

# Principled Bayesian Workflow: Practicing Safe Bayes.

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So now you can Bayes - need to do it safely (especially in a regulatory environment).

- ▶ Motivated by courses I gave in Health Canada in 2010/11 - “resulted” in less than critical review of submissions.
- ▶ Today is largely to provide motivation to read and work through material/programs developed by Michael Betancourt on Principled Bayesian Workflow.
- ▶ GitHub for Rstan
- ▶ GitHub for Pystan
- ▶ I'll do a largely conceptual introduction and work through a “toyed down” example. Files on Github.

## My relevant background.

- ▶ Provided statistical support and mentor-ship to research fellows at the University of Toronto and Toronto Hospital (1985-1998).
- ▶ Had no degree in statistics but still taught one course and gave multiple tutorials.
- ▶ Was recruited to the Ottawa Hospital (1998-2001).
- ▶ Did a DPhil in Statistics at Oxford (2001-2007).
- ▶ Visited the Statistical Sciences Department at Duke University, *the world's leading center for Bayesian statistics* (2007-2008).
- ▶ Joined Health Canada 2009, gave Bayesian Courses in 2010/11.
- ▶ Gave webinars on Bayes for the American Statistical Society and CSEB in 2012 (first reviewed positively, not the second).
- ▶ Currently an author on Statistical Modeling, Causal Inference, and Social Science a mostly Bayesian perspective blog.

# Conceptual review of Bayes.

Seeing all of Bayes all at once.

- ▶ Need to go back and focus on basics - not always popular.
- ▶ Once told that they hide the best techniques in martial arts in the beginners form.
- ▶ Two stage simulation is not a cheap trick but rather a way to *really, really* understand (first came up with it in 2004).
- ▶ Andrew Gelman blogged numerous times - in Bayes you write down the model and simply watch it work (crickets. . .).
- ▶ Or as Michael Betancourt put it: Work it, Make it, Do it, Makes Us Harder, Better, Faster, Stronger.
- ▶ Numerous (wild?) claims about Bayes (ASA webinar).
- ▶ **Take no one's word for it!**

# Conceptual review of statistics.

Seeing all of statistics all at once.

- ▶ In both Bayes and Frequentist statistics the holy grail is clearly seeing what repeatedly would happen.
- ▶ When trying to learn from observations like the ones you have in hand.
- ▶ Helpful? metaphors of discerning what cast the shadows just from the shadows.
- ▶ Think of learning about an object just from the shadows it casts while being unable to look directly at the object. We see those shadows but really are only interested in what is casting them. They may look very scary, but the object casting them maybe mice.
- ▶ My toy example today will be annual faint rates in business leaders “rounds” at Toronto Hospital where 3 out of 9 fainted the one year I heard about it.

## Strategy: Generate realistic Fake Universes to better grasp reality.

- ▶ Make something where you can easily “see” what would repeatedly happen.
- ▶ In analytical chemistry you can repeatedly spike test tubes with know trace amounts of X.
- ▶ In statistics we can't repeatedly spike humans with say know faint rates.
- ▶ So we have to represent faint rates abstractly - make a fake universe were you set the rate.
- ▶ Most statisticians prefer to use probability models (and math) to do that.
- ▶ Today almost anyone can use probability models (and simulation!!!) to do that.
- ▶ Two stage simulation is the just the Bayesian version of generating realistic? fake universes!

# Why make fake universes of faint rates and occurrences?

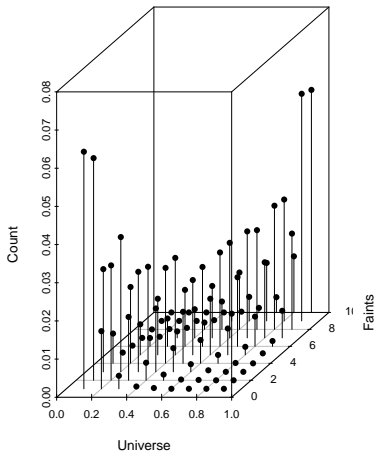
Two stage simulation, simulate the faint rate and then number of faints given that rate.

- ▶ Once the joint model is set - the prior and data generating model - Bayes is deductive.
- ▶ The joint model is the first premise, the data is the second premise and the posterior is the conclusion.
- ▶ Assuming adequate sampling from the posterior - the quality of what follows totally depends the premises.
- ▶ *Can be very bad if the premises are deficient (either joint model or data).*

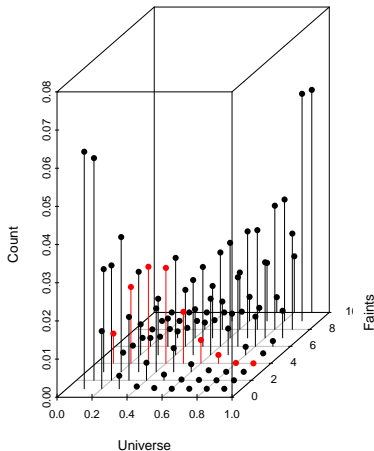
# Lets try to see the adequacy of joint model issue in plots.

First premise on the left - do they adequately represent our grasp of reality (as in a artist's pencil sketch)? Note the conclusion in red on the right is clearly contained in the premises (QED).

Set of Fake Universes for Fainting.



Note the posterior it implies, given 3 out of 9

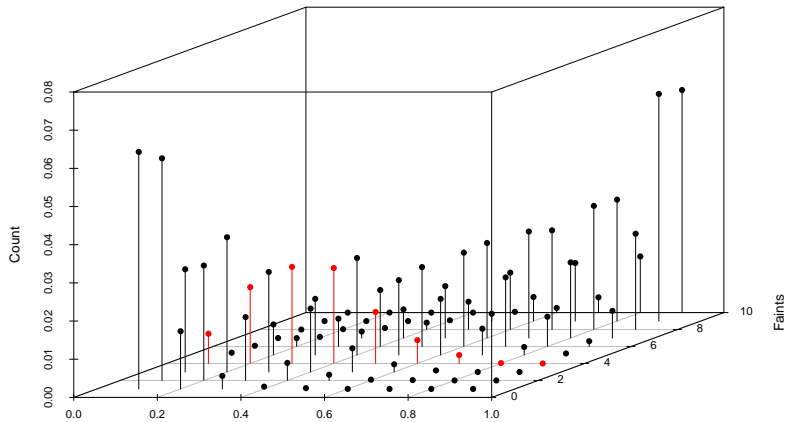




## Consistent with domain expertise?

Note the 9 fake universes on the x axis and 10 possible faint totals on the y axis with how often on the z axis. A proportional set of fake universes and faint totals that we judge could be encountered in topics *like this* (Bayes/Frequency reference sets).

**The proportional subset of universes most compatible with our universe shown in red**



But so much more to make of the set of fake universes generated.

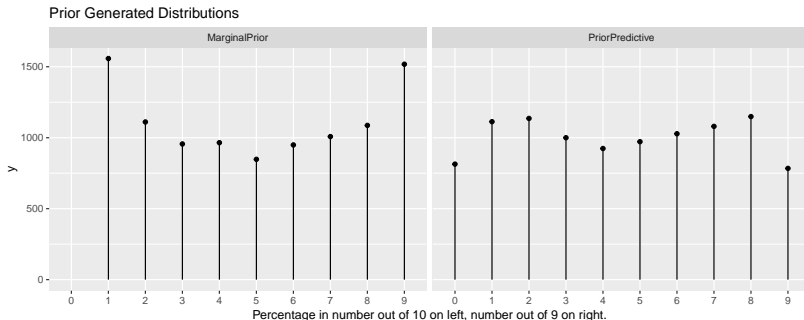
##	TrueProp1	3 Faints	TrueProp2	Not 3 Faints
## 1	0.3	3	0.1	0
## 2	0.3	3	0.9	8
## 3	0.6	3	0.7	7

- ▶ Samples on the left are valid posterior samples - right?
- ▶ What are the ones on the right a valid posterior sample from?
- ▶ But now you know how to get many posterior samples using MCMC.
- ▶ If the MCMC is valid the ranks of two stage sample with the MCMC sampleS (say 500) should be uniform.
- ▶ Now we can check the MCMC sampling for every point in the fake universes generated (true parameter, valid sample).

## Things to consider on the last slides.

- ▶ Are these fake universes and repeatedly happening faint totals roughly like our universe?
- ▶ Biggest fail so far may be air density greater than concrete in air quality modeling.
- ▶ **If not too wrong, provides an ample laboratory to evaluate the performance of the given joint model and data!**
- ▶ Often a good idea to use different joint model to create *wronger* fake universes for the joint model been used in the analysis.

# Marginal views and disagreements on views.



- ▶ Prior predictive distribution is on the right - seem realistic?
- ▶ Marginal prior distribution over all universes is on the left - do these parameters seem realistic.
- ▶ Some argue only look at the first (which has advantages especially with many parameters) while I would look at both.

## Three tasks in principled Bayesian workflow to be safer.

- ▶ Three tasks in Bayesian Workflow - assess adequacy of joint model, posterior sampling and data.
- ▶ Assessing the adequacy of joint model has only become practical in the past 5 or so years.
- ▶ Around 2012 even experts were refusing (even saying not kosher in Bayes).
- ▶ Still some resistance to assessing the adequacy of the data?
- ▶ Lots can and did go wrong and it remains the most challenging aspect (and an incentive to just ignore it?).

# What two stage fake Bayesian universe generation enables.

Makes safe Bayes practical (was not until recently!).

- ▶ Domain Expertise Consistency: Is our model consistent with our domain expertise?
- ▶ Computational Faithfulness: Are our computational tools sufficient to accurately fit the model?
- ▶ Model Sensitivity: How do we expect our inferences to perform over the possible fake universes?
- ▶ (Model Adequacy: Is our model rich enough to capture the relevant structure of our universe?)

## Some early experience.

- ▶ Used someone's published Bayesian model - obtained positive posterior probabilities for negative proportions.
- ▶ The priors they had specified were independent even though proportions were dependent.
- ▶ Contacted the author - they were unconcerned.
- ▶ Claimed it was a result of my not having adequate data and so no need to worry.
- ▶ Sometimes it is hard to know when you have adequate data but one of the premises is obviously false and *should* be fixed.
- ▶ (This sort of thing was be done a lot in early 2000s given limitations).

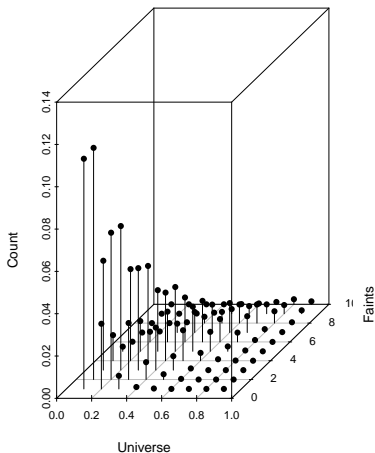
## More early experience.

- ▶ One of the first I heard about without an obvious error was from Peter F. Thall at M.D. Anderson Cancer Center.
- ▶ Not obvious at the time perhaps given the lack of awareness of the dangers of (silly?) default flat priors.
- ▶ In 2011 I emailed him and got back *It seems that we are both in the How to tell if you have a prior with undesirable properties business. I got into this years ago when I was doing CRM dose finding with the model  $\text{logit}\{\text{Prob}(\text{toxicity} \mid \text{dose})\} = a + b \log(\text{dose})$  and just assumed noninformative normal priors on  $a$  and  $b$ , with variance = 100. The resulting CRM design does strange things with the first 3 to 6 patients. It drove me and my programmer crazy for a month.*
- ▶ See Stan Prior Choice Recommendations wiki.

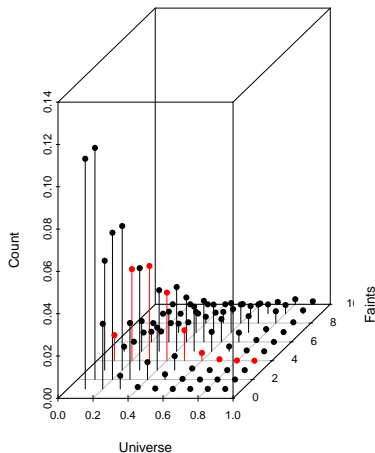


# A more sensible set of fake universes?

Set of Fake Universes for Fainting.

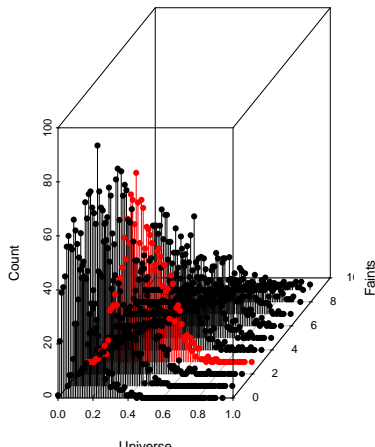
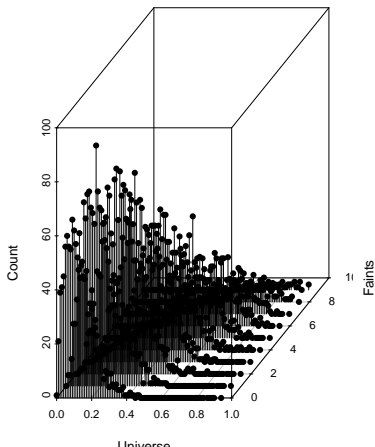


Note the posterior it implies, given 3 out of 9

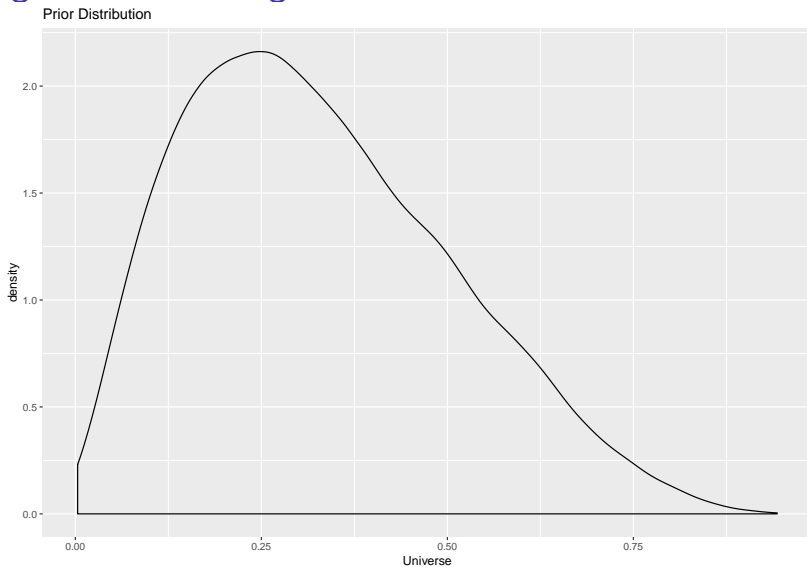


## Domain Expertise Consistency: Is our model consistent with our domain expertise?

Now use Rstan which has prior predictive functions (used to have to leave out data ~ statement in model block).

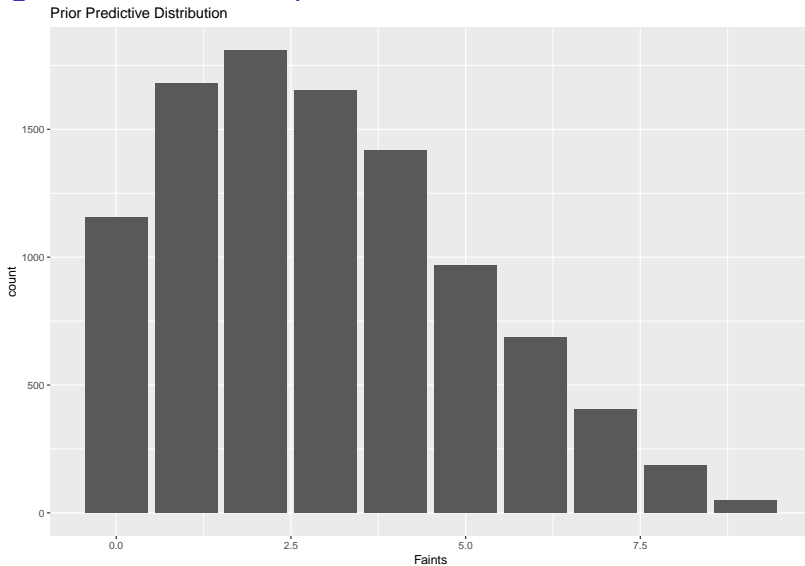


# Marginal views: Marginal Prior



- Do these parameters seem realistic?

# Marginal views: Prior predictive distribution.



- Prior predictive distribution - these observations seem realistic?

# Computational Faithfulness: Are our computational tools sufficient to accurately fit the model?

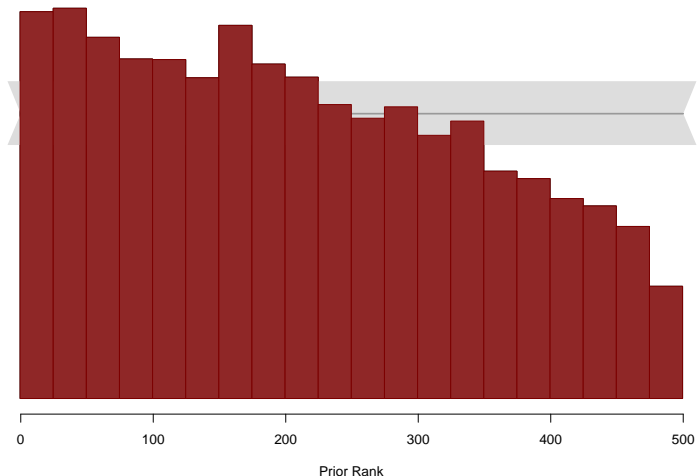
Lets look at some samples of the two stage simulation

##	Universe	Faints	Universe	Faints
## 1	0.2418439	3	0.09257722	0
## 2	0.4519268	3	0.19122446	0
## 14	0.3327996	3	0.25878004	4

- ▶ If the MCMC is valid the ranks of two stage sample with the MCMC sampleS should be uniform.
- ▶ Now we can check the MCMC sampling for every point in the fake universes generated (true parameter, valid sample).

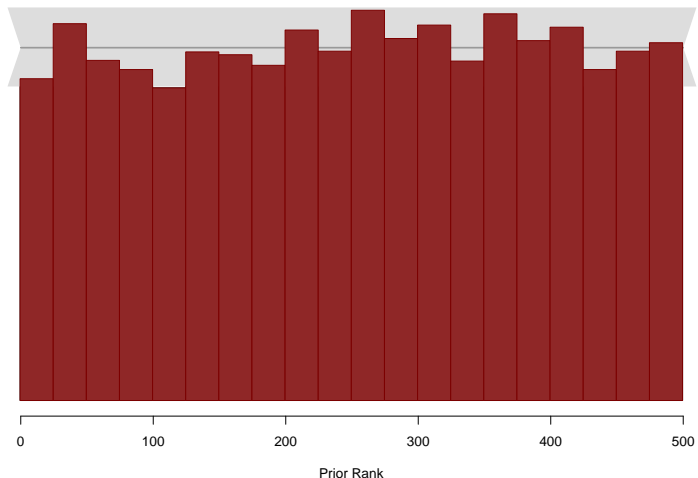
## Computational Faithfulness - are ranks uniform?

Lets obtain 500 MCMC samples for each two stage sample in Rstan.  
What went wrong? My priors did not match.



## Now, lets do that correctly in Stan

Lets obtain 500 MCMC samples for each two stage sample in Rstan using same priors.



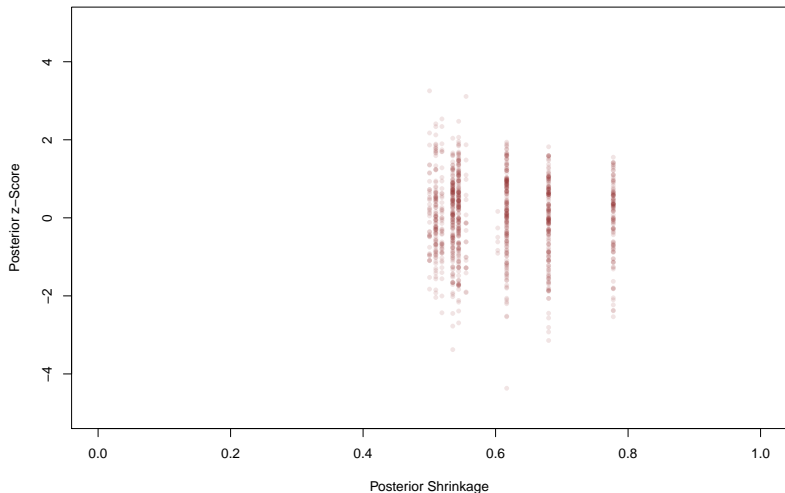
Now actually you should first run a set of utility diagnostic checks.

```
warning_code <- ensemble_output[1,]  
if (sum(warning_code) != 0) {  
  print ("Some simulated posterior fits in the joint ensemble")  
  for (r in 1:R) {  
    if (warning_code[r] != 0) {  
      print(sprintf('Replication %s of %s', r, R))  
      util$parse_warning_code(warning_code[r])  
      print(sprintf('Simulated theta = %s', simu_thetas[r]))  
      print(" ")  
    }  
  }  
} else {  
  print ("No posterior fits in the joint ensemble encountered")  
}
```

```
## [1] "No posterior fits in the joint ensemble encountered"
```



Model Sensitivity: Generic Posterior Shrinkage ( $1 - \text{post.var}/\text{prior.var}$ ) and z-Scores  $((\text{post.mean} - \text{truth})/\text{post.sd})$



## Model Sensitivity: More focused assessments.

For instance type 1 and 2 errors.

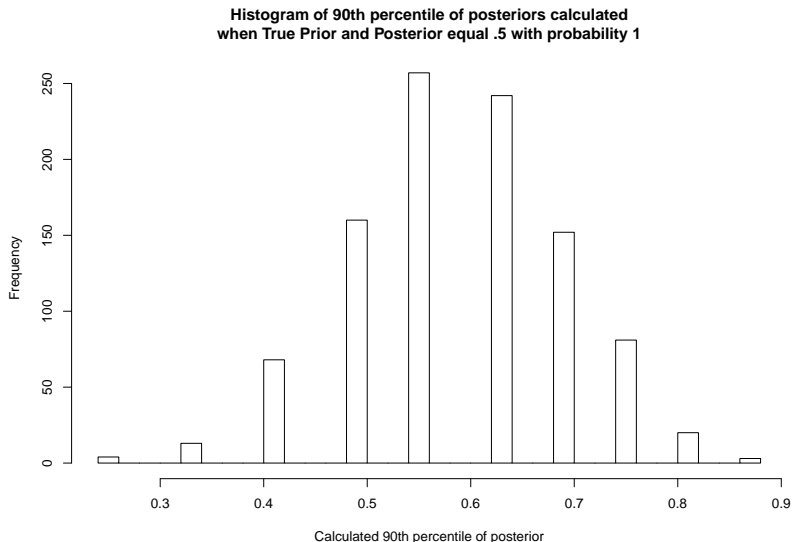
- ▶ FDA 2010 guidance for doing simulations to obtain operating characteristics – FDA usually recommends you provide simulations of your trial at the planning (or IDE) stage. This will facilitate FDA's assessment of the operating characteristics of the Bayesian trial; specifically, the type I and type II error rates.
- ▶ Actually carried out a feasibility study of calculating the type I and type II error rates with a major pharma consulting firm and Yongmin Yu at Health Canada in 2012.

Pick one universe and repeatedly sample and see what happens.

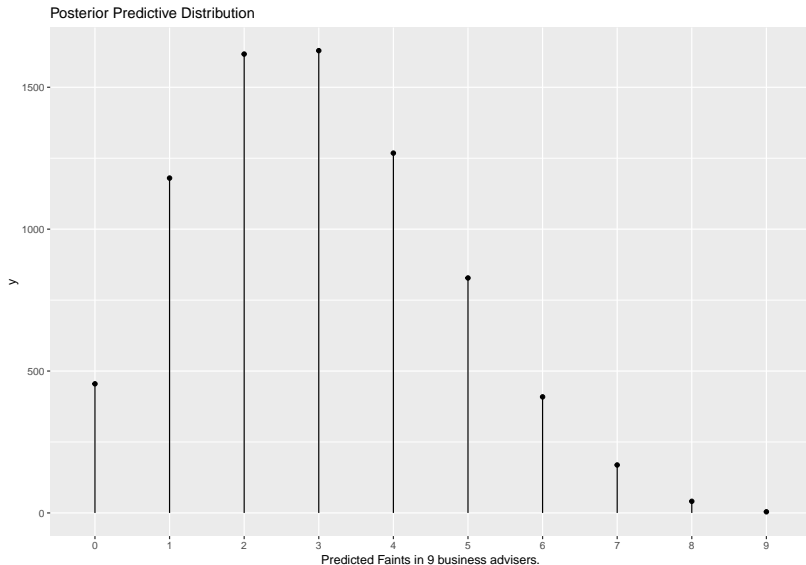
- ▶ To re-use my toy example think needing to know if rate not  $\geq 50\%$ .
- ▶ Set the probability of fainting to .5 and simulate data just for that universe.
- ▶ Type 1 error? See how often -
- ▶  $(\text{post\_mean\_theta} - .5) / \text{post\_sd\_theta} < -1.65$ .
- ▶ Calculate the 90th percentile of posterior probability.
- ▶ See how often that is claimed  $< 50\%$ .

# What repeatedly happens in this fake universe.

```
## [1] "Type 1 Error = 0.085"
```

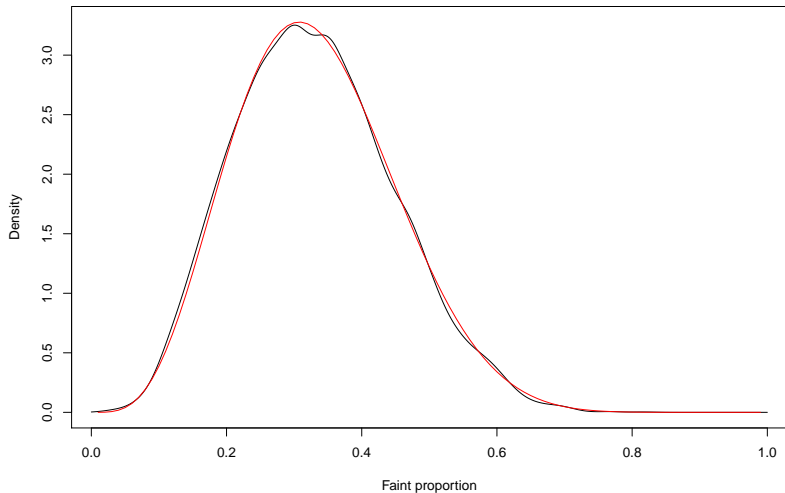


# Model Adequacy: Checking the posterior itself.



# Posterior plot.

Density estimate from posterior sample against closed form posterior (in red)



# Checking accuracy of posterior quantiles.

Check Stan website if a concern.

