

STA305/1004 - Class 7

January 30, 2017

Today's Class

- ▶ Case study on power poses study: study replication and power
- ▶ Sample size and power in studies with two proportions
- ▶ Calculating power via simulation
- ▶ Introduction to causal inference

Can power poses significantly change outcomes in your life?



Professor Amy Cuddy

Can power poses significantly change outcomes in your life?

Cuddy's study methods:

- ▶ Randomly assigned 42 participants to the high-power pose or the low-power-pose condition.
- ▶ Participants believed that the study was about the science of physiological recordings and was focused on how placement of electrocardiography electrodes above and below the heart could influence data collection.
- ▶ Participants' bodies were posed by an experimenter into high-power or low-power poses. Each participant held two poses for 1 min each.
- ▶ Participants' risk taking was measured with a gambling task; feelings of power were measured with self-reports.
- ▶ Saliva samples, which were used to test cortisol and testosterone levels, were taken before and approximately 17 min after the power-pose manipulation.

(Carney, Cuddy, Yap, 2010)

Can power poses significantly change outcomes in your life?

Cuddy's study results:

As hypothesized, high-power poses caused an increase in testosterone compared with low-power poses, which caused a decrease in testosterone, $F(1, 39) = 4.29$, $p < .05$; $r = .34$. Also as hypothesized, high-power poses caused a decrease in cortisol compared with low-power poses, which caused an increase in cortisol, $F(1, 38) = 7.45$, $p < .02$; $r = .43$

Can power poses significantly change outcomes in your life?

- ▶ The study was replicated by Ranehill et al. (2015)
- ▶ An initial power analysis based on the effect sizes in Carney et al. (power = 0.8, $\alpha = .05$) indicated that a sample size of 100 participants would be suitable.

```
library(pwr)  
pwr.t.test(d=0.6,power = 0.8)
```

Two-sample t test power calculation

```
      n = 44.58579  
      d = 0.6  
sig.level = 0.05  
  power = 0.8  
alternative = two.sided
```

NOTE: n is number in *each* group

Can power poses significantly change outcomes in your life?

- ▶ Ranehill et al. study used a sample of 200 participants to increase reliability.
- ▶ This study found none of the significant differences found in Cuddy's study.
- ▶ The replication study obtained very precise estimates of the effects.
- ▶ What happened?

Can power poses significantly change outcomes in your life?

- ▶ Sampling theory predicts that the variation between samples is proportional to $\frac{1}{\sqrt{n}}$.
- ▶ In small samples we can expect variability.
- ▶ Many researchers often expect that these samples will be more similar than sampling theory predicts.

Study replication

Type I

Type II

Suppose that you have run an experiment on 20 subjects, and have obtained a significant result from a two-sided z-test ($H_0 : \mu = 0$ vs. $H_1 : \mu \neq 0$) which confirms your theory ($z = 2.23$, $p < 0.05$, two-tailed). The researcher is planning to run the same experiment on an additional 10 subjects. What is the probability that the results will be significant at the 5% level by a one-tailed test ($H_1 : \mu > 0$), separately for this group?

STUDY 1: $H_0: \mu = 0$ vs $H_1: \mu \neq 0$

$$z = 2.23 = \frac{\bar{X} - 0}{\sigma/\sqrt{20}} \Rightarrow \bar{X} = 2.23 \sigma/\sqrt{20}$$

Study 2: Take $\mu = \bar{X} = 2.23 \sigma/\sqrt{20}$

Reject at 5% level H_0

$$\frac{\bar{X} - 0}{\sigma/\sqrt{10}} \geq 1.645 \Rightarrow \bar{X} \geq 1.645 \frac{\sigma}{\sqrt{10}}$$

$H_A: \mu = 2.23 \sigma/\sqrt{20}$ from Study 1.

$$\begin{aligned} \text{power} &= P\left(\bar{X} \geq 1.645 \frac{\sigma}{\sqrt{10}}, \text{ when } \mu = 2.23 \frac{\sigma}{\sqrt{20}}\right) \\ &= P\left(\frac{\bar{X} - \mu}{\sigma/\sqrt{10}} \geq \frac{1.645 \frac{\sigma}{\sqrt{10}} - \mu}{\sigma/\sqrt{10}}\right) \quad \begin{array}{l} \text{use} \\ \mu = 2.23 \frac{\sigma}{\sqrt{20}} \end{array} \\ &= P(Z \geq 0.068) = 0.47 \end{aligned}$$

Comparing Proportions for Binary Outcomes

- ▶ In many clinical trials, the primary endpoint is dichotomous, for example, whether a patient has responded to the treatment, or whether a patient has experienced toxicity.
- ▶ Consider a two-arm randomized trial with binary outcomes. Let p_1 denote the response rate of the experimental drug, p_2 as that of the standard drug, and the difference is $\theta = p_1 - p_2$.

Comparing Proportions for Binary Outcomes

Let Y_{ik} be the binary outcome for subject i in arm k ; that is,

$$Y_{ik} = \begin{cases} 1 & \text{with probability } p_k \\ 0 & \text{with probability } 1 - p_k, \end{cases} \quad \text{Bernoulli}$$

for $i = 1, \dots, n_k$ and $k = 1, 2$. The sum of independent and identically distributed Bernoulli random variables has a binomial distribution,

$$\sum_{i=1}^{n_k} Y_{ik} \sim \text{Bin}(n_k, p_k), \quad k = 1, 2.$$

(Yin, pg. 173-174)

Comparing Proportions for Binary Outcomes

The sample proportion for group k is

$$\hat{p}_k = \bar{Y}_k = (1/n_k) \sum_{i=1}^{n_k} Y_{ik}, \quad k = 1, 2,$$

and $E(\bar{Y}_k) = p_k$ and $Var(\bar{Y}_k) = \frac{p_k(1-p_k)}{n_k}$.

The goal of the clinical trial is to determine if there is a difference between the two groups using a binary endpoint. That is we want to test $H_0 : \theta = 0$ versus $H_0 : \theta \neq 0$.

The test statistic (assuming that H_0 is true) is:

$$T = \frac{\hat{p}_1 - \hat{p}_2}{\sqrt{p_1(1-p_1)/n_1 + p_2(1-p_2)/n_2}} \sim N(0, 1),$$

Comparing Proportions for Binary Outcomes

The test rejects at level α if and only if

$$|T| \geq z_{\alpha/2}.$$

Using the same argument as the case with continuous endpoints and ignoring terms smaller than $\alpha/2$ we can solve for β

$$\beta \approx \Phi \left(z_{\alpha/2} - \frac{|\theta_1|}{\sqrt{p_1(1-p_1)/n_1 + p_2(1-p_2)/n_2}} \right).$$

Comparing Proportions for Binary Outcomes

Using this formula to solve for sample size. If $n_1 = r \cdot n_2$ then

$$n_2 = \frac{(z_{\alpha/2} + z_{\beta})^2}{\theta^2} (p_1(1 - p_1)/r + p_2(1 - p_2)) .$$

Comparing Proportions for Binary Outcomes

- ▶ The built-in R function `power.prop.test()` can be used to calculate sample size or power.
- ▶ For example suppose that the standard treatment for a disease has a response rate of 20%, and an experimental treatment is anticipated to have a response rate of 28%.
- ▶ The researchers want both arms to have an equal number of subjects. How many patients should be enrolled if the study will conduct a two-sided test at the 5% level with 80% power?

```
power.prop.test(p1 = 0.2,p2 = 0.25,power = 0.8)
```

Two-sample comparison of proportions power calculation

```
      n = 1093.739
    p1 = 0.2
    p2 = 0.25
sig.level = 0.05
  power = 0.8
alternative = two.sided
```

NOTE: n is number in *each* group

Calculating Power by Simulation

- ▶ If the test statistic and distribution of the test statistic are known then the power of the test can be calculated via simulation.
- ▶ Consider a two-sample t-test with 30 subjects per group and the standard deviation of the clinical outcome is known to be 1.
- ▶ What is the power of the test $H_0 : \mu_1 - \mu_2 = 0$ versus $H_0 : \mu_1 - \mu_2 = 0.5$, at the 5% significance level?
- ▶ The power is the proportion of times that the test correctly rejects the null hypothesis in repeated sampling.

Calculating Power by Simulation

We can simulate a single study using the `rnorm()` command. Let's assume that $n_1 = n_2 = 30$, $\mu_1 = 3.5$, $\mu_2 = 3$, $\sigma = 1$, $\alpha = 0.05$.

```
set.seed(2301)
t.test(rnorm(30,mean=3.5,sd=1),rnorm(30,mean=3,sd=1),var.equal = T)
```

Two Sample t-test

```
data:  rnorm(30, mean = 3.5, sd = 1) and rnorm(30, mean = 3, sd = 1)
t = 2.1462, df = 58, p-value = 0.03605
alternative hypothesis: true difference in means is not equal to 0
95 percent confidence interval:
 0.03458122 0.99248595
sample estimates:
mean of x mean of y
 3.339362  2.825828
```

Should you reject H_0 ?

Calculating Power by Simulation

- ▶ Suppose that 10 studies are simulated.
- ▶ What proportion of these 10 studies will reject the null hypothesis at the 5% level?
- ▶ To investigate how many times the two-sample t-test will reject at the 5% level the `replicate()` command will be used to generate 10 studies and calculate the p-value in each study.
- ▶ It will still be assumed that
$$n_1 = n_2 = 30, \mu_1 = 3.5, \mu_2 = 3, \sigma = 1, \alpha = 0.05.$$

```
set.seed(2301)
pvals <- replicate(10,t.test(rnorm(30,mean=3.5,sd=1),
                             rnorm(30,mean=3,sd=1),
                             var.equal = T)$p.value)
pvals # print out 10 p-values
```

```
[1] 0.03604893 0.15477655 0.01777959 0.40851999 0.34580930 0.11131007
[7] 0.14788381 0.00317709 0.09452230 0.39173723
```

```
#power is the number of times the test rejects at the 5% level
sum(pvals<=0.05)/10
```

```
[1] 0.3
```

Calculating Power by Simulation

But, since we only simulated 10 studies the estimate of power will have a large standard error. So let's try simulating 10,000 studies so that we can obtain a more precise estimate of power.

```
set.seed(2301)
pvals <- replicate(10000,t.test(rnorm(30,mean=3.5,sd=1),
                                rnorm(30,mean=3,sd=1),
                                var.equal = T)$p.value)
sum(pvals<=0.05)/10000
```

```
[1] 0.4881
```

Calculating Power by Simulation

This is much closer to the theoretical power obtained from `power.t.test()`.

```
power.t.test(n = 30,delta = 0.5,sd = 1,sig.level = 0.05)
```

Two-sample t test power calculation

```
      n = 30
  delta = 0.5
      sd = 1
sig.level = 0.05
   power = 0.477841
alternative = two.sided
```

NOTE: n is number in *each* group

Calculating Power by Simulation

- ▶ The built-in R functions `power.t.test()` and `power.prop.test()` don't have an option for calculating power where there is unequal allocation of subjects between groups.
- ▶ These built-in functions don't have an option to investigate power if other assumptions don't hold (e.g., normality).
- ▶ One option is to simulate power for the scenarios that are of interest. Another option is to write your own function using the formula derived above.

Calculating Power by Simulation

- ▶ Suppose the standard treatment for a disease has a response rate of 20%, and an experimental treatment is anticipated to have a response rate of 28%.
- ▶ The researchers want both arms to have an equal number of subjects.
- ▶ A power calculation above revealed that the study will require 1094 patients for 80% power.
- ▶ What would happen to the power if the researchers put 1500 patients in the experimental arm and 500 patients in the control arm?

Calculating Power by Simulation

- ▶ The number of subjects in the experimental arm that have a positive response to treatment will be an observation from a $Bin(1500, 0.28)$.
- ▶ The number of subjects that have a positive response to the standard treatment will be an observation from a $Bin(500, 0.2)$.
- ▶ We can obtain simulated responses from these distributions using the `rbinom()` command in R.

```
set.seed(2301)  
rbinom(1, 1500, 0.28)
```

```
[1] 403
```

```
rbinom(1, 500, 0.20)
```

```
[1] 89
```


Calculating Power by Simulation

- The p-value for this simulated study can be obtained using `prop.test()`.

```
set.seed(2301)
prop.test(x=c(rbinom(1,1500,0.28),rbinom(1,500,0.20)),
          n=c(1500,500),correct = F)
```

2-sample test for equality of proportions without continuity
correction

```
data:  c(rbinom(1, 1500, 0.28), rbinom(1, 500, 0.2)) out of c(1500, 500)
X-squared = 16.62, df = 1, p-value = 4.568e-05
alternative hypothesis: two.sided
95 percent confidence interval:
 0.05032654 0.13100680
sample estimates:
   prop 1    prop 2 
0.2686667 0.1780000
```

Calculating Power by Simulation

- ▶ A power simulation repeats this process a large number of times.
- ▶ In the example below we simulate 10,000 hypothetical studies to calculate power.

```
set.seed(2301)
pvals <- replicate(10000,
  prop.test(x=c(rbinom(n = 1,size = 1500,prob = 0.25),
               rbinom(n=1,size=500,prob=0.20)),
            n=c(1500,500),correct=F)$p.value)
sum(pvals<=0.05)/10000
```

[1] 0.6231

If the researchers decide to have a 3:1 allocation ratio of patients in the treatment to control arm then the power will be _____?