BMEG 802 – Advanced Biomedical Experimental Design and Analysis

Bayesian Statistics

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Recap

- Maximum Likelihood Estimation (MLE)
 - Probability Distribution Function
 - Likelihood function
 - 3 Ways to find the Maximum Likelihood Estimation
 - Analytical (Calculus)
 - Brute Force (Grid Search)
 - Optimization (Gradient Descent)

Today

Bayesian Statistics

- Derivation from Set Theory
- Point Probabilities
 - priors, likelihood, posteriors
- Continuous Probabilites
 - Analytical (Conjugate Priors)
 - Numerical

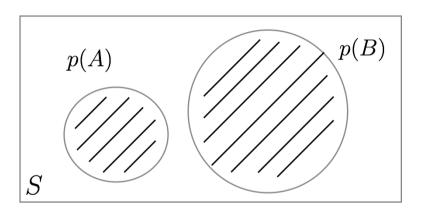
Bayesian vs. Frequentist

- Data are treated as fixed observations vs. data (sample) treated as a random variable
- Models (parameters) are treated as random variables vs. models (population parameters) are treated as fixed quantities
- we compute the probability of all models vs. we compute the probability of one model (H₀)
- we end up with a richer understanding of relative probability of all models vs. we make a decision (reject H_0 or not)

Notation

- S = sample space (all possible outcomes)
- p(A) = probability of event A
- $A \cup B =$ union of events A and B
- $A \cap B =$ intersection of events A and B
- p(B|A) = probability of B given A
- p(A') or $p(A^C)$ or $p(\bar{A}) = \text{complement probability of } p(A)$

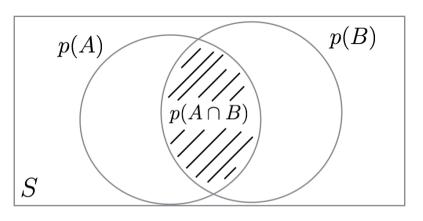
Mutually Exclusive (Disjoint Probability)



$$p(A \cup B) = p(A) + p(B)$$

0.7 = 0.4 + 0.3

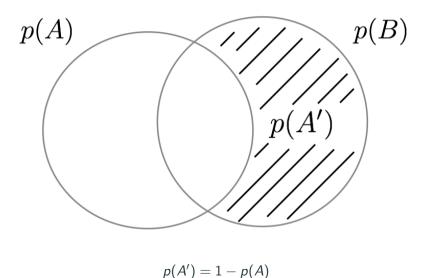
Joint Probability



$$p(A \cup B) = p(A) + p(B) - p(A \cap B)$$
$$0.5 = 0.4 + 0.3 - 0.2$$

• the probability of two events occuring simultaneously

Complement Probability

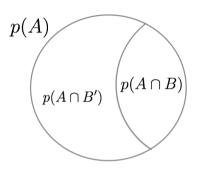


Marginal Probability

H	Red	Yellow	Green	Marginal probability P(H)
Not Hit	0.198	0.09	0.14	0.428
Hit	0.002	0.01	0.56	0.572
Total	0.2	0.1	0.7	1

- Probability of a single event occurring (hit), independent of other events (light)
- e.g., probabilities of getting in an accident at an intersection irrespective of lights
- note: joint probabilities in each cell

Marginal Probability



$$p(A \cap B') = p(A) - p(A \cap B)$$

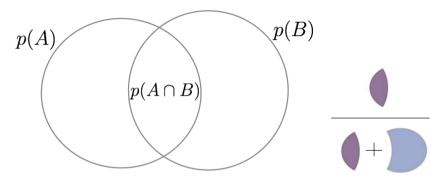
The marginal p(A) or p(B) is found by summating their disjoint parts.

$$p(A) = p(A \cap B) + p(A \cap B')$$
, and similarly $p(B) = p(A \cap B) + p(A' \cap B)$

Conditional Probability

- p(accepted) = 0.3
- p(funding|accepted) = 0.43
- $p(funding \cap accepted) = p(funding | accepted) \cdot p(accepted)$
- $p(funding \cap accepted) = 0.43 \cdot 0.3 = 0.13$
- Probability that an event occurs given that another specific event has already occurred

Conditional Probability



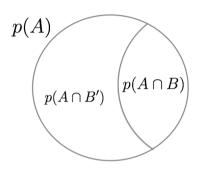
$$p(A \cap B) = p(B|A) \cdot p(A)$$

$$p(B|A) = \frac{p(A \cap B)}{p(A)}$$

$$p(A \cap B) = p(A|B) \cdot p(B) \text{ (in terms of B)}$$

$$p(A|B) = \frac{p(A \cap B)}{p(B)}$$

Conditional Probability Complements



$$p(A \cap B') = p(B'|A) \cdot p(A)$$

Other friendly complements:
 $p(A' \cap B) = p(B|A') \cdot p(A')$
 $p(A' \cap B') = p(B'|A') \cdot p(A')$

Good News!

Bayes' Theorem is simply a conditional probability!

Deriving Bayes' Theorem

Remember:

- $p(A|B) = \frac{p(A \cap B)}{p(B)}$, (eq.1)(slide 11)
- $p(A \cap B) = p(B|A) \cdot p(A), (eq.2) (slide 11)$

Substitute (eq.2) into (eq.1):

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}, (eq.3)$$

That's it!

In terms of statistical models:

$$p(model|data) = \frac{p(data|model) \cdot p(model)}{p(data)}$$

Handy Dandy Steps for Point Estimates

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}, (eq.3)$$
Calculate $p(B)$ by using its marginal probability
$$p(B) = p(A \cap B) + p(A' \cap B), (eq.4) \text{ (slide 9)}$$

Handy Dandy Steps for Point Estimates

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}, (eq.3)$$

$$Calculate \ p(B) \ by \ using \ its \ marginal \ probability$$

$$p(B) = p(A \cap B) + p(A' \cap B), (eq.4) \text{ (slide 9)}$$

$$Substitute \ (eq.4) \ into \ (eq.3)$$

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(A \cap B) + p(A' \cap B)}, (eq.5)$$

Handy Dandy Steps for Point Estimates

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}, (eq.3)$$

Calculate p(B) by using its marginal probability

$$p(B) = p(A \cap B) + p(A' \cap B), (eq.4)$$
(slide 9)

Substitute (eq.4) into (eq.3)

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(A \cap B) + p(A' \cap B)}, (eq.5)$$

Since,

$$p(A \cap B) = p(B|A) \cdot p(A), (eq.6) \text{ (slide 11)}$$

$$p(A' \cap B) = p(B|A') \cdot p(A'), (eq.7) \text{(slide 12)}$$

Substitute (eq.6) and (eq.7) into (eq.5)

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B|A) \cdot p(A) + p(B|A') \cdot p(A')}, (eq.8)$$

Why Bayesian???

Powerful way to continually account for new evidence given prior beliefs

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}$$
x Likelihood
(new evidence)
Prior
Posterior

POINT PROBABILITES

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Powerful way to continually account for new evidence given prior beliefs

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B|A) \cdot p(A) + p(B|A') \cdot p(A')}$$
x Likelihood
(new evidence)
Prior
Posterior

p(B) = marginal probability (e.g., true positive & false positive tests) classic example: A = preg, A' = not preg, B = + test, B' = - test

Let's say you take a COVID test and it comes out positive. What is the probability that you have COVID?

- Lets assume our initial, prior guess on whether we have covid based on an exposure is 40%, p(+covid) = 0.4.
- The probability of having a positive test given you have covid is 75%, $p(+test \mid + covid) = 0.75$. (i.e., test sensitivity)
- The probability of having a positive test given you do NOT have covid is 25%: $p(+test \mid -covid) = 0.25$. (i.e., 1 test specificity)
- We observe a + test. What is the probability that you have COVID, $p(+covid \mid + test)$?
- note: these are fictitious numbers

$$p(+covid \mid + test) = \frac{p(+test \mid + covid) \cdot p(+covid)}{p(+test \mid + covid) \cdot p(+covid) + p(+test \mid - covid) \cdot p(-covid)}$$

Knowns:

$$p(+covid) = 0.4$$

 $p(+test \mid + covid) = 0.75$
 $p(+test \mid - covid) = 0.25$

Unknowns:

$$p(-covid) = ?$$

 $p(+covid \mid + test) = ?$

Knowns:

$$p(+covid) = 0.4$$

 $p(+test \mid + covid) = 0.75$
 $p(+test \mid - covid) = 0.25$

Unknowns:

$$p(-covid) = 0.6; (1 - 0.4)$$

 $p(+covid \mid + test) = ?$

$$p(+covid \mid + test) = \frac{p(+test \mid + covid) \cdot p(+covid)}{p(+test \mid + covid) \cdot p(+covid) + p(+test \mid - covid) \cdot p(-covid)}$$
$$p(+covid \mid + test) = 0.67 = \frac{0.75 \cdot 0.4}{0.75 \cdot 0.4 + 0.25 \cdot 0.6}$$

Why Bayesian?

Powerful way to continually account for new evidence given prior beliefs



p(A|B) becomes p(A) on the next iteration!

Updating

- updating the model (i.e., take another test). Seems like an appropriate thing to do in science
- when new data are gathered, we can re-evaluate a hypothesis
- we do not begin anew (ignorant) each time we ask a question
- previous research provides us information about the merits of the hypothesis
- the posterior from the previous model becomes the prior for the new model

Let's continue from our previous example. We take another test and it comes out positive. What is our probability of having covid given another positive test?

Knowns:

$$p(+covid) = 0.67$$

 $p(+test \mid + covid) = 0.75$
 $p(+test \mid - covid) = 0.25$

Unknowns:

$$p(-covid) = ?$$

 $p(+covid \mid + test) = ?$

Knowns:

$$p(+covid) = 0.67$$

 $p(+test \mid + covid) = 0.75$
 $p(+test \mid - covid) = 0.25$

Unknowns:

$$p(-covid) = 0.33; (1 - 0.67)$$

 $p(+covid \mid + test) = ?$

$$p(+covid \mid + test) = \frac{p(+test \mid + covid) \cdot p(+covid)}{p(+test \mid + covid) \cdot p(+covid) + p(+test \mid - covid) \cdot p(-covid)}$$
$$p(+covid \mid + test) = 0.86 = \frac{0.75 \cdot 0.67}{0.75 \cdot 0.67 + 0.25 \cdot 0.33}$$

Let's keep going and pretend we observed 5 spikes in row from our initial belief of 40%. Calculating p(open|spike) for each iteration leads to:

0.67

0.86

0.95

0.98

0.99

Influence of PRIOR beliefs

How much of our prediction is influenced by our prior belief that the hand is open or not?

Prior	Posterior
0.1	0.25
0.2	0.43
0.3	0.56
0.4	0.67
0.5	0.75
0.6	0.82
0.7	0.88
8.0	0.92
0.9	0.96

CONTINUOUS PROBABILITIES

Bayes with Probability Distributions

- in previous example, the likelihood and prior were both single quantities (point probabilities)
- typically Bayesian approaches use full probability distributions
- essentially allows us to evaluate probability of a whole range of possible models, at once

Back to the Bayesics

$$p(A|B) = \frac{p(B|A) \cdot p(A)}{p(B)}$$

$$p(\theta|y)d\theta = \frac{p(y|\theta) \cdot p(\theta)d\theta}{\int_{a}^{b} p(y|\theta) \cdot p(\theta)d\theta}$$

$$p(\theta|y) = \frac{p(y|\theta) \cdot p(\theta)}{\int_{a}^{b} p(y|\theta) \cdot p(\theta)d\theta}$$

Marginal probability a normalization constant (sum of prior and likelihood)

$$p(\theta|y) \propto p(y|\theta) \cdot p(\theta)$$

 $More\ details\ here:\ https://ocw.mit.edu/courses/mathematics/18-05-introduction-to-probability-and-statistics-spring-2014/readings/MIT18_05S14_Reading13a.pdf$

Bayes' Theorem

$$p(\theta|y) \propto \mathcal{L}(y|\theta) \cdot p(\theta)$$
x Likelihood
(new evidence)
Prior
Update Belief

posterior, likelihood, prior can all be defined with probability distributions

Continuous Probability Example

- Let's revisit the coin flipping example.
- Is the coin fair (w = 0.5)?
 - model: some proposed process by which the outcome of our coin flip is determined.
 - Binomial Distribution
 - data: k = 2 heads (# of successes), n = 3 flips

Likelihood Function

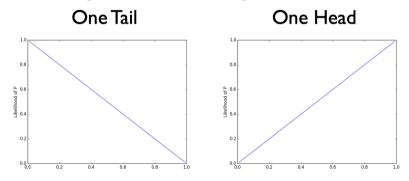
$$\mathcal{L}(w|n,y) = \frac{n!}{y!(n-y)!} w^y (1-w)^{n-y}$$

Likelihood of W for a single toss

w = 0 represents the coin is perfectly weighted towards Tails

w=1 represents the coin is perfectly weighted towards Heads

First, lets consider a single toss of Tails and a single toss of Heads



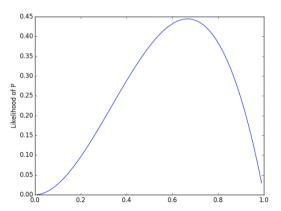
Note: x-axis is w

Likelihood of P for 3 tosses

w = 0 represents the coin is perfectly weighted towards Tails

w = 1 represents the coin is perfectly weighted towards Heads

2 Heads, 1 Tails



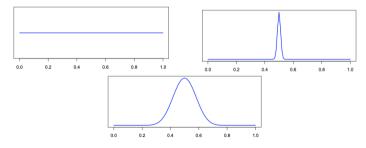
The Prior

$$p(\theta|y) \propto \mathcal{L}(y|\theta) \cdot p(\theta)$$

- What is our prior belief? Is the coin weighted or not?
 - "uninformative"
 - flat prior all values of w are equally likely (Bayes / Laplace):
 - Others Jeffrey's prior, reference priors, maximum entropy
 - Informative
 - we have some previous experience / evidence

The Prior Cont'd

- 1. I have no clue what P is (flat prior)
- 2. Every coin we have seen in the past has been fair
- 3. Most coins have been relatively fair



Calculating the Posterior

- Analytical
- Numerical

Analytical

Find a Conjugate Prior

- 1. IF, the posterior and prior are the same type of distribution, they are conjugate distributions
- 2. THEN, the prior is a conjugate prior to the likelihood function
- 3. If we have a conjugate prior we can use hyperparameters to solve the posterior
- 4. Hyperparameters solved for many distributions: Conjugate Priors Wikipedia
- The Binomical distribution conjugate prior is the Beta distribution
- The Normal distribution conjugate prior is the Normal distribution

Analytical

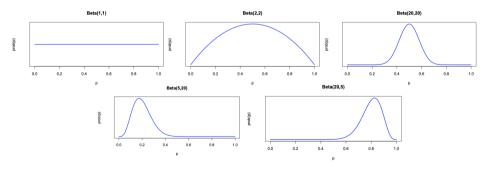
- Back to our coin flipping example.
- Our Likelihood function is the Binomial Distribution
- Binomial distribution's conjugate prior = Beta distribution

$$p(w|\alpha,\beta) = \frac{1}{B(\alpha,\beta)} w^{\alpha-1} (1-w)^{\beta-1}$$

Beta Distribution Conjugate Prior

$$p(w|\alpha,\beta) = \frac{1}{B(\alpha,\beta)} w^{\alpha-1} (1-w)^{\beta-1}$$

Beta distribution's range [0,1] convenient for our prior w somewhere between 0 and 1



*note: x-axis is w

Analytically Calculating the Posterior

- If we have a conjugate prior
- Then, the posterior is calculated from parameters used in the likelihood and prior (called hyperparameters)

Analytically Calculating the Posterior Cont'd

Prior:
$$p(w|\alpha,\beta) = \frac{1}{B(\alpha,\beta)} w^{\alpha-1} (1-w)^{\beta-1}$$

• parameters are: α, β

Likelihood:
$$\mathcal{L}(w|n,y) = \frac{n!}{y!(n-y)!} w^y (1-w)^{n-y}$$

• parameters are: k, n

Posterior = Likelihood * Prior => after some heavy calculus / algebra

Posterior:
$$p(w|\alpha_h, \beta_h) = \frac{1}{B(\alpha_h, \beta_h)} w^{\alpha_h - 1} (1 - w)^{\beta_h - 1}$$

- hyperparameters are: α_h, β_h
- $\bullet \quad \alpha_h = k + \alpha$
- $\beta_h = n k + \beta$

Let's update based on each toss

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coin flip: n = 3 trials, k = 2 success
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- 1st toss = Heads
- 2nd toss = Tails
- 3rd toss = Heads

Let's assume a flat prior

all w equal

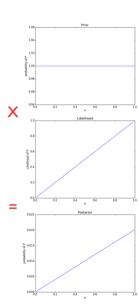
Toss One - Heads

Prior:
$$p(P|\alpha,\beta); \alpha=1,\beta=1$$

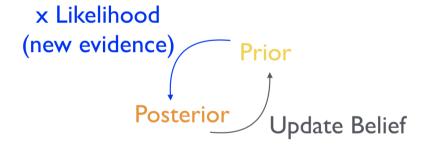
Likelihood:
$$L(P|k, n); k = 1, n = 1$$

Posterior: $p(P|\alpha_h, \beta_h); \alpha_h = 2, \beta_h = 1$

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Same Update Procedure

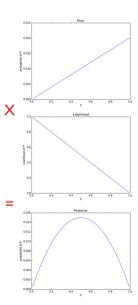


Toss Two - Tails

Prior:
$$p(P|\alpha,\beta); \alpha=2, \beta=1$$

Likelihood:
$$L(P|k, n); k = 0, n = 1$$

Posterior: $p(P|\alpha_h, \beta_h)$; $\alpha_h = 2, \beta_h = 2$

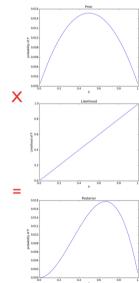


Toss Three - Heads

Prior:
$$p(P|\alpha, \beta); \alpha = 2, \beta = 2$$

Likelihood:
$$L(P|k, n); k = 1, n = 1$$

Posterior: $p(P|\alpha_h, \beta_h); \alpha_h = 3, \beta_h = 2$

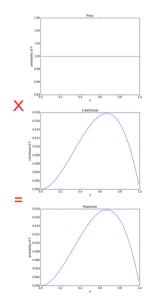


OR, in a Single Step (all tosses already made)

Prior:
$$p(P|\alpha, \beta); \alpha = 1, \beta = 1$$

Likelihood:
$$L(P|k, n); k = 2, n = 3$$

Posterior: $p(P|\alpha_h, \beta_h); \alpha_h = 3, \beta_h = 2$

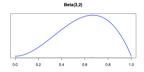


Effect of Prior



Analytically Describing the Posterior

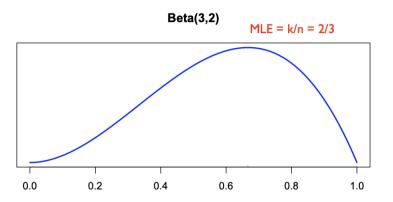
Graphically:



Summary Statistics:

- expression for mean, variance, mode, etc.
- mean $= \frac{\alpha}{\alpha + \beta}$
- variance $=\frac{\alpha\beta}{(\alpha+\beta)^2(\alpha+\beta+1)}$

Bayesian vs. MLE



Bayesian: the posterior tells us the probability of all possible w's

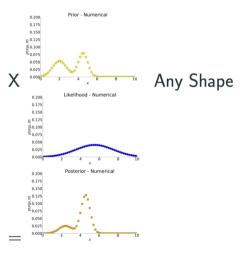
MLE (frequentist approach): w is 0.667

does not incorporate prior information

Numerically Calculating the Posterior

- Option 1: Grid approximation by discretizing the prior
- Option II: Markov Chain Monte Carlo (MCMC)

Grid Approximation



Tradeoff between grid coarseness and speed

Criticisms of Bayesian Approach

- The prior: too much "subjectivity"?
- Data fixed, models (parameters) random
- Often difficult to find analytical solutions

Advantages of Bayesian Approach

- Bayesian approach allows for incorporating previous findings in a principled way
- frequentist involves testing only one hypothesis (model): the null hypothesis ...
 Bayesian estimates probability of all models (parameter values)
- interval estimates (and other such measures of posterior) have a clearer meaning than CIs in frequentist approaches
- in Bayesian approach we get full posterior distribution, a much richer picture than just a mean \pm CI or s.e.

Applications

- Linear Regression: https://statswithr.github.io/book/introduction-to-bayesian-regression.html
 - pretty heavy stuff...
- Kalman Filters
- Multisensory Integration, Illusions, Sensorimotor Adaptation, etc.
- Bayes Factors
- etc.

Next Week

Markov Chain Monte Carlo (MCMC)

sampling the posterior