# Lecture 02: Statistical Inference for Binomial Parameters 

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## Inference for a probability

- Phase II cancer clinical trials are usually designed to see if a new, single treatment produces favorable results (proportion of success), when compared to a known, 'industry standard').
- If the new treatment produces good results, then further testing will be done in a Phase III study, in which patients will be randomized to the new treatment or the 'industry standard'.
- In particular, $n$ independent patients on the study are given just one treatment, and the outcome for each patient is usually

$$
Y_{i}=\left\{\begin{array}{l}
1 \text { if new treatment shrinks tumor (success) } \\
0 \text { if new treatment does not shrinks tumor (failure) }
\end{array}\right.
$$

$i=1, \ldots, n$

- For example, suppose $n=30$ subjects are given Polen Springs water, and the tumor shrinks in 5 subjects.
- The goal of the study is to estimate the probability of success, get a confidence interval for it, or perform a test about it.
- Suppose we are interested in testing

$$
H_{0}: p=.5
$$

where .5 is the probability of success on the "industry standard"
As discussed in the previous lecture, there are three ML approaches we can consider.

- Wald Test (non-null standard error)
- Score Test (null standard error)
- Likelihood Ratio test


## Wald Test

For the hypotheses

$$
\begin{aligned}
& H_{0}: p=p_{0} \\
& H_{A}: p \neq p_{0}
\end{aligned}
$$

The Wald statistic can be written as

$$
\begin{aligned}
z_{W} & =\frac{\widehat{p}-p_{0}}{S E} \\
& =\frac{\widehat{p}-p_{0}}{\sqrt{\widehat{p}(1-\widehat{p}) / n}}
\end{aligned}
$$

## Score Test

Agresti equations 1.8 and 1.9 yield

$$
\begin{aligned}
u\left(p_{0}\right) & =\frac{y}{p_{0}}-\frac{n-y}{1-p_{0}} \\
\iota\left(p_{0}\right) & =\frac{n}{p_{0}\left(1-p_{0}\right)} \\
z_{S} & =\frac{u\left(p_{0}\right)}{\left[\iota\left(p_{0}\right)\right]^{1 / 2}} \\
& =\frac{(\text { some algebra })}{\sqrt{p}-p_{0}} \\
& =\frac{\sqrt{p_{0}\left(1-p_{0}\right) / n}}{l}
\end{aligned}
$$

## Application of Wald and Score Tests

- Suppose we are interested in testing

$$
\mathrm{H}_{0}: p=.5
$$

- Suppose $Y=2$ and $n=10$ so $\widehat{p}=.2$
- Then,

$$
Z_{W}=\frac{(.2-.5)}{\sqrt{.2(1-.8) / 10}}=-2.37171
$$

and

$$
Z_{S}=\frac{(.2-.5)}{\sqrt{.5(1-.5) / 10}}=-1.89737
$$

- Here, $Z_{W}>Z_{S}$ and at the $\alpha=0.05$ level, the statistical conclusion would differ.


## Notes about $Z_{W}$ and $Z_{S}$

- Under the null, $Z_{W}$ and $Z_{S}$ are both approximately $N(0,1)$. However, $Z_{S}$ 's sampling distribution is closer to the standard normal than $Z_{W}$ so it is generally preferred.
- When testing

$$
\begin{gathered}
\mathrm{H}_{0}: p=.5 \\
\left|Z_{W}\right| \geq\left|Z_{S}\right|
\end{gathered}
$$

i.e.,

$$
\left|\frac{(\widehat{p}-.5)}{\sqrt{\widehat{p}(1-\widehat{p}) / n}}\right| \geq\left|\frac{(\widehat{p}-.5)}{\sqrt{.5(1-.5) / n}}\right|
$$

- Why ? Note that

$$
\widehat{p}(1-\widehat{p}) \leq .5(1-.5),
$$

i.e., $p(1-p)$ takes on its maximum value at $p=.5$ :

| p | .10 | .20 | .30 | .40 | .50 | .60 | .70 | .80 | .90 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| $\mathrm{p}(1-\mathrm{p})$ | .09 | .16 | .21 | .24 | .25 | .24 | .21 | .16 | .09 |

- Since the denominator of $Z_{W}$ is always less than the denominator of $Z_{S},\left|Z_{W}\right| \geq\left|Z_{S}\right|$
- Under the null, $p=.5$,

$$
\widehat{p}(1-\widehat{p}) \approx .5(1-.5)
$$

so

$$
\left|Z_{S}\right| \approx\left|Z_{W}\right|
$$

- However, under the alternative,

$$
\mathrm{H}_{A}: p \neq .5
$$

$Z_{S}$ and $Z_{W}$ could be very different, and, since

$$
\left|Z_{W}\right| \geq\left|Z_{S}\right|
$$

the test based on $Z_{W}$ is more powerful (when testing against a null of 0.5 ).

- For the general test

$$
\mathrm{H}_{0}: p=p_{o},
$$

for a specified value $p_{o}$, the two test statistics are

$$
Z_{S}=\frac{\left(\widehat{p}-p_{o}\right)}{\sqrt{p_{o}\left(1-p_{o}\right) / n}}
$$

and

$$
Z_{W}=\frac{\left(\widehat{p}-p_{o}\right)}{\sqrt{\widehat{p}(1-\widehat{p}) / n}}
$$

- For this general test, there is no strict rule that

$$
\left|Z_{W}\right| \geq\left|Z_{S}\right|
$$

## Likelihood-Ratio Test

- It can be shown that

$$
2 \log \left\{\frac{L\left(\widehat{p} \mid \mathrm{H}_{A}\right)}{L\left(p_{o} \mid \mathrm{H}_{0}\right)}\right\}=2\left[\log L\left(\widehat{p} \mid \mathrm{H}_{A}\right)-\log L\left(p_{o} \mid \mathrm{H}_{0}\right)\right] \sim \chi_{1}^{2}
$$

where

$$
L\left(\widehat{p} \mid \mathrm{H}_{A}\right)
$$

is the likelihood after replacing $p$ by its estimate, $\widehat{p}$, under the alternative $\left(\mathrm{H}_{A}\right)$, and

$$
L\left(p_{o} \mid \mathrm{H}_{0}\right)
$$

is the likelihood after replacing $p$ by its specified value, $p_{o}$, under the null $\left(\mathrm{H}_{0}\right)$.

## Likelihood Ratio for Binomial Data

- For the binomial, recall that the log-likelihood equals

$$
\log L(p)=\log \binom{n}{y}+y \log p+(n-y) \log (1-p)
$$

- Suppose we are interested in testing

$$
\mathrm{H}_{0}: p=.5 \quad \text { versus } \quad \mathrm{H}_{0}: p \neq .5
$$

- The likelihood ratio statistic generally only is for a two-sided alternative (recall it is $\chi^{2}$ based)
- Under the alternative,

$$
\log L\left(\widehat{p} \mid \mathrm{H}_{A}\right)=\log \binom{n}{y}+y \log \widehat{p}+(n-y) \log (1-\widehat{p}),
$$

- Under the null,

$$
\log L\left(.5 \mid \mathrm{H}_{0}\right)=\log \binom{n}{y}+y \log .5+(n-y) \log (1-.5)
$$

Then, the likelihood ratio statistic is

$$
\begin{aligned}
2\left[\log L\left(\widehat{p} \mid \mathrm{H}_{A}\right)-\log L\left(p_{o} \mid \mathrm{H}_{0}\right)\right]= & 2\left[\log \binom{n}{y}+y \log \hat{p}+(n-y) \log (1-\widehat{p})\right] \\
& -2\left[\log \binom{n}{y}+y \log .5+(n-y) \log (1-.5)\right] \\
= & 2\left[y \log \left(\frac{\widehat{p}}{.5}\right)+(n-y) \log \left(\frac{1-\widehat{p}}{1-.5}\right)\right] \\
= & 2\left[y \log \left(\frac{y}{.5 n}\right)+(n-y) \log \left(\frac{n-y}{(1-.5) n}\right)\right]
\end{aligned}
$$

which is approximately $\chi_{1}^{2}$

## Example

- Recall from previous example, $Y=2$ and $n=10$ so $\widehat{p}=.2$
- Then, the Likelihood Ratio Statistic is

$$
2\left[2 \log \left(\frac{.2}{.5}\right)+(8) \log \left(\frac{.8}{.5}\right)\right]=3.85490(p=0.049601)
$$

- Recall, both $Z_{W}$ and $Z_{S}$ are $\mathrm{N}(0,1)$, and the square of a $\mathrm{N}(0,1)$ is a chi-square 1 df .
- Then, the Likelihood ratio statistic is on the same scale as $Z_{W}^{2}$ and $Z_{S}^{2}$, since both $Z_{W}^{2}$ and $Z_{S}^{2}$ are chi-square 1 df
- For this example

$$
Z_{S}^{2}=\left[\frac{(.2-.5)}{\sqrt{.5(1-.5) / 10}}\right]^{2}=3.6
$$

and

$$
Z_{W}^{2}=\left[\frac{(.2-.5)}{\sqrt{.2(1-.8) / 10}}\right]^{2}=5.625
$$

- The Likelihood Ratio Statistic is between $Z_{S}^{2}$ and $Z_{W}^{2}$.


## Likelihood Ratio Statistic

For the general test

$$
\mathrm{H}_{0}: p=p_{o},
$$

the Likelihood Ratio Statistic is

$$
2\left[y \log \left(\frac{\widehat{p}}{p_{o}}\right)+(n-y) \log \left(\frac{1-\widehat{p}}{1-p_{o}}\right)\right] \sim \chi_{1}^{2}
$$

asymptotically under the null.

## Large Sample Confidence Intervals

- In large samples, since

$$
\widehat{p} \sim N\left(p, \frac{p(1-p)}{n}\right),
$$

we can obtain a $95 \%$ confidence interval for $p$ with

$$
\widehat{p} \pm 1.96 \sqrt{\frac{\widehat{p}(1-\widehat{p})}{n}}
$$

- However, since $0 \leq p \leq 1$, we would want the endpoints of the confidence interval to be in $[0,1]$, but the endpoints of this confidence interval are not restricted to be in $[0,1]$.
- When $p$ is close to 0 or 1 (so that $\widehat{p}$ will usually be close to 0 or 1 ), and/or in small samples, we could get endpoints outside of $[0,1]$. The solution would be the truncate the interval endpoint at 0 or 1.


## Example

- Suppose $n=10$, and $Y=1$, then

$$
\widehat{p}=\frac{1}{10}=.1
$$

and the $95 \%$ confidence interval is

$$
\begin{gathered}
\widehat{p} \pm 1.96 \sqrt{\frac{\widehat{p}(1-\widehat{p})}{n}}, \\
.1 \pm 1.96 \sqrt{\frac{.1(1-.1)}{10}}, \\
{[-.086, .2867]}
\end{gathered}
$$

- After truncating, you get,

$$
[0, .2867]
$$

## Exact Test Statistics and Confidence Intervals

Unfortunately, many of the phase II trails have small samples, and the above asymptotic test statistics and confidence intervals have very poor properties in small samples. (A 95\% confidence interval may only have $80 \%$ coverage-See Figure 1.3 in Agresti).

In this situation, 'Exact test statistics and Confidence Intervals' can be obtained.

## One-sided Exact Test Statistic

- The historical norm for the clinical trial you are doing is $50 \%$, so you want to test if the response rate of the new treatment is greater then $50 \%$.
- In general, you want to test

$$
\mathrm{H}_{0}: p=p_{o}=0.5
$$

versus

$$
\mathrm{H}_{A}: p>p_{o}=0.5
$$

- The test statistic

$$
Y=\text { the number of successes out of } n \text { trials }
$$

Suppose you observe $y_{o b s}$ successes ;

Under the null hypothesis,

$$
n \widehat{p}=Y \sim \operatorname{Bin}\left(n, p_{o}\right),
$$

i.e.,

$$
P\left(Y=y \mid \mathrm{H}_{0}: p=p_{o}\right)=\binom{n}{y} p_{o}^{y}\left(1-p_{o}\right)^{n-y}
$$

- When would you tend to reject $\mathrm{H}_{0}: p=p_{o}$ in favor of $\mathrm{H}_{A}: p>p_{o}$


## Answer

Under $\mathrm{H}_{0}: p=p_{o}$, you would expect $\widehat{p} \approx p_{o}$ ( $Y \approx n p_{o}$ )
Under $\mathrm{H}_{A}: p>p_{o}$, you would expect $\widehat{p}>p_{o}$
( $Y>n p_{o}$ )
i.e., you would expect $Y$ to be 'large' under the alternative.

## Exact one-sided $p$-value

- If you observe $y_{o b s}$ successes, the exact one-sided $p$-value is the probability of getting the observed $y_{o b s}$ plus any larger (more extreme) $Y$

$$
\begin{aligned}
p-\text { value } & =P\left(Y \geq y_{o b s} \mid \mathrm{H}_{0}: p=p_{o}\right) \\
& =\sum_{j=y_{o b s}}^{n}\binom{n}{j} p_{o}^{j}\left(1-p_{o}\right)^{n-j}
\end{aligned}
$$

## Other one-sided exact $p$-value

- You want to test

$$
\mathrm{H}_{0}: p=p_{o}
$$

versus

$$
\mathrm{H}_{A}: p<p_{o}
$$

- The exact $p$-value is the probability of getting the observed $y_{o b s}$ plus any smaller (more extreme) $y$

$$
\begin{aligned}
p-\text { value } & =P\left(Y \leq y_{o b s} \mid \mathrm{H}_{0}: p=p_{o}\right) \\
& =\sum_{j=0}^{y_{o b s}}\binom{n}{j} p_{o}^{j}\left(1-p_{o}\right)^{n-j}
\end{aligned}
$$

## Two-sided exact $p$-value

- The general definition of a 2 -sided exact $p$-value is

$$
P\left[\begin{array}{l|l}
\text { seeing a result as likely or } \\
\text { less likely than the observed result }
\end{array} \quad \mathrm{H}_{0}\right]
$$

It is easy to calculate a 2 -sided $p$-value for a symmetric distribution, such as $Z \sim N(0,1)$. Suppose you observe $z>0$,


## Symmetric distributions

- If the distribution is symmetric with mean 0 , e.g., normal, then the exact 2 -sided $p$-value is

$$
p-\text { value }=2 \cdot P(Z \geq|z|)
$$

when $z$ is positive or negative.

- In general, if the distribution is symmetric, but not necessarily centered at 0 , then the exact 2 -sided $p$-value is

$$
p-\text { value }=2 \cdot \min \left\{P\left(Y \geq y_{o b s}\right), P\left(Y \leq y_{o b s}\right)\right\}
$$

- Now, consider a symmetric binomial. For example, suppose $n=4$ and $p_{o}=.5$, then,

```
Binomial PDF for N=4 and P=0.5
Number of
\begin{tabular}{cccc} 
Successes & \(P(Y=y)\) & \(P(Y<=y)\) & \(P(Y>=y)\) \\
0 & 0.0625 & 0.0625 & 1.0000 \\
1 & 0.2500 & 0.3125 & 0.9375 \\
2 & 0.3750 & 0.6875 & 0.6875 \\
3 & 0.2500 & 0.9375 & 0.3125 \\
4 & 0.0625 & 1.0000 & 0.0625
\end{tabular}
```

Suppose you observed $y_{o b s}=4$, then the exact two-sided $p$-value would be

$$
\begin{aligned}
p-\text { value } & =2 \cdot \min \left\{P\left(Y \geq y_{o b s}\right), P\left(Y \leq y_{o b s}\right)\right\} \\
& =2 \cdot \min \{P(Y \geq 4), P(Y \leq 4)\} \\
& =2 \cdot \min \{.0625,1\} \\
& =2(.0625) \\
& =.125
\end{aligned}
$$

- The two-sided exact $p$-value is trickier when the binomial distribution is not symmetric
- For the binomial data, the exact 2 -sided $p$-value is

- Essentially the sum of all probabilities such that $P\left(Y=y \mid P_{0}\right) \leq P\left(y_{o b s} \mid P_{0}\right)$


## In general, to calculate the 2 -sided $p$-value

1. Calculate the probability of the observed result under the null

$$
\pi=P\left(Y=y_{o b s} \mid p=p_{o}\right)=\binom{n}{y_{o b s}} p_{o}^{y_{o b s}}\left(1-p_{o}\right)^{n-y_{o b s}}
$$

2. Calculate the probabilities of all $n+1$ values that $Y$ can take on:

$$
\pi_{j}=P\left(Y=j \mid p=p_{o}\right)=\binom{n}{j} p_{o}^{j}\left(1-p_{o}\right)^{n-j}
$$

$j=0, \ldots, n$.
3. Sum the probabilities $\pi_{j}$ in (2.) that are less than or equal to the observed probability $\pi$ in (1.)

$$
\begin{gathered}
p-\text { value }=\sum_{j=0}^{n} \pi_{j} I\left(\pi_{j} \leq \pi\right) \text { where } \\
I\left(\pi_{j} \leq \pi\right)=\left\{\begin{array}{l}
1 \text { if } \pi_{j} \leq \pi \\
0 \text { if } \pi_{j}>\pi
\end{array}\right.
\end{gathered}
$$

## Example

- Suppose $n=5$, you hypothesize $p=.4$ and we observe $y=3$ successes.
- Then, the PMF for this binomial is

| Binomial PMF (probability mass function) for $\mathrm{N}=5$ and $\mathrm{P}=0.4$ |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: |
| Number of |  |  |  |  |
| Successes | $\mathrm{P}(\mathrm{Y}=\mathrm{y})$ | $\mathrm{P}(\mathrm{Y}<=\mathrm{y})$ | $P(Y>=y)$ |  |
| 0 | 0.07776 | 0.07776 | 1.00000 |  |
| 1 | 0.25920 | 0.33696 | 0.92224 |  |
| 2 | 0.34560 | 0.68256 | 0.66304 |  |
| 3 | 0.23040 | 0.91296 | 0.31744 | <----Y obs |
| 4 | 0.07680 | 0.98976 | 0.08704 |  |
| 5 | 0.01024 | 1.00000 | 0.01024 |  |

## Exact P-Value by Hand

- Step 1: Determine $P\left(Y=3 \mid n=5, P_{0}=.4\right)$. In this case $P(Y=3)=.2304$.
- Step 2: Calculate Table (see previous slide)
- Step 3: Sum probabilities less than or equal to the one observed in step 1. When $Y \in\{0,3,4,5\}, P(Y) \leq 0.2304$.

| ALTERNATIVE | EXACT | PROBS |
| :--- | :--- | :--- |
|  |  |  |
| $\mathrm{H}_{A}: p>.4$ | .317 | $P[Y \geq 3]$ |
| $\mathrm{H}_{A}: p<.4$ | .913 | $P[Y \leq 3]$ |
|  |  |  |
| $\mathrm{H}_{A}: p \neq .4$ | .395 | $P[Y \geq 3]+$ |
|  |  | $P[Y=0]$ |

## Comparison to Large Sample Inference

Note that the exact and asymptotic do not agree very well:

|  |  | LARGE |
| :--- | :--- | :--- |
| ALTERNATIVE | EXACT | SAMPLE |
| $\mathrm{H}_{A}: p>.4$ | .317 | .181 |
| $\mathrm{H}_{A}: p<.4$ | .913 | .819 |
| $\mathrm{H}_{A}: p \neq .4$ | .395 | .361 |

## Calculations by Computer

We will look at calculations by

1. STATA (best)
2. $R$ (good)
3. SAS (surprisingly poor)

## STATA

The following STATA code will calculate the exact $p$-value for you

```
From within STATA at the dot, type
bitesti 5 3 .4
----------Output------------
```



To perform an exact binomial test, you have to use binom.test function at the R prompt as below

```
> binom.test(3,5,0.5,alternative="two.sided",conf.level=0.95) # R
code
and the output looks like
```

```
    Exact binomial test
```

    Exact binomial test
    data: 3 and 5, number of successes = 3, number of trials = 5,
data: 3 and 5, number of successes = 3, number of trials = 5,
p-value = 1 alternative hypothesis: true probability of success is
p-value = 1 alternative hypothesis: true probability of success is
not equal to 0.5 95 percent confidence interval: 0.1466328
not equal to 0.5 95 percent confidence interval: 0.1466328
0.9472550 sample estimates: probability of success 0.6

```
0.9472550 sample estimates: probability of success 0.6
```

This gets a score of good since the output is not as descriptive as the STATA output.

## SAS

```
Unfortunately, SAS Proc Freq gives the wrong 2-sided p-value
data one;
    input outcome $ count;
cards;
1succ 3
2fail 2
;
proc freq data=one;
    tables outcome / binomial(p=.4);
    weight count;
    exact binomial;
run;
```

```
---------Output-----------
Binomial Proportion
for outcome = 1succ
    Test of HO: Proportion = 0.4
\begin{tabular}{ll} 
ASE under HO & 0.2191 \\
\(Z\) & 0.9129 \\
One-sided \(\operatorname{Pr}>\mathrm{Z}\) & 0.1807 \\
Two-sided \(\operatorname{Pr}>|\mathrm{Z}|\) & 0.3613
\end{tabular}
Exact Test
One-sided Pr >= P 0.3174
Two-sided = 2 * One-sided 0.6349
Sample Size = 5
```


## Better Approximation using the normal distribution

- Because $Y$ is discrete, a 'continuity-correction' is often applied to the normal approximation to more closely approximate the exact $p$-value.
- To make a discrete distribution look more approximately continuous, the probability function is drawn such that $P(Y=y)$ is a rectangle centered at $y$ with width 1 , and height $P(Y=y)$, i.e.,
- The area under the curve between $y-0.5$ and $y+0.5$ equals

$$
[(y+0.5)-(y-0.5)] \cdot P(Y=y)=1 \cdot P(Y=y)
$$

For example, suppose as before, we have $n=5$ and $p_{o}=.4$,.

Then on the probability curve,

$$
P(Y \geq y) \approx P(Y \geq y-.5)
$$

which, using the continuity corrected normal approximation is

$$
P\left(\left.Z \geq \frac{(y-.5)-n p_{o}}{\sqrt{n p_{o}\left(1-p_{o}\right)}} \right\rvert\, \mathrm{H}_{0}: p=p_{o} ; Z \sim N(0,1)\right)
$$

and

$$
P(Y \leq y) \approx P(Y \leq y+.5)
$$

which, using the continuity corrected normal approximation

$$
P\left(\left.Z \leq \frac{(y+.5)-n p_{o}}{\sqrt{n p_{o}\left(1-p_{o}\right)}} \right\rvert\, \mathrm{H}_{0}: p=p_{o} ; Z \sim N(0,1)\right)
$$

With the continuity correction, the above $p$-values become

| ALTERNATIVE | EXACT | CARGE <br> SAMPLE | Corrected <br> LARGE |
| ---: | ---: | ---: | ---: |
| SAMPLE |  |  |  |

Then, even with the small sample size of $n=5$, the continuity correction does a good job of approximating the exact $p$-value.

Also, as $n \rightarrow \infty$, the exact and asymptotic are equivalent under the null; so for large $n$, you might as well use the asymptotic.

However, given the computational power available, you can easily calculate the exact p -value.

## Exact Confidence Interval

A $(1-\alpha)$ confidence interval for $p$ is of the form

$$
\left[p_{L}, p_{U}\right],
$$

where $p_{L}$ and $p_{U}$ are random variables such that

$$
P\left[p_{L} \leq p \leq p_{U}\right]=1-\alpha
$$

For example, for a large sample $95 \%$ confidence interval,

$$
p_{L}=\widehat{p}-1.96 \sqrt{\frac{\widehat{p}(1-\widehat{p})}{n}},
$$

and

$$
p_{U}=\widehat{p}+1.96 \sqrt{\frac{\widehat{p}(1-\widehat{p})}{n}},
$$

It's kind of complicated, but it can be shown that, to obtain a $95 \%$ exact confidence interval

$$
\left[p_{L}, p_{U}\right]
$$

the endpoints $p_{L}$ and $p_{U}$ satisfy

$$
\begin{aligned}
\alpha / 2=.025 & =P\left(Y \geq y_{o b s} \mid p=p_{L}\right) \\
& =\sum_{j=y_{o b s}}^{n}\binom{n}{j} p_{L}^{j}\left(1-p_{L}\right)^{n-j},
\end{aligned}
$$

and

$$
\begin{aligned}
\alpha / 2=.025 & =P\left(Y \leq y_{o b s} \mid p=p_{U}\right) \\
& \left.=\sum_{j=0}^{y_{o b s}}\binom{n}{j} p_{U}^{j}\left(1-p_{U}\right)\right)^{n-j}
\end{aligned}
$$

In these formulas, we know $\alpha / 2=.025$ and we know $y_{o b s}$ and $n$. Then, we solve for the unknowns $p_{L}$ and $p_{U}$.
Can figure out $p_{L}$ and $p_{U}$ by plugging different values for $p_{L}$ and $p_{U}$ until we find the values that make $\alpha / 2=.025$

- Luckily, this is implemented on the computer, so we don't have to do it by hand.
- Because of relationship between hypothesis testing and confidence intervals, to calculate the exact confidence interval, we are actually setting the exact one-sided $p$-values to $\alpha / 2$ for testing $\mathrm{H}_{o}: p=p_{o}$ and solving for $p_{L}$ and $p_{U}$.
- In particular, we find $p_{L}$ and $p_{U}$ to make these $p$-values equal to $\alpha / 2$.


## Example

- Suppose $n=5$ and $y_{o b s}=4$, and we want a $95 \%$ confidence interval. ( $\alpha=.05$, $\alpha / 2=.025)$.
- Then, the lower point, $p_{L}$ of the exact confidence interval $\left[p_{L}, p_{U}\right]$ is the value $p_{L}$ such that

$$
\alpha / 2=.025=P\left[Y \geq 4 \mid p=p_{L}\right]=\sum_{j=4}^{5}\binom{5}{j} p_{L}^{j}\left(1-p_{L}\right)^{n-j},
$$

- If you don't have a computer program to do this, you can try "trial" and error for $p_{L}$

$$
\begin{array}{ll}
p_{L} & P\left(Y \geq 4 \mid p=p_{L}\right) \\
.240 & 0.013404 \\
.275 & 0.022305 \\
.2836 & .025006^{*} \approx .025
\end{array}
$$

- Then, $p_{L} \approx .2836$.
- Similarly, the upper point, $p_{U}$ of the exact confidence interval $\left[p_{L}, p_{U}\right.$ ] is the value $p_{U}$ such that

$$
\alpha / 2=.025=P\left[Y \leq 4 \mid p=p_{U}\right]=\sum_{j=0}^{4}\binom{5}{j} p_{U}^{j}\left(1-p_{U}\right)^{n-j},
$$

- Similarly, you can try "trial" and error for the $p_{U}$

| $p_{U}$ | $P\left(Y \leq 4 \mid p=p_{U}\right)$ |
| :--- | :--- |
| .95 | 0.22622 |
| .99 | 0.049010 |
| .994944 | $0.025026^{*} \approx .025$ |

## Using STATA

The following STATA code will calculate the exact binomial confidence interval for you

- cii 54



## Using SAS

```
data one;
    input outcome $ count;
cards;
1succ 4
2fail 1
;
proc freq data=one;
    tables outcome / binomial;
    weight count;
run;
```


## Binomial Proportion

```
Proportion 0.8000
ASE 0.1789
95% Lower Conf Limit 0.4494
95% Upper Conf Limit 1.0000
Exact Conf Limits
95% Lower Conf Limit 0.2836
95% Upper Conf Limit 0.9949
Test of HO: Proportion = 0.5
ASE under HO 0.2236
Z 1.3416
One-sided Pr > Z 0.0899
Two-sided Pr > |Z| 0.1797
Sample Size = 5
```


## Comparing the exact and large sample

- Then, the two sided confidence intervals are

> LARGE
> SAMPLE
> (NORMAL)
EXACT
[.449,1]

- We had to truncate the upper limit based on using $\widehat{p}$ at 1 .
- The exact Cl is not symmetric about $\widehat{p}=\frac{4}{5}=.8$, whereas the the confidence interval based on $\widehat{p}$ would be if not truncated.
- Suggestion; if $Y<5$, and/or $n<30$, use exact; for large $Y$ and $n$, you can use whatever you like, it is expected that they would be almost identical.


## Exact limits based on F Distribution

- While software would be the tool of choice (I doubt anyone still calculates exact binomial confidence limits by hand), there is a distributional relationship among the Binomial and F distributions.
- In particular $P_{L}$ and $P_{U}$ can be found using the following formulae

$$
P_{L}=\frac{y_{o b s}}{y_{o b s}+\left(n-y_{o b s}+1\right) F_{2\left(n-y_{o b s}+1\right), 2 \cdot y_{o b s}, 1-\alpha / 2}}
$$

- and

$$
P_{U}=\frac{\left(y_{o b s}+1\right) \cdot F_{2 \cdot\left(y_{o b s}+1\right), 2 \cdot\left(n-y_{o b s}\right), 1-\alpha / 2}}{\left(n-y_{o b s}\right)+\left(y_{o b s}+1\right) \cdot F_{2 \cdot\left(y_{o b s}+1\right), 2 \cdot\left(n-y_{o b s}\right), 1-\alpha / 2}}
$$

## Example using F-dist

- Thus, using our example of $n=5$ and $y_{o b s}=4$

$$
\begin{aligned}
P_{L} & =\frac{y_{o b s}}{y_{o b s}+\left(n-y_{o b s}+1\right) F_{2\left(n-y_{o b s}+1\right), 2 \cdot y_{o b s}, 1-\alpha / 2}} \\
& =\frac{4}{4+2 F_{4,8,0.975}} \\
& =\frac{4}{4+2 \cdot 5.0526} \\
& =0.2836
\end{aligned}
$$

- and

$$
\begin{aligned}
P_{U} & =\frac{\left(y_{o b s}+1\right) \cdot F_{2 \cdot\left(y_{o b s}+1\right), 2 \cdot\left(n-y_{o b s}\right), 1-\alpha / 2}}{\left(n-y_{o b s}\right)+\left(y_{o b s}+1\right) \cdot F_{2} \cdot\left(y_{o b s}+1\right), 2 \cdot\left(n-y_{o b s}\right), 1-\alpha / 2} \\
& =\frac{5 \cdot F_{10,2,0.975}^{1+5 \cdot F_{10,2,0.975}}}{} \\
& =\frac{5 \cdot 39.39797}{1+5 \cdot 39.39797} \\
& =0.9949
\end{aligned}
$$

- Therefore, our 95\% exact confidence interval for $p$ is [0.2836, 0.9949] as was observed previously
\%macro mybinomialpdf(p,n);
dm "output" clear; dm "log" clear; options nodate nocenter nonumber;
data myexample;

$$
\begin{aligned}
& \text { do } i=0 \text { to } \& n ; \\
& \text { prob }=\text { PDF ("BINOMIAL", i, } \& p, \& n) ; \\
& \text { cdf }=\operatorname{CDF}(\text { "BINOMIAL", } i, \& p, \& n) ; \\
& \text { mlcdfprob }=1-c d f+p r o b ;
\end{aligned}
$$

output;
end;
label i $=$ "Number of $*$ Successes";
label prob $=\quad " P(Y=y) \quad " ;$
label cdf $=\quad$ "P(Y<=y)";
label m1cdfprob="P(Y>=y)";
run;
title "Binomial PDF for $N=\& n$ and $P=\& p$ "; proc print noobs label split="*"; run;
\%mend mybinomialpdf;
\%mybinomialpdf(0.4,5);

