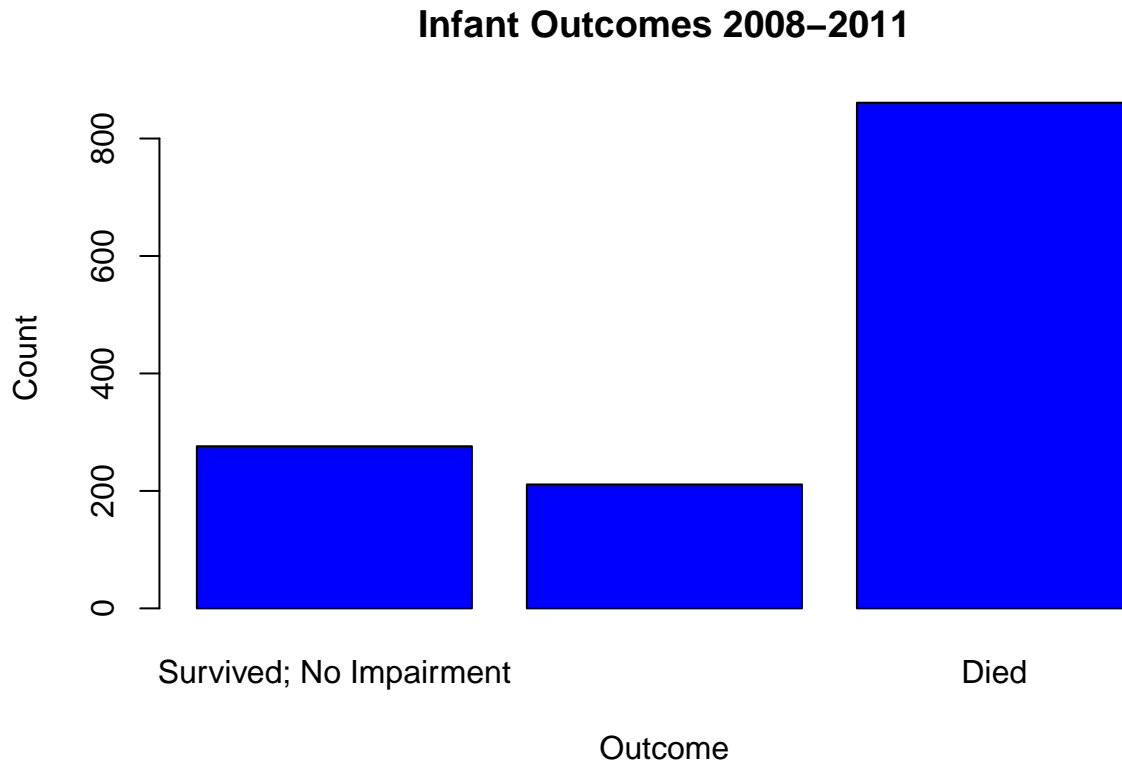


Homework 1, Due September 13 at 11:45am by uploading .Rmd and .pdf files on Sakai

Provide clearly documented code in an R markdown document where appropriate. Points will be deducted in each question if results are not reproducible. Whenever possible, provide statistical evidence to support your answers.

1. Analysis of periviable neonates (8 points)

In a 2017 New England Journal of Medicine paper, Duke researchers examined outcomes of infants born at 22-24 weeks of gestation. Of 1348 infants in the study born at these gestational ages in 2008-2011, 861 died, 211 survived with neurodevelopmental impairment, and 276 survived without neurodevelopmental impairment.



Suppose historical rates of all three outcomes (from the early 2000's) are 15% survival without impairment, 15% survival with impairment, and 70% death. Estimate the proportions of infants surviving without impairment, surviving with impairment, and dying, providing point and interval estimates. Evaluate the hypothesis that the current rates are equal to the historical rates of 15, 15, and 70%, respectively, using the multinomial distribution. Display results using informative graphics.

Solution

I use the conjugate prior of Dirichlet($\alpha_1, \alpha_2, \alpha_3$). I use three different sets of α to demonstrate varying levels of prior certainty. In a situation like this study, we have a total sample of 1348 ($n = 1348$), which is a good sample size, but when we compare this with historical averages and we are interested in comparing current rates versus past rates, we should have a relatively strong prior representing historical knowledge.

We understand that under this Dirichlet-multinomial framework, the posterior distribution will follow a Dirichlet($n_1 + \alpha_1, n_2 + \alpha_2, n_3 + \alpha_3$).

```
n1 = 276
n2 = 211
n3 = 861
prior.mat = matrix(c(15,15,70,202,202,944,750,750,3500), nrow = 3, byrow = FALSE)
colnames(prior.mat) = c("rel_weak", "equal", "strong")
rownames(prior.mat) = c("alpha_1", "alpha_2", "alpha_3")
```

```
post.mat = prior.mat
post.mat[1,1:3] = post.mat[1,1:3]+n1
post.mat[2,1:3] = post.mat[2,1:3]+n2
post.mat[3,1:3] = post.mat[3,1:3]+n3
post.mat
```

```
##           rel_weak equal strong
## alpha_1      291   478   1026
## alpha_2      226   413    961
## alpha_3      931  1805   4361
```

```
kable(prior.mat)
```

	rel_weak	equal	strong
alpha_1	15	202	750
alpha_2	15	202	750
alpha_3	70	944	3500

```
kable(post.mat)
```

	rel_weak	equal	strong
alpha_1	291	478	1026
alpha_2	226	413	961
alpha_3	931	1805	4361

```
n.reps = 1e06
post.alpha1 = rdirichlet(n.reps, post.mat[,1])
post.alpha2 = rdirichlet(n.reps, post.mat[,2])
post.alpha3 = rdirichlet(n.reps, post.mat[,3])

par(mfrow = c(1,3))
plot(density(post.alpha1[,1]), xlim = c(.10,.75), lwd = 2, main = expression("Posterior of ", theta), xlab = "theta", ylab = "density")
lines(density(post.alpha1[,2]), col = "red", lwd = 2)
lines(density(post.alpha1[,3]), col = "blue", lwd = 2)

plot(density(post.alpha2[,1]), xlim = c(.10,.75), lwd = 2, main = expression("Posterior of ", theta), xlab = "theta", ylab = "density")
lines(density(post.alpha2[,2]), col = "red", lwd = 2)
lines(density(post.alpha2[,3]), col = "blue", lwd = 2)

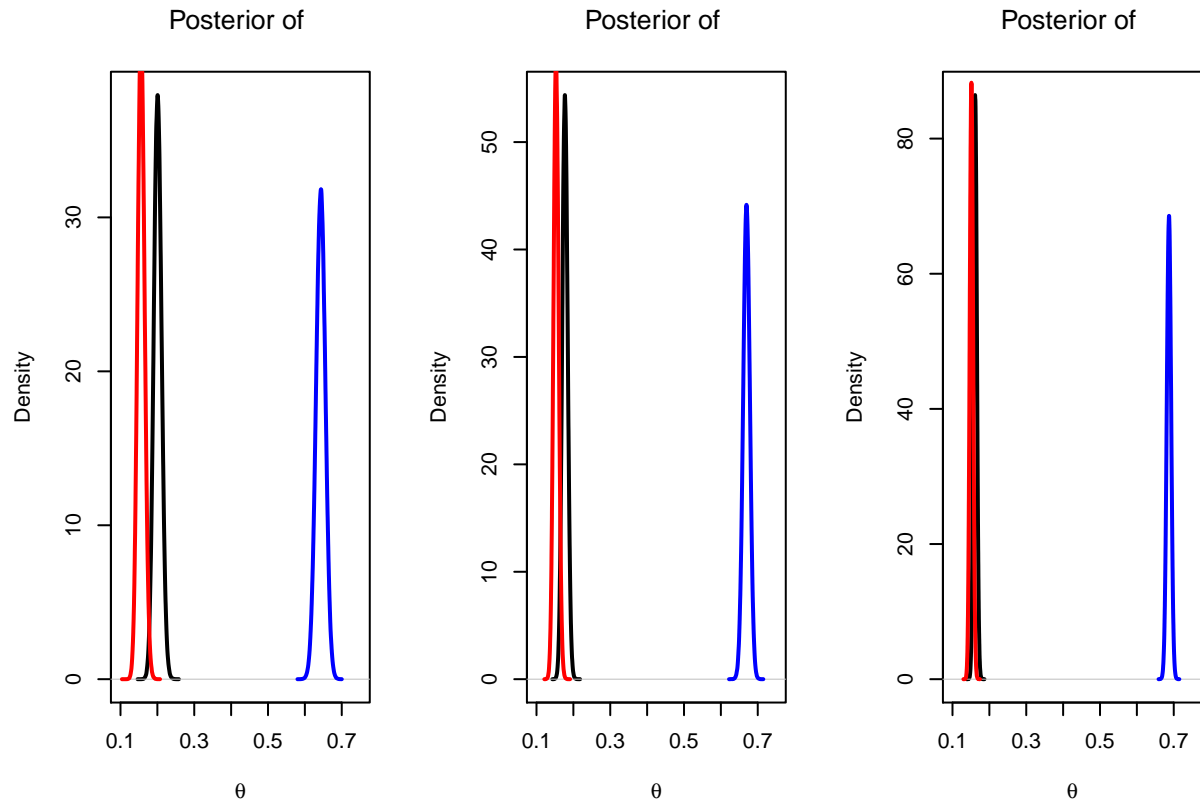
plot(density(post.alpha3[,1]), xlim = c(.10,.75), lwd = 2, main = expression("Posterior of ", theta), xlab = "theta", ylab = "density")
lines(density(post.alpha3[,2]), col = "red", lwd = 2)
lines(density(post.alpha3[,3]), col = "blue", lwd = 2)
```

```

lines(density(post.alpha2[,2]), col = "red", lwd = 2)
lines(density(post.alpha2[,3]), col = "blue", lwd = 2)

plot(density(post.alpha3[,1]), xlim = c(.10,.75), lwd = 2, main = expression("Posterior of ",theta), xlab = expression(theta), ylab = "Density")
lines(density(post.alpha3[,2]), col = "red", lwd = 2)
lines(density(post.alpha3[,3]), col = "blue", lwd = 2)

```



```

exp.mat = matrix(NA, nrow= 3,ncol = 3)
colnames(exp.mat) = colnames(prior.mat)
rownames(exp.mat) = rownames(prior.mat)
exp.mat[1,1:3] = apply(post.alpha1,2,mean)
exp.mat[2,1:3] =apply(post.alpha2,2,mean)
exp.mat[3,1:3]= apply(post.alpha3,2,mean)

quant.mat = matrix(NA,nrow = 3,ncol = 6)
rownames(quant.mat) = rownames(prior.mat)
quant.mat[1,1:6] = apply(post.alpha1,2, FUN = quantile, c(.025,.975))
quant.mat[2,1:6] = apply(post.alpha2,2, FUN = quantile, c(.025,.975))
quant.mat[3,1:6] = apply(post.alpha3,2, FUN = quantile, c(.025,.975))

kable(exp.mat)

```

	rel_weak	equal	strong
alpha_1	0.2009709	0.1560776	0.6429515
alpha_2	0.1772926	0.1531886	0.6695188

	rel_weak	equal	strong
alpha_3	0.1616243	0.1513851	0.6869905

```
kable(quant.mat)
```

alpha_1	0.1807612	0.2219870	0.1378481	0.1752130	0.6180870	0.6674183
alpha_2	0.1631100	0.1919227	0.1398398	0.1670192	0.6516638	0.6871992
alpha_3	0.1526826	0.1707797	0.1426814	0.1603030	0.6755291	0.6983516

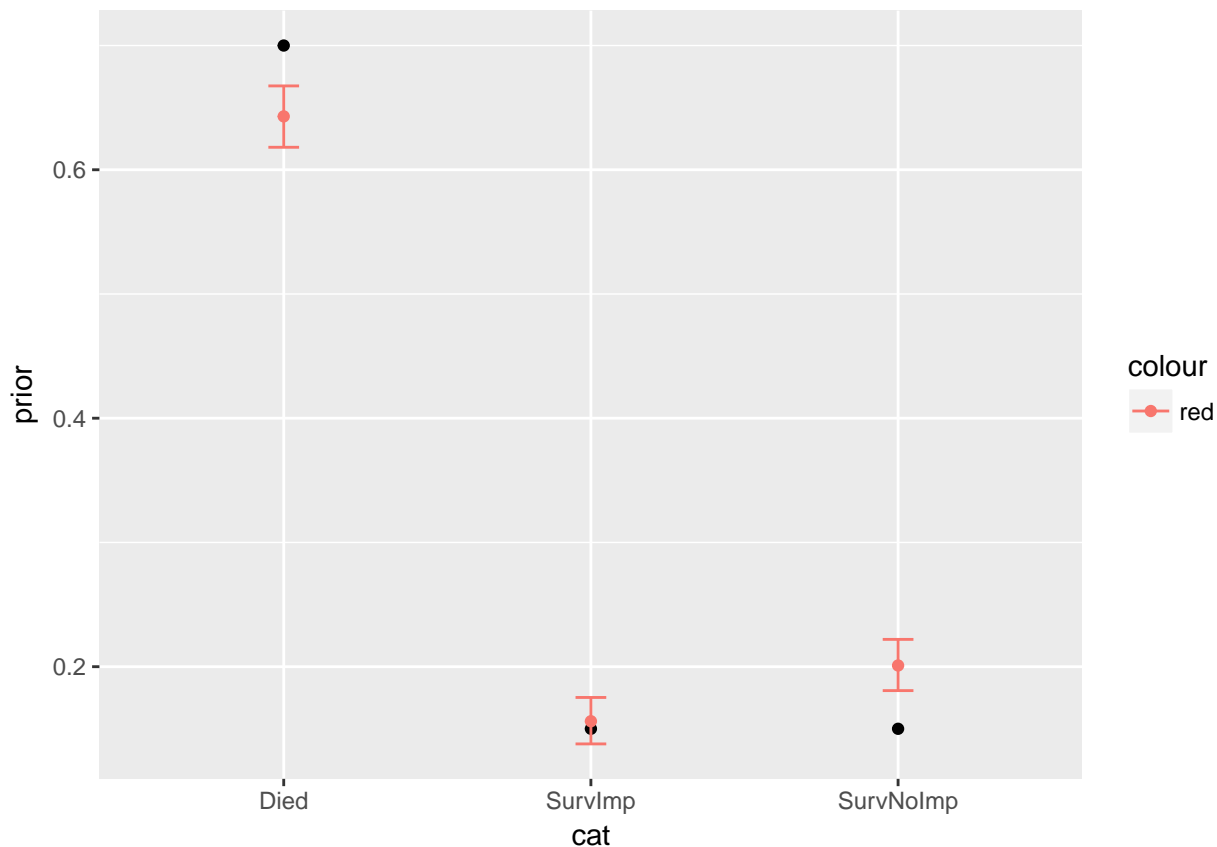
It seems clear that this does not represent historical rates and there is a difference. None of the historical rates are in the credible intervals for the data under any of my priors.

Comments from TA: -2. You should describe your methods and results. You present all your code and output. I need to check all your code to figure out what you intended to do. -1. Your graph is not informative. One may want to show that the observed data is against the null hypothesis. You only plot the posteriors under different priors but did not compare them with the null.

Re-do this with just one prior and compare it to the null hypothesis

```
n1 = 276
n2 = 211
n3 = 861
alpha1 = alpha2 = 15; alpha3 = 70
post.alpha1 = n1 + alpha1
post.alpha2 = n2 + alpha2
post.alpha3 = n3 + alpha3

n.reps = 1e06
post1.alpha = rdirichlet(n.reps,c(post.alpha1,post.alpha2,post.alpha3))
cred.int = apply(post1.alpha,2,FUN = quantile ,c(.025,.975))
#plot(c("SurvNoImp", "SurvImp", "Died"),c(.15,.15,.70),xlim = c("SurvNoImp", "SurvImp", "Died") ,ylab = )
d = data.frame(
  cat = c("SurvNoImp", "SurvImp", "Died"),
  prior = c(.15,.15,.70),
  low = cred.int[1,],
  upper = cred.int[2,],
  mean = apply(post1.alpha,2, FUN = mean)
)
ggplot(d,aes(x = cat,y =prior)) + geom_point() +
  geom_errorbar(width = .1,aes(ymin = low,ymax = upper, col = "red")) +
  geom_point(aes(x = cat,y = mean, col = "red"))
```



Now clearly, under this prior, when we compare it to the historic rates, it seems as the the data disprove Died and SurvNoImp, but it does not disprove SurvImp. The SurvImp credible interval does contain .15, which is the historic rate. However, the credible intervals for Died and SurvNoImp does not contain the historic rate.

2. Agresti 1.7 (12 points total)

Complete the given parts and add part f, given below. In a crossover trial comparing a new drug to a standard, π denotes the probability that the new one is judged better. It is desired to estimate π and to test $H_0 : \pi = 0.50$ against $H_a : \pi \neq 0.50$. In 20 independent observations, the new drug is better each time.

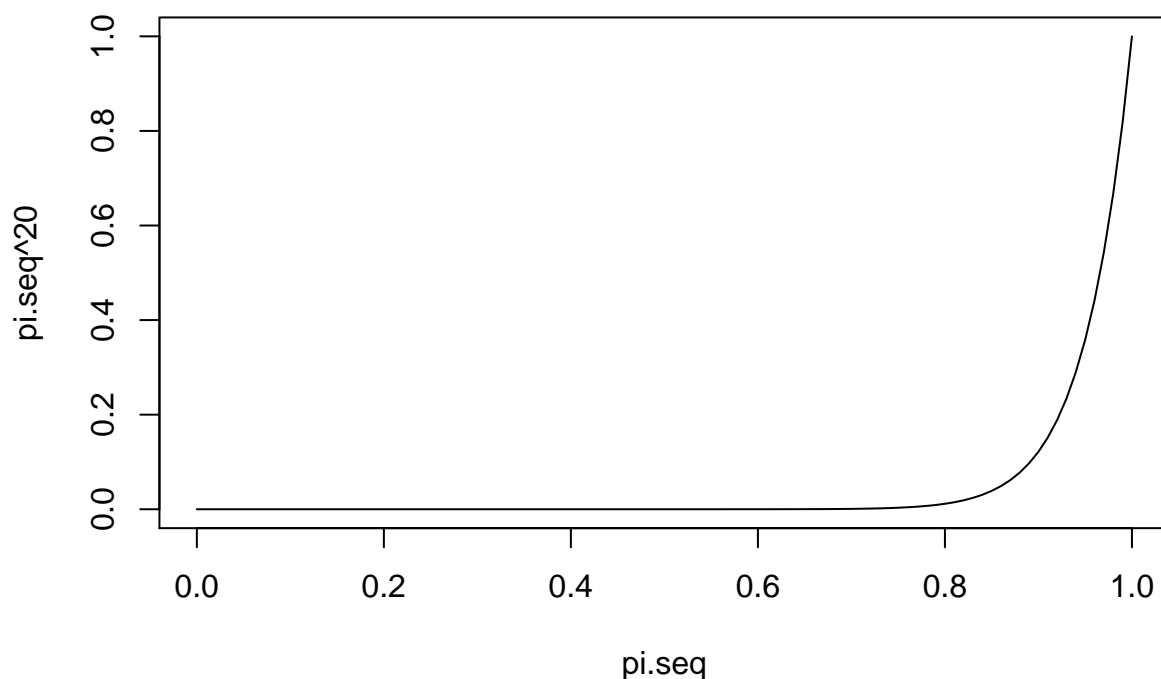
2.a. (2 points)

Plot the likelihood function. Is it close to the quadratic shape that large-sample normal approximations utilize?

Solution

The likelihood is π^{20} , which is not quadratic.

```
pi.seq = seq(0,1,by = .01)
plot(pi.seq,pi.seq^20, type = "l")
```



Comments -1. You plot x^2 but not $x^{(20)}$. obviously. Why did I do this?

This was a dumb mistake. I think I just never changed it.

2.b. (2 points)

Give the ML estimate of π . Conduct a Wald test and construct a 95% Wald CI for π . Are these sensible?

Solution

The ML estimate of $\pi = 1$, which is clearly not very sensible.

```
wald.est = (1 - .5) / sqrt(1*(1-1)/20)
ci = 1 + c(-1,1)*1.96*(sqrt(1*(1-1)/20))
wald.est
```

```
## [1] Inf
```

```
ci
```

```
## [1] 1 1
```

```
new.ci = c(ci[1],1)
```

Clearly the estimate and confidence interval are not sensible.

2.c. (2 points)

Conduct a score test, reporting the p-value. Construct a 95% score confidence interval. Interpret.

Solution

```
score.est = (1 - .5) / sqrt(.5*.5 / 20)
score.ci = 1 + c(-1,1)*1.96*sqrt(.5*.5 / 20)
(1 - pnorm(score.est))*2
```

```
## [1] 7.744216e-06
```

The p-value is extremely small. We are 95% confident that the true value for π is between 0.7808653, 1.2191347.

Comments: -1.5. This is a two-sided test and the p-value should be 2 times the p-value you reported. The method for calculating CI is wrong. When compute CI by inverting a test, you cannot assume a true value of the parameter. Check section 1.4.2 in the textbook or sample solution.

```
score.test = prop.test(x=20,n=20,correct = F)
score.test$p.value
```

```
## [1] 7.744216e-06
```

```
score.test$conf.int
```

```
## [1] 0.8388748 1.0000000
## attr("conf.level")
## [1] 0.95
```

2.d. (2 points)

Conduct a likelihood-ratio test and construct a likelihood-based 95% confidence interval. Interpret.

Solution

```
likel.test = 2*(20*log(2))
likel.test
```

```
## [1] 27.72589
```

```
like.ci = c(exp(-1.96^2 / 40) ,exp(1.96^2 / 40))
like.ci
```

```
## [1] 0.9084277 1.1008031
```

```
like.int = c(like.ci[1],1)
```

We are 95% confident that the true probability that a successive trial will be better than the previous is between (0.9084277, 1).

Comments: -0.5. Your final CI is correct but your derivation is not correct. It may not be a symmetric interval. Check section 1.4.2 in the textbook or sample solution.

```
exp(-qchisq(.95,1)/40)
```

```
## [1] 0.9084309
```

2.e.(2 points)

Construct an exact binomial test. Interpret.

Solution

```
binom.test(20,20)

##
## Exact binomial test
##
## data: 20 and 20
## number of successes = 20, number of trials = 20, p-value =
## 1.907e-06
## alternative hypothesis: true probability of success is not equal to 0.5
## 95 percent confidence interval:
## 0.8315665 1.0000000
## sample estimates:
## probability of success
## 1
p.val = binom.test(20,20)$p.value
p.val

## [1] 1.907349e-06
```

Under the exact binomial test, the probability of observing 20 consecutive tests is $1.907e - 06$, which is very low.

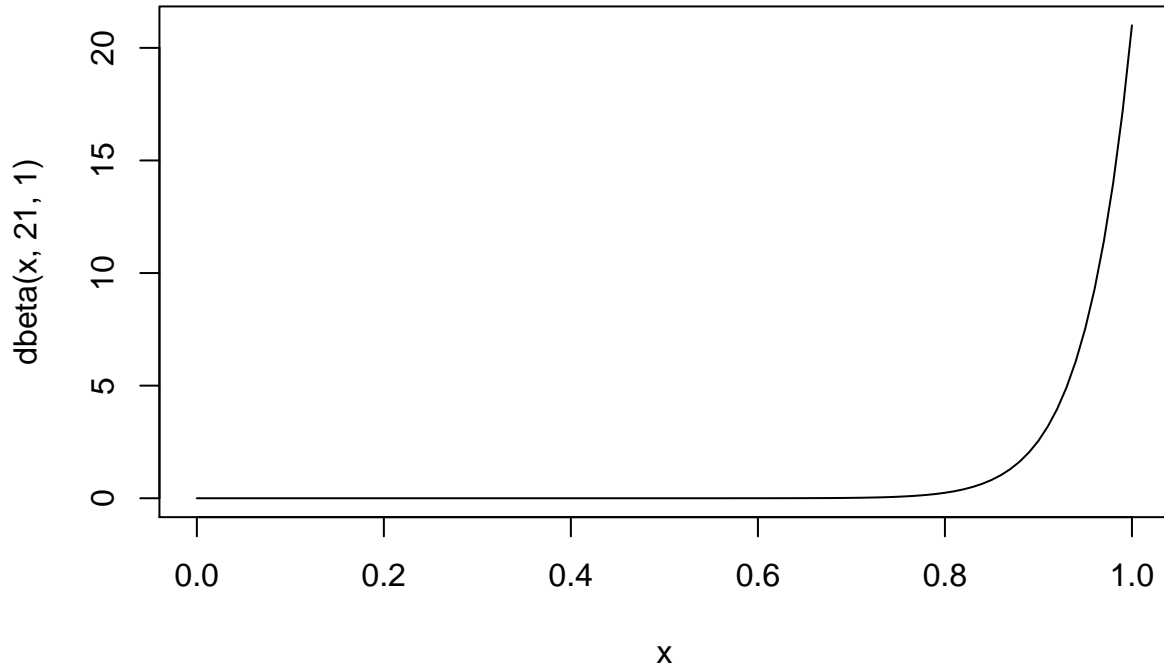
2.f. (2 points)

Estimate a 95% HPD interval for π when a $U[0,1]$ prior is used. Interpret.

Solution

The $U[0,1]$ prior is equivalent to a $\text{beta}(1,1)$ distribution, and thus $\pi|x \sim \text{beta}(x+1, n-x+1) = \text{beta}(21,1)$. If we look at the plot shown of this posterior distribution. It is clear that the HPD will involve 1 as the upper limit.

```
curve(dbeta(x,21,1))
```

```
int = betaHPD(21,1.000000001)
```

Because this is an HPD, we can speak probabilistically, which means that there is .95 probability that the probability the next trial is better than the first is between (0.8670541, 1).

3. Agresti 1.41 (10 points)

For the Dirichlet prior for multinomial probabilities, show the Bayesian estimator of the posterior expectation is a weighted average of the prior mean and sample proportions as in Agresti (1.19).

Solution

We know that $E(\theta_j) = \frac{\alpha_j}{\sum \alpha_k}$ and the sample proportion of a multinomial distribution is $\hat{\theta} = \frac{n_j}{n}$. We will use the following to show that the posterior mean $E(\theta_j | n_1, \dots, n_j) = \frac{n_j + \alpha_j}{n + \sum \alpha_k}$ is a weighted average of the above values.

$$\begin{aligned}
E(\theta_j | n_1, \dots, n_j) &= \frac{n_j + \alpha_j}{n + \sum \alpha_k} \\
&= w \frac{\alpha_j}{\sum \alpha_k} + (1 - w) \frac{n_j}{n} \\
&= \frac{\sum \alpha_k}{n + \sum \alpha_k} \frac{\alpha_j}{\sum \alpha_k} + \frac{n}{n + \sum \alpha_k} \frac{n_j}{n} \\
&= w \frac{\alpha_j}{\sum \alpha_k} + (1 - w) \frac{n_j}{n} \quad \text{where } w = \frac{\sum \alpha_k}{n + \sum \alpha_k}
\end{aligned}$$

4. Agresti 2.26 (10 points)

Show that the OR and RR need not be similar when π_i is close to 1.0 for both groups.

Solution

```

pi1 = .999
pi2 = .99
or = (pi1 * (1 - pi2)) / (pi2 * (1 - pi1))
rr = (pi1 / pi2)

med.pi1 = .499
med.pi2 = .49
or.med = (med.pi1 * (1 - med.pi2)) / (med.pi2 * (1 - med.pi1))
rr.med = med.pi1 / med.pi2

c(or, rr)

```

```
## [1] 10.090909 1.009091
```

```
c(or.med, rr.med)
```

```
## [1] 1.036661 1.018367
```

Although this isn't a formal proof or explanation yet, it is clear that when π_i is near 1, any difference between the values explodes. However, the same difference at medium values of π_i does not yield nearly as strong a difference, and the two estimates seem similar. But it is clear they are not similar near the boundary of $\pi = 1$. Also, it is obvious that where π_i is near 0, OR and RR will be very similar.

5. Agresti 2.28 (10 points)

In comparing new and standard treatments with success probabilities π_1 and π_2 , the number needed to treat (NNT) is the number of patients that would need to be treated with the new treatment instead of the standard in order for one patient to benefit. Explain why a natural estimate of this is $\frac{1}{\pi_1 - \pi_2}$.

Solution

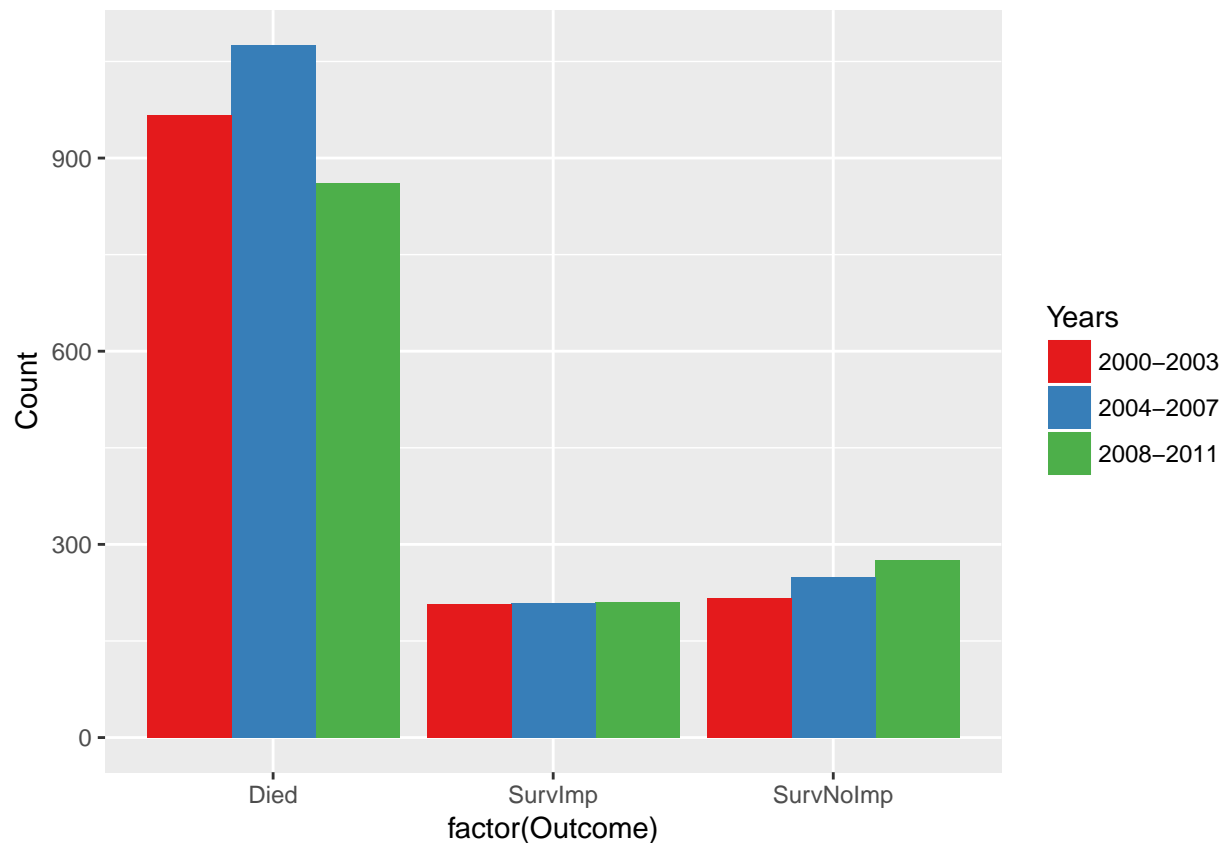
$\frac{1}{\pi_1 - \pi_2}$ is a natural estimate of this because NNT is the reciprocal of the risk difference, which is $\pi_1 - \pi_2$. The estimator for this risk difference is $\widehat{RR} = \hat{\pi}_1 - \hat{\pi}_2$, and thus it would make sense that the estimator for the reciprocal of this would be $\frac{1}{\widehat{RR}} = \frac{1}{\hat{\pi}_1 - \hat{\pi}_2}$.

Comments: -8. I did not see why using $1/RR$ is reasonable. If you think it is reasonable, you need to explain why it is reasonable.

Let n denote the number needed to treat and $\hat{\pi}_1$ and $\hat{\pi}_2$ are estimates of π_1 and π_2 respectively. Then if the standard treatments are used on n patients, we may expect $n\hat{\pi}_2$ patients recover. Similarly, we expect $n\hat{\pi}_1$ patients to recover using the new treatment. Therefore, if supposing π_1 is larger than π_2 , in order to get one more recovery patient, we need to have $n\hat{\pi}_1 - n\hat{\pi}_2 = 1$, which is $n = \frac{1}{\hat{\pi}_1 - \hat{\pi}_2}$.

6. Periviable infants (12 points total)

Recall the data on outcomes of periviable infants. Of 1348 infants in the study born at these gestational ages in 2008-2011, 861 died, 211 survived with neurodevelopmental impairment, and 276 survived without neurodevelopmental impairment. In 2000-2003, of 1391 infants born at these gestational ages, 217 survived with no impairment, 207 survived with impairment, and 967 died. In 2004-2007, of 1535 total infants, 250 survived without impairment, 209 survived with impairment, and 1076 died.



6.a. Nominal variables (6 points)

Provide a formal test of the hypothesis that time period is not associated with outcome. Interpret results of your test in clear language that could be used in a scientific publication. If there is a change over time, clearly describe the nature of that change.

Solution

We will use a χ^2 test for independence on the cells to test for the association between these nominal time variables.

```
tot.sum = sum(Neonate$Count)
row.sum = c(sum(Neonate$Count[Neonate$Years == "2000-2003"]),
             sum(Neonate$Count[Neonate$Years == "2004-2007"]),
             sum(Neonate$Count[Neonate$Years == "2008-2011"]))
row.sum.tab = rep(row.sum,c(3,3,3))
col.sum = c(sum(Neonate$Count[Neonate$Outcome == "SurvNoImp"]),
             sum(Neonate$Count[Neonate$Outcome == "SurvImp"]),
             sum(Neonate$Count[Neonate$Outcome == "Died"]))
col.sum.tab = rep(col.sum,3)
Neonate = cbind(Neonate,row.sum.tab,col.sum.tab)
obs.mat = matrix(Neonate$Count, nrow = 3, byrow = TRUE)
exp.counts = row.sum.tab / sum(Neonate$Count) * col.sum.tab / sum(Neonate$Count)
exp.mat = matrix(exp.counts,nrow = 3, byrow = TRUE)
colnames(exp.mat) = colnames(obs.mat) = c("SurvNoImp","SurvImp","Died")
rownames(exp.mat) = rownames(obs.mat) = c("2000-2003","2004-2007","2008-2011")
Neonate = cbind(Neonate,exp.counts)
test.chi = sum((Neonate$Count - Neonate$exp.counts)^2 / Neonate$exp.counts)
test.chi

## [1] 17.95398
1 - pchisq(test.chi , (3-1)*(3-1))

## [1] 0.00125992
```

According to our test statistic and subsequent p-value, it seems like there are some dependencies between these time intervals and the life outcome of the periviable neonates. LARGE SAMPLE. NONE are SPARSE

Comments: -2. You should describe the nature of the change.

The data provides evidence that time period is associated with outcome. However, we cannot tell how outcome is associated with time period from the test itself. ### 6.b. Ordinal variables (6 points) Now assume that the outcomes are ordered so that death is the worst outcome and survival without impairment is the best outcome. Conduct a linear trend test using scores (1,2,3) for time (1=2000-2003) and outcome (3=dead, 2=impaired, 1=survived and not impaired) and interpret the results. Do the results change if we use the outcome weights 100=dead, 10=impaired, 1=survived not impaired? If so, explain how and why the results differ. Comment on the appropriateness of the scores for both variables.

Solution

```
year.vec = c(rep(1,sum(obs.mat[1,1:3])),
             rep(2,sum(obs.mat[2,1:3])),
             rep(3,sum(obs.mat[3,1:3])))
)
out.vec = c(rep(c(1,2,3),(obs.mat[1,])),
            rep(c(1,2,3),(obs.mat[2,])),
            rep(c(1,2,3),(obs.mat[3,])))
)
cor.val = cor(year.vec,out.vec)
m2 = (cor.val^2) * (sum(Neonate$Count) -1)
cor.val

## [1] -0.05416482
```

```
m2
```

```
## [1] 12.53625
```

```
1 - pchisq(m2,1)
```

```
## [1] 0.0003991334
```

Above, we conduct the linear trend test, which is clearly significant as well. When we change the values of the outcome, from (1,2,3) to (1,10,100), the result is still significant, but less so. The p-value is 0.0012 as compared to 0.000399 from the previous.

```
year.vec = c(rep(1,sum(obs.mat[1,1:3])),
              rep(2,sum(obs.mat[2,1:3])),
              rep(3,sum(obs.mat[3,1:3]))
            )
out.vec = c(rep(c(1,10,100),(obs.mat[1,])),
            rep(c(1,10,100),(obs.mat[2,])),
            rep(c(1,10,100),(obs.mat[3,]))
          )
cor.val = cor(year.vec,out.vec)
m2 = (cor.val^2) * (sum(Neonate$Count) -1)
cor.val
```

```
## [1] -0.04948508
```

```
m2
```

```
## [1] 10.46361
```

```
1 - pchisq(m2,1)
```

```
## [1] 0.001217493
```

Comments: -2. You need to comment on the appropriateness of scores.

The choice of scores depends on the context. In some situations, these three outcomes or time periods may be considered to have equal distances and then our first choice of scores appears to be more appropriate. However, in some medical experiments, death may be considered much more serious than other two outcomes. Hence assigning a high score to death is more sensible. Therefore, the choice of scores can only be justified under specific context. All conclusions based on the analysis are valid under the specific choice of scores and may change if adopting other scores. When the trend is significant, it may be relatively robust to the choice of scores.

7. Oral contraceptives and thromboembolism (8 points)

We consider a retrospective matched pair case-control study of thromboembolism (formation of a blood clot that breaks loose and is carried by the bloodstream to plug another vessel) and oral contraceptive use. The cases were 175 women of reproductive age, discharged alive from 43 hospitals in 5 cities after experiencing idiopathic thrombophlebitis, pulmonary embolism, or cerebral thrombosis or embolism. Controls were females matched with their cases for hospital, residence, time of hospitalization, race, age, marital status, parity, and insurance pay status. The question of interest is whether cases are more likely than controls to use oral contraceptives.

Of the 175 matched pairs, in 10 pairs both cases and controls used oral contraceptives (OC's), in 95 pairs neither used OC's, in 13 pairs only the control used OC's, and in 57 pairs only the case used OC's.

Using the appropriate test, evaluate the null hypothesis that the proportion of cases exposed to OC's is the same as the proportion of controls who are exposed, and provide the p-value for this test. Calculate the OR

and exact 95% CI comparing the odds of thromboembolism in OC users to those in non-OC users. Interpret your results.

Solution

```
mcnem.res = mcnemar.exact(matrix(c(10,57,13,95),byrow = TRUE, nrow = 2))
mcnem.res

##
## Exact McNemar test (with central confidence intervals)
##
## data: matrix(c(10, 57, 13, 95), byrow = TRUE, nrow = 2)
## b = 57, c = 13, p-value = 1.029e-07
## alternative hypothesis: true odds ratio is not equal to 1
## 95 percent confidence interval:
##  2.371377 8.731311
## sample estimates:
## odds ratio
##  4.384615

mcnem.p = mcnem.res$p.value
mcnem.est = mcnem.res$estimate
mcnem.confint = mcnem.res$conf.int
```

To test the null hypothesis that the proportion of cases exposed to OC's is the same as the proportion of controls who are exposed, we will use McNemar's test for paired data. Using this test, we get the p-value of $(1.0289941 \times 10^{-7})$, which is significant. Also, we get an estimate of the odds-ratio of 4.3846154 with a confidence interval of (2.3713765, 8.7313106). This means that we are 95% confident that use of an oral contraceptive is associated with an 2.37 to 8.73 increase of odds of a thromboembolism.

8. Passive smoking and cancer risk (15 points)

A landmark study examined the association between passive smoking and cancer in a group of 509 individuals with cancer and 489 cancer-free (unmatched) controls. Researchers defined passive smoking as exposure to the cigarette smoke of a partner who smoked at least one cigarette per day for at least 6 months. One potential confounding variable is smoking by the participants (i.e., personal smoking) because personal smoking is related to both cancer risk and partner smoking. Therefore, it is important to consider personal smoking when examining the relationship between passive smoking and cancer risk.

Among cases, 120 were exposed to passive smoke but did not smoke themselves; 161 were smokers also exposed to passive smoke; 111 had no exposure from passive or active smoking; and 117 were active smokers who were not exposed to passive smoke. Among controls, 80 were exposed to passive smoke but did not smoke themselves; 130 were smokers also exposed to passive smoke; 155 had no exposure from passive or active smoking; and 124 were active smokers who were not exposed to passive smoke.

Calculate the conditional odds ratios comparing the odds of being a case by passive smoking exposure status, conditional on active smoking status (active smoker or nonsmoker). Evaluate whether case status and passive smoking are conditionally independent given active smoking status. Is the association between passive smoking and case status homogeneous across categories of active smoking status? Provide statistical evidence to support your answers.

Solution

```
smoker.mat = matrix(c(161,117,130,124),byrow= TRUE,nrow = 2)
non.smoker.mat = matrix(c(120,111,80,155),byrow = TRUE,nrow = 2)
colnames(smoker.mat) = colnames(non.smoker.mat) = c("Exposure","Not-exposed")
rownames(smoker.mat) = rownames(non.smoker.mat) = c("Case","Control")
smoker.mat
```

```
##           Exposure Not-exposed
## Case           161           117
## Control        130           124
```

```
non.smoker.mat
```

```
##           Exposure Not-exposed
## Case           120           111
## Control         80           155
```

To test the conditional independence of the the groups of cancer-free individuals and individuals with cancers, we will use the Cochran-Mantel-Haenszel test, which is defined as the following $\frac{[\sum_k (n_{11k} - \mu_{11k})]^2}{\sum_k \text{var}(n_{11k})}$, where $\mu_{11k} = \frac{n_{1+k}n_{1+k}}{n_k}$ and $\text{var}(n_{11k}) = \frac{n_{1+k}n_{1+k}n_{2+k}n_{2+k}}{n_k^2(n_k-1)}$. To begin, in order for this test to be valid, there must be unidirectionality among the odds-ratios. In our cases $OR_1 = \frac{(161)(124)}{(130)(117)} = 1.313$ and $OR_2 = \frac{(120)(155)}{(80)(111)} = 2.095$ are both greater than 1. Now note that $\mu_{11,1} = \frac{(278)(291)}{532} = 152.06$ and $\mu_{11,2} = \frac{(231)(200)}{466} = 99.14$. Also, the $\text{var}(n_{11,1}) = \frac{(278)(254)(291)(241)}{532^2(532-1)} = 32.95$ and $\text{var}(n_{11,2}) = \frac{(231)(235)(200)(266)}{466^2(466-1)} = 99.14$. So now using these values in the Cochran-Mantel-Haenszel test, the CSH = $\frac{[(161-152.06)+(120-99.14)]^2}{(32.95+28.6)} = 14.428$, which will follow a χ_1^2 . This means that the p-value according to this test is 0.000146. According to this, it seems like these are not conditionally independent because this would mean that assuming case status and pasive smoking are conditionally independent, the probability of obtaining a test statistics as extreme or more extreme between these control and case groups is .000146.

To test the homogeneous association, we will use the Breslow-Day test, which tests whether the odds-ratios between the different groups are equal, which would indicate homogeneity.

```
bres.array = abind(smoker.mat,non.smoker.mat,along = 3)
BreslowDayTest(bres.array)
```

```
##
## Breslow-Day test on Homogeneity of Odds Ratios
##
## data: bres.array
## X-squared = 3.2737, df = 1, p-value = 0.0704
```

```
BreslowDayTest(bres.array,correct = TRUE)
```

```
##
## Breslow-Day Test on Homogeneity of Odds Ratios (with Tarone
## correction)
##
## data: bres.array
## X-squared = 3.2736, df = 1, p-value = 0.0704
```

According to the Breslow-Day test without correction, the p-value is 0.074. According to this, there isn't evidence of non-homogeneity between the two groups of patients. We can't conclude that the groups aren't homogeneous. Thus, in this case, we can say that conditioning on smoking leads to conditional indepedence, but there isn't homogeneity between the smoking levels.

Comments: `mantelhaen.test()` can be used for CMH test.

```
mantelhaen.test(bres.array,correct = F)

##
## Mantel-Haenszel chi-squared test without continuity correction
##
## data:  bres.array
## Mantel-Haenszel X-squared = 14.422, df = 1, p-value = 0.0001461
## alternative hypothesis: true common odds ratio is not equal to 1
## 95 percent confidence interval:
##  1.263955 2.090024
## sample estimates:
## common odds ratio
##          1.625329
```

9. Exercise without pain! (15 points)

An athletics company produces a new model of running shoe that includes a harder material in the insert to correct for overpronation. The company is worried that the material will cause heel tenderness due to some loss of cushioning with the strike of each step. The company enrolled 87 runners and asked them about occasional heel tenderness at study entry and then again after using the new shoe for one month. The company would like to know whether the proportion of runners with occasional heel tenderness is the same before the study and after using the study shoe for one month.

Analyze the data and produce a professional report (as part of this document) providing statistical evidence that addresses the question of interest to the company. Data are in the file `dontbeaheel.xlsx`.

Solution

```
heel = read.xlsx("C:/Users/Zachary/Desktop/Fall_2017_PRojects/STA_841/STA_841_cat/HW01/dontbeaheel.xlsx")
table(heel$Occasion,heel$Pain)

##
##           No Yes
## After  53  34
## Before 63  24

before = heel$Pain[heel$Occasion == "Before"]
after = heel$Pain[heel$Occasion == "After"]
id = unique(heel$ID)
new.mat = as.data.frame(cbind(id,before,after))
table(new.mat$before,new.mat$after)

##
##           No Yes
## No   48  15
## Yes  5   19

cont.mat = matrix(c(19,5,15,48),byrow = TRUE, nrow = 2)
rownames(cont.mat) = colnames(cont.mat) = c("yes","no")

mcnem.res = mcnemar.exact(matrix(c(19,5,15,48),byrow = TRUE, nrow = 2))
mcnem.res
```



```
##
## Exact McNemar test (with central confidence intervals)
##
## data: matrix(c(19, 5, 15, 48), byrow = TRUE, nrow = 2)
## b = 5, c = 15, p-value = 0.04139
## alternative hypothesis: true odds ratio is not equal to 1
## 95 percent confidence interval:
## 0.0947764 0.9648136
## sample estimates:
## odds ratio
## 0.3333333

mcnem.p = mcnem.res$p.value
mcnem.est = mcnem.res$estimate
mcnem.confint = mcnem.res$conf.int
```

We seek to understand whether or not the newly produced running shoe causes heel tenderness in its users. This new shoe loses some cushioning previously models had, and we will analyze a study of 87 runners to understand the effect of this change in model. This design of this study is a classic design where we collect whether or not an individual experiences heel pain, both before and after they use the new model of shoe. Through this design, we have both a baseline and follow-up to understand how their pain can change based on these new shoes. Specifically, we will be interested in the discordant pairs, which are the individuals whose pain experience changes from before and after the shoe use (i.e. pain before and none after, no pain before but pain after). We will use McNemar's test to analyze these discordant pairs.

McNemar's test leverages these discordant pairs, which are the individuals of interest because it is not important to us if an individual's pain experience does not change. This would mean that these shoes do not have an effect on the individual for better or for worse. McNemar's test statistic $X^2 = \frac{(b-c)^2}{b+c}$ compares these discordant pairs and measures how far b is from c . If there is not a difference from before and after the new shoe use, then $b - c = 0$, which would mean $X^2 = 0$. When we perform this analysis, we know that 5 individual experienced pain before and no pain after while 15 experienced no pain before but pain after.

Using these numbers, we can find a p-value and a confidence interval for the odds-ratio. The p-value in our case is equal to 0.0413895. This means that we would reject the hypothesis that there is no difference from before and after the new shoe use. According to the results of the confidence interval, we are 95% confident that the new shoe is associated with 0.095 to 0.964 decreased odds of experiencing heel pain. This would indicate that the shoe is even associated with less heel pain, instead of more, which was initially suspected.

Comments: -3. Your explanation for odds ratio and CI is wrong. According to your formulation of data. Your odds ratio is actually the odds for getting heel pain before study is 0.333 as that after study. Therefore, a conclusion may be the new shoes may cause heel pain.

I have it reversed.