# HESP 180 REVIEW

# Block 1 Defining Science, Research, and Epidemiology

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### CLINICAL EPIDEMIOLOGY LECTURES

Block One: Defining Science, Research, and Epidemiology

Lecture 1: Syllabus and the Etymology of "Science"

Lecture 2: Fundamentals of Science

Office Hours (Week 1)

Lecture 3: Fundamentals of Study Design

Lecture 4: Developing a Research Question

Lecture 5: From Bench to Bedside

Lecture 6: Introduction to Epidemiology

Office Hours (Week 2)

**Lecture 7**: Epidemiological Tools and Questions

Lecture 8: Drawing Inappropriate Conclusions

Office Hours (Week 3)

Lecture 9: Current Trends in Epidemiology

Lecture 10: Future of Epidemiology

Lecture 11: Review of Block One

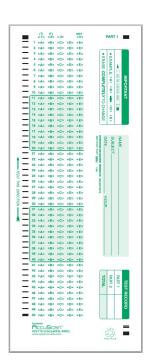
Review Slides (PDF)







# Block 1 Defining Science, Research, and Epidemiology



#### **EXAM ONE**

Scientific method, history of epidemiology, and fundamentals of evidence-based practice. The material you could be tested on includes everything presented during Lectures 1-10, especially (but not exclusively) what was focused on in Lecture 11... and potentially some material from Chapter 6 of the ACSM book.

Not on an actual Scantron, but all multiple choice.









# Block 1 Defining Science, Research, and Epidemiology

Everything that you could potentially be accountable for

Lecture 2:

ACSM Research Methods: Chapter 1 (1-8)

Lecture 3:

ACSM Research Methods: Chapter 5 (73-90)

ACSM Research Methods: Chapter 7 (121-141)

Epidemiology 101: Chapter 6 (105-119) (2nd Edition: Chapter 7)

Lecture 4:

Epidemiology 101: Chapter 4 (65-87) (2<sup>nd</sup> Edition: Chapter 5)

ACSM Research Methods: Chapter 3 (29-49)

Lecture 5:

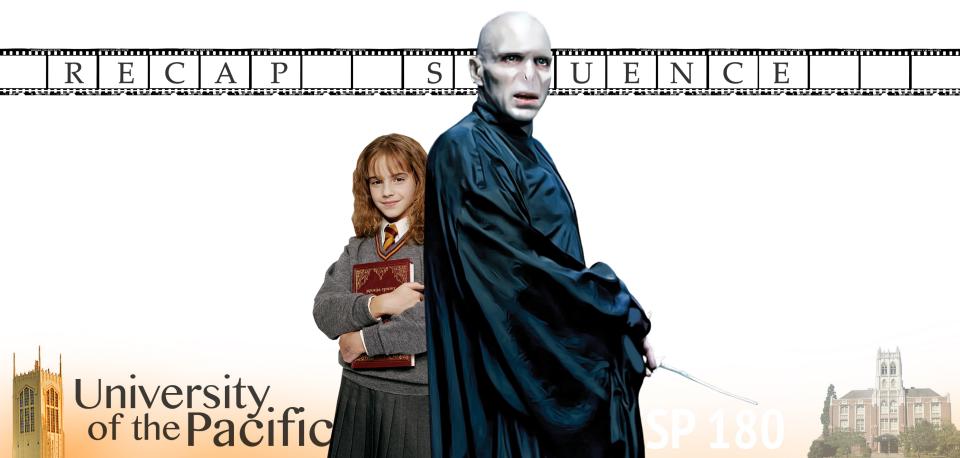
ACSM Research Methods: Chapter 6 (93-118)

**Lecture 6:** 

Epidemiology 101: Chapter 1 (1-23)



For the exam, I'll be drawing from everything we've talked about so far (Lectures 2–10). I've compiled the most important slides here; if you know all of this stuff, you're off to a really good start.

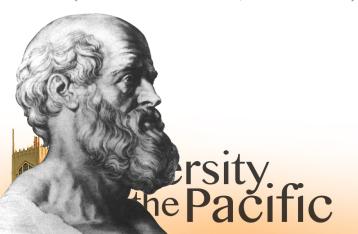


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**Epidemiology:** A long time ago, Hippocrates discussed exposures and outcomes, and offered us some epidemiological nomenclature: *epidemic* and *endemic*. In the 17<sup>th</sup> century, "political arithmetic" entered public health, with the work of demographers such as John Graunt (1620–1674). In the middle of the 19<sup>th</sup> century, John Snow establishes epidemiology as a powerful tool for evaluating relationships between exposures and outcomes.



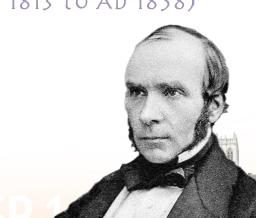
(460BC-ISH TO 370 BC-ISH)





JOHN SNOW

(AD 1813 TO AD 1858)



**Epidemiology:** In the years after John Snow, epidemiology developed and deformed itself. It became a mighty weapon against ignorance, but was often discharged with terrible aim.

Table 1. We have found 12 papers in which claims coming from observational studies were tested in randomised clinical trials. Many of the trials are quite large. In most of the observational studies multiple claims were tested, often in factorial designs, e.g. vitamin D and calcium individually and together along with a placebo group. Note that none of the claims replicated in the direction claimed in the observational studies and that there was statistical significance in the opposite direction five times

ID no.	Pos.	Neg.	No. of claims	Treatment(s)	Reference
1	0	1	3	Vit E, beta-carotene	NEJM 1994; <b>330</b> : 1029–1035
2	0	3	4	Hormone Replacement Ther.	JAMA 2003; 289: 2651-2662, 2663-2672, 2673-2684
3	0	1	2	Vit E, beta-carotene	JNCI 2005; 97: 481-488
4	0	0	3	Vit E	JAMA 2005; <b>293</b> : 1338-1347
5	0	0	3	Low Fat	JAMA. 2006; 295: 655-666
6	0	0	3	Vit D, Calcium	NEJM 2006; <b>354</b> : 669-683
7	0	0	2	Folic acid, Vit B6, B12	NEJM 2006; <b>354</b> : 2764-2772
8	0	0	2	Low Fat	JAMA 2007; <b>298</b> : 289-298
9	0	0	12	Vit C, Vit E, beta-carotene	Arch Intern Med 2007; 167: 1610-1618
10	0	0	12	Vit C, Vit E	JAMA 2008; <b>300</b> : 2123-2133
11	0	0	3	Vit E, Selenium	JAMA 2009; <b>301</b> : 39-51
12	0	0	3	HRT + Vitamins	JAMA 2002; <b>288</b> : 2431–2440
Totals	0	5	52		

significance

september2011

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Deming, data and observational studies
A process out of control and needing fixing







## Good

### The New york Times

**BUSINESS DAY** 

### What Wal-Mart Knows About Customers' Habits

By CONSTANCE L. HAYS NOV. 14, 2004



## **Good & Bad**



### **Positive Side of Neutral**

Table 1. We have found 12 papers in which claims coming from observational studies were tested in randomised clinical trials. Many of the trials are quite large. In most of the observational studies multiple claims were tested, often in factorial designs, e.g. vitamin D and calcium individually and together along with a placebo group. Note that none of the claims replicated in the direction claimed in the observational studies and that there was statistical significance in the opposite direction five times

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significance

september2011

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Deming, data and observational studies
A process out of control and needing fixing







## Good











## **Mostly Good**

nature	Vol 457 19 February 2009 doi:10.1038/nature07	634
Detecting influe query data	nza epidemics using search engir	ıe
Jeremy Ginsberg <sup>1</sup> , Matthew H. Mol	ebbi $^1$ , Rajan S. Patel $^1$ , Lynnette Brammer $^2$ , Mark S. Smolinski $^1$ & Larry Brilli	ant <sup>1</sup>
	California 94043, USA <sup>2</sup> Centers for Disease Control and Prevention, 1600 Clifton Road, NE, Atlanta, Georgia 3033	3, USA
	Google Search I'm Feeling Lucky	

The final model was validated on 42 points per region of previously untested data from 2007 to 2008, which were excluded from all previous steps. Estimates generated for these 42 points obtained a mean correlation of 0.97 (min = 0.92, max = 0.99, n = 9 regions) with the CDC-observed ILI percentages.

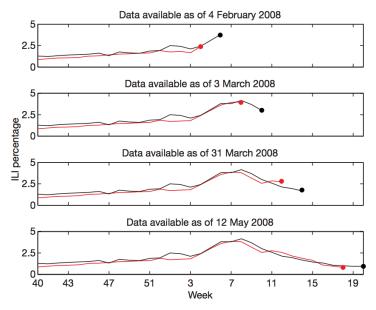


Figure 3 | ILI percentages estimated by our model (black) and provided by the CDC (red) in the mid-Atlantic region, showing data available at four points in the 2007-2008 influenza season.





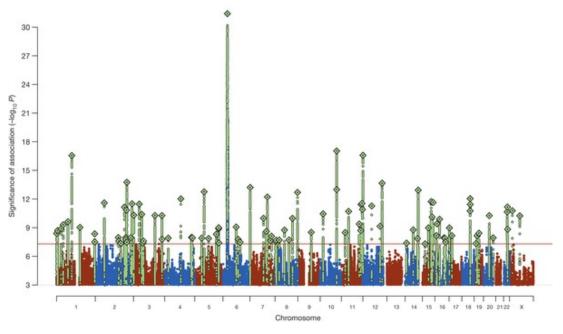


## **Mostly Good**





## Mostly Good, but...



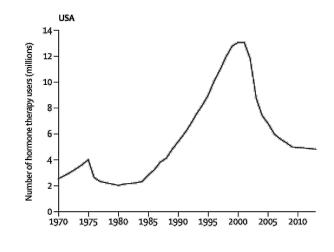
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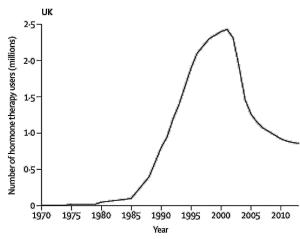
Biological insights from 108 schizophrenia-associated genetic loci



University of the Pacific

## A Bit More Good Than Bad





sity.



## **Barely Leaning Bad**













## **Positive Side of Neutral**

## Maternal Grandmother's Smoking Correlated With Autism Diagnosis

40.5K SHARES







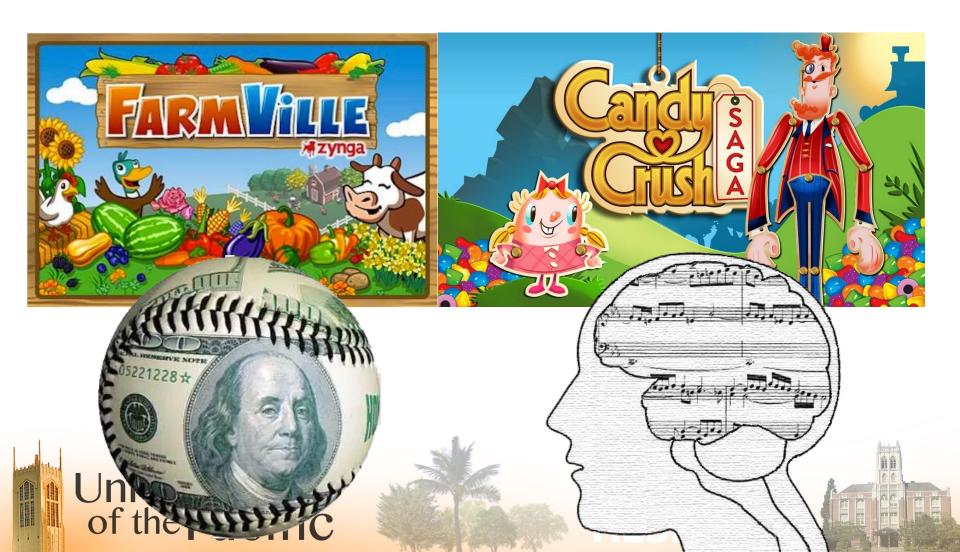






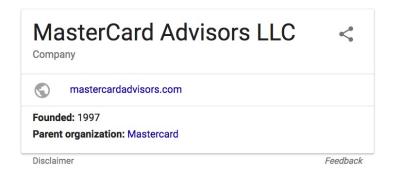


## **Good (Excepting Artistic Value)**



## **Mostly Good**





MasterCard Advisors is a division of the company that aggregates and analyzes tons of transactions (>65 billion) from tons of card holders (>1 billion) in lots of countries (>250) to understand and predict consumer trends.

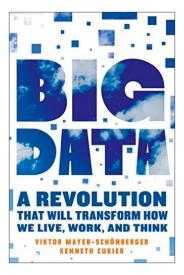






## **Mostly Good**





### **7** IMPLICATIONS

within an hour.

Analysts identified a trend: People filled up their gas tanks around 4 in the afternoon had a high likelihood of spending \$35 to \$50 at a restaurant or grocery store

127

How can that finding be used?

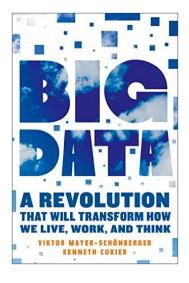






## **Mostly Good**





#### **7** IMPLICATIONS

127

Sell the information to gas stations so they can sell advertising to restaurants and stores, printing coupons on the back of gas receipts.

Epidemiology is true here, but all it's doing is characterizing patterns, not outlining cause and effect.







## Mostly Good, But...



Fitbit is tracking your sleep. Your steps. Your heart rate. Not just for you.

The Health Insurance Portability and Accountability Act (HIPAA) was written in 1996. Its writers didn't anticipate the iPhone or the FitBit. The collecting and sharing of data doesn't fall under HIPAA restrictions. The Federal Food, Drug, and Cosmetic Act discusses devices, but not of this variety. Your information is being analyzed, just like your Amazon purchases.

## **Mostly Good**









## **Mostly Good**

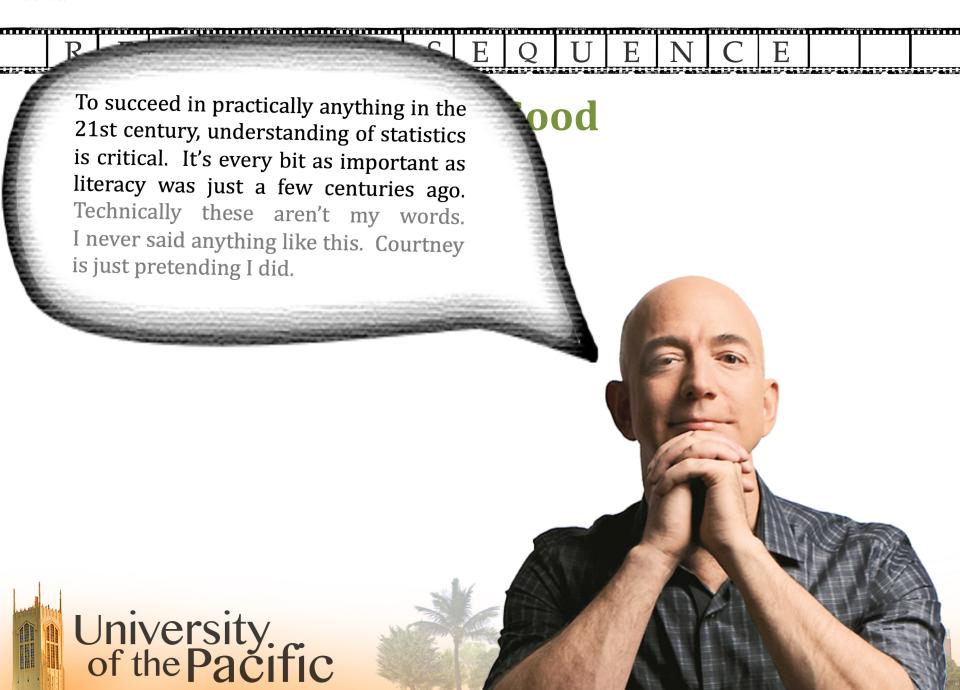
Once upon a time, there were in-house reviewers. Then, there wasn't anymore. Algorithmic recommendations recommend better. And that dries way (way) more sales. What are those algorithms doing?

(Hint: epidemiolog......)

Today, those recommendations and other algorithm-driven personalization features create about a third of Amazon's sales.







## **Mostly Bad**



### **Probability and Preemptive Punishment**

In the year 2054, there's a specialized police department called "PreCrime". They imprisoned people for crimes that they were predicted to commit pretty soon.

(It was some weird, hairless clairvoyant women-ish things, and not data, but data can do it too... sort of.)







## **Mostly Bad**



**Probability and Preemptive Punishment** 

Non-cinematic world:

Parole boards in more than half of the states use data analyses to predict whether to release someone from prison or keep them locked up.

Certain precincts (e.g., LA) use predictive policing. Algorithms predict people and places that are more likely to need policing. So the police are a bit more vigilant with those regions and people.







## **Depends How You Use It**

### **Probability and Preemptive Punishment**

Richard Berk (a full professor of stats and criminology at Upenn) makes predictions on whether a person who gets released on parole will then kill someone or be killed. Using his predictors, he does so with ~75% accuracy.





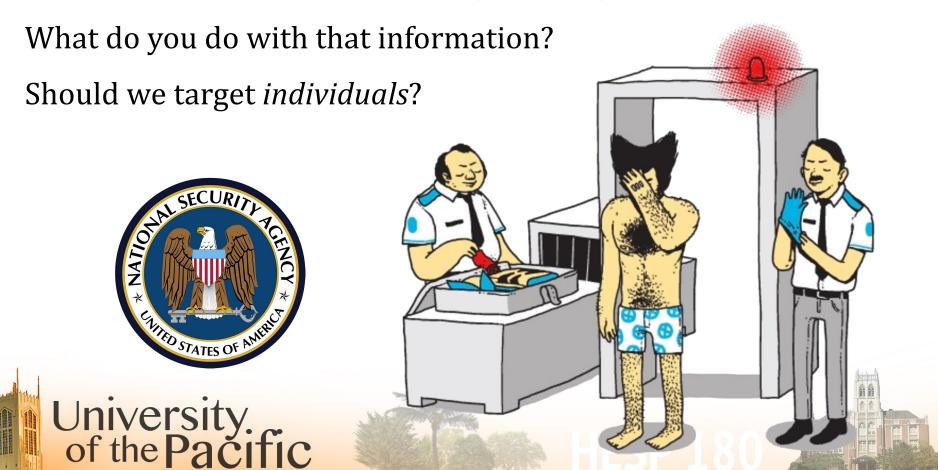


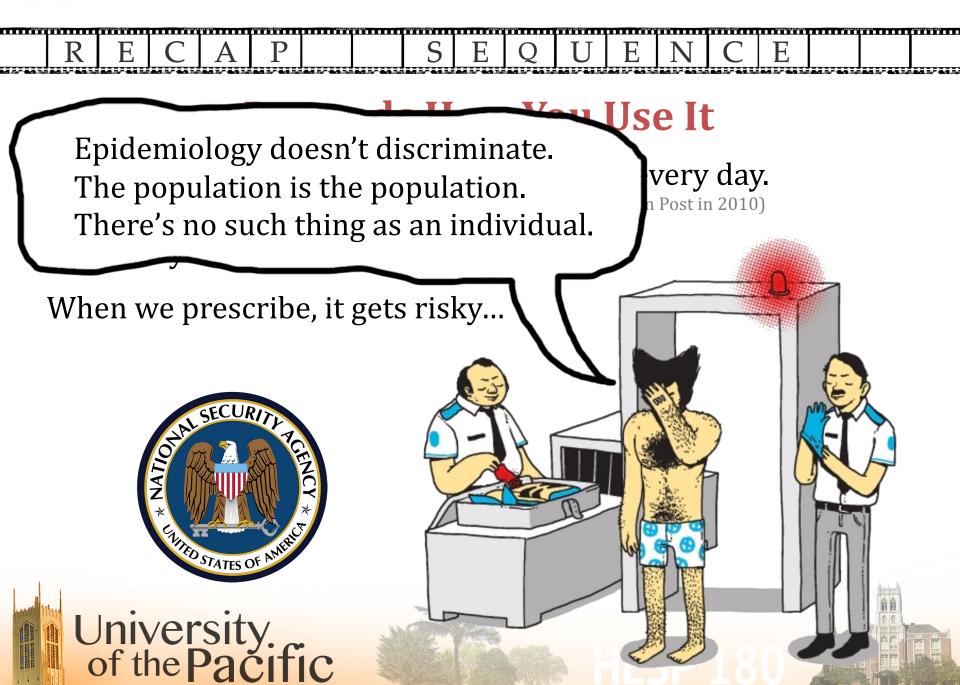


## **Depends How You Use It**

NSA intercepts 1.7 billion communications every day.

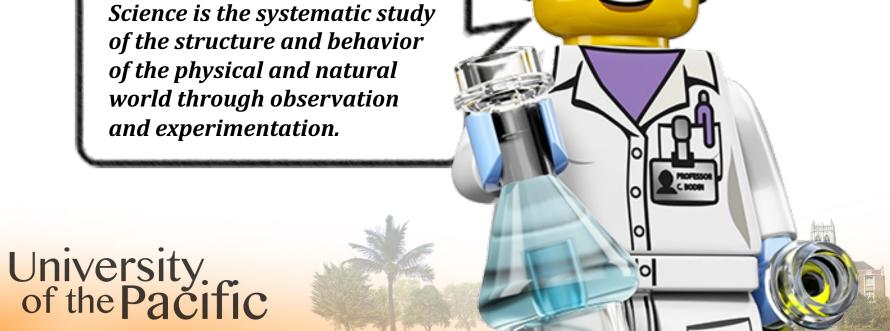
(according to the Washington Post in 2010)





### What's science?

of the structure and behavior of the physical and natural world through observation and experimentation.



What's science?

Why is science important?

How is science typically communicated?

1988. Not famous. Nobody liked this.



**1993.** Ridiculously simple. Sold millions.







## RECAP SEQUENCE

What's science?

Why is science important?

How is science typically communicated?

Why is this (or why is it not) a problem?

How do you fix it?

What's the purpose of research?

What's a good place to start with research?









### Is an egg a fruit or a vegetable?

Party asked 9 mins ago - 6 4 days left to answer

im pretty sure they grow on trees but also they sometimes grow in the ground so is it a fruit?? or a vegetable???



### Can games bite my ears?

Games asked 5 days ago - 👌 3 days left to answer

Can games bite my ears if I wear earphones while playing?

What's a good place to start with research?







Worth asking: What am I good at?
What's a useful thing to know?

I have decided to research what others have already researched. And I've decided to do it in exactly the same way so that my work adds nothing. I want to be boring and pointless.

I'm horrible at working with people. Everyone deeply hates me. I should do research that involves lots and lots of human interaction!

**HESP 18** 

### FROM BENCH TO BEDSIDE

Although each of these examples occupies a different stage in the bench-to-bedside continuum, the respective researchers have a similar set of challenges to overcome. First, they must identify the purpose of their research (as well as the end user it proposes to affect). Only then can one properly decide how the data are to be collected. Once the data are collected, the nature of those data must guide the choice of malysis. Statistics is really a process of coping with uncertainty.

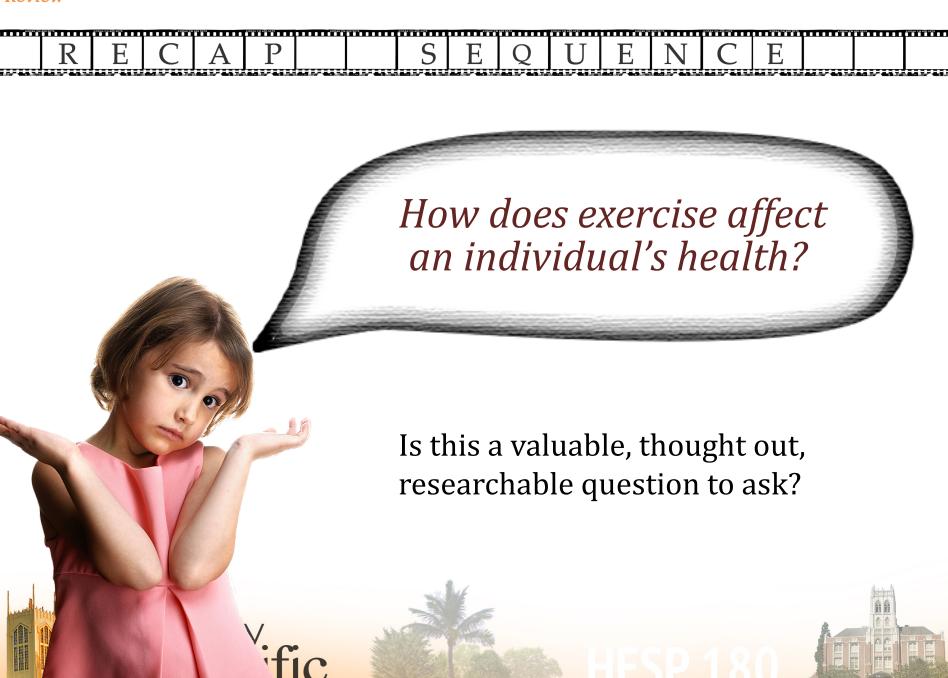






First, they must identify the purpose of their research (as well as the end user it proposes to affect).







How does eight weeks of thrice weekly aerobic exercise (40 min sessions on a cycle ergometer at 60% VO<sub>2</sub> max) affect serum lipoprotein levels (HDLs, LDLs) of post-menopausal women?

University of the Pacific

**HESP 180** 





**Variable:** A *thing* in research that has different values. This includes just about anything that can be measured.

**Independent Variable:** This is the measurable thing you're controlling or manipulating to produce an effect.

**Dependent Variable:** This is the measurable thing that *gets* affected *by* the independent variable.



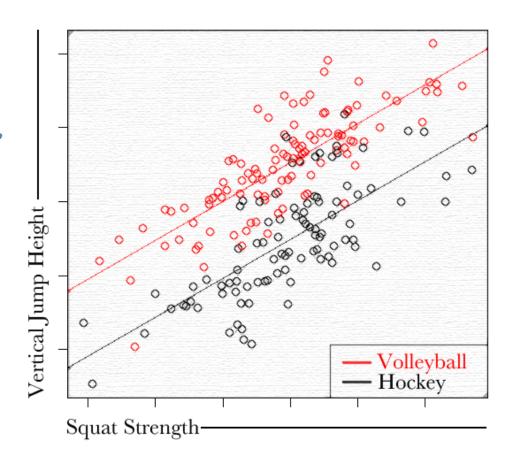


### **Scattered results**



Variables vary, so they have "variance."

In other words, the results will be very scattered.









#### **Perfect results**

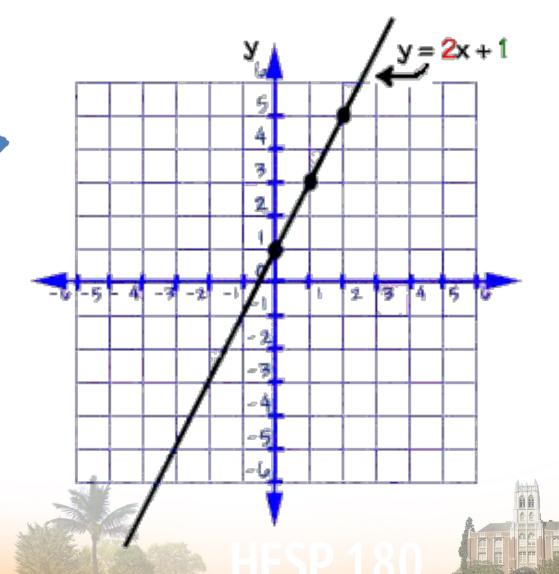


The equation of a straight line (i.e., a "linear equation") is written as:

$$y = mx + b$$

m =slope of the line b = y-intercept.

X and Y (the two variables) would have a perfect correlation there.

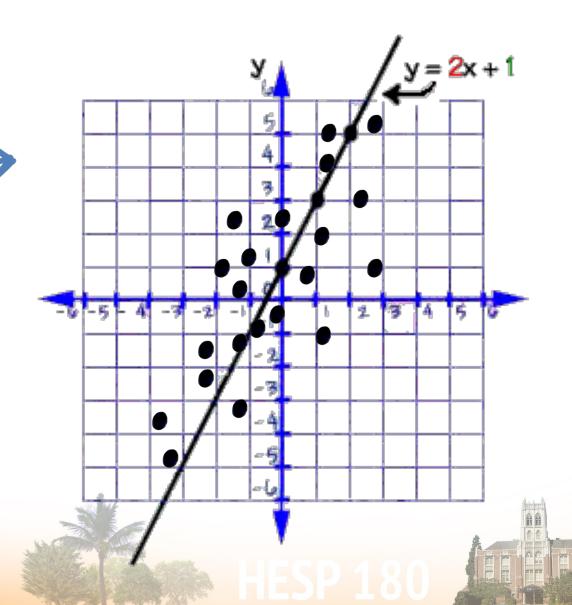




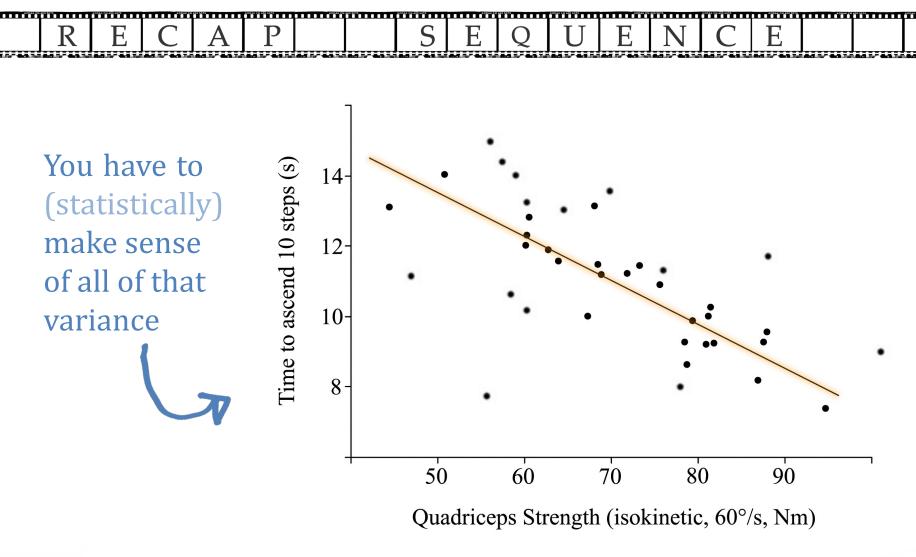
# **Imperfect results**

# High school algebra:

In nature (in reality) there are very few perfect correlations. Almost everything is scattered.



University of the Pacific





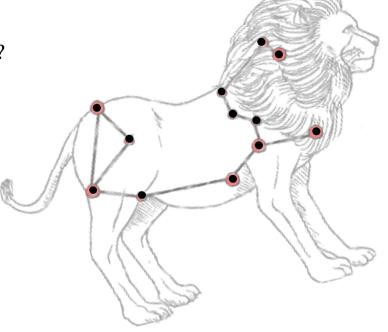




Variance: how spread out are the values? "How scattered are the data points?"

This question requires a reference point.

This question requires a reference point.





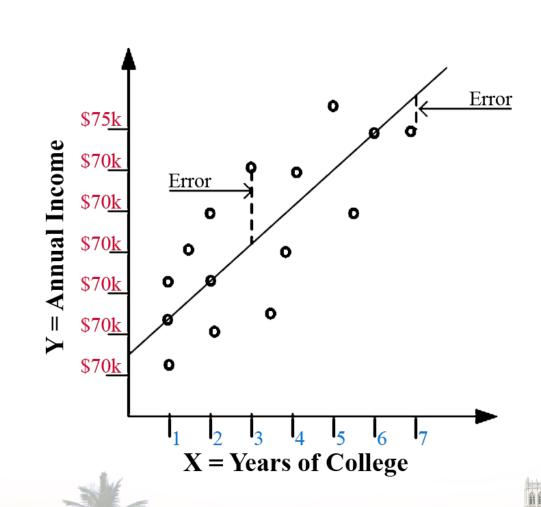




How far is it to Disneyland? Variance: how sprea "How scattered are This question require Bill Herrin Director of SIS

#### variance:

How far are the data scattered from the mean?







Very, very occasionally, your results won't be all that scattered:









Binary outcome: You live or you die. Two categorical options.













# **Types of variables:**

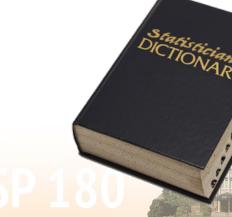
Categorical variables. Dichotomous, nominal, ordinal.

**Discrete random variables.** These are countable. 1, 2, 3, 4, 5, etc. Goals in soccer, runs in baseball, times you've eaten donkey meat, etc. Infinite possible numbers, but they're all integers.

**Continuous random variables.** These values can't be counted. 1.2492, 3¾, 32.12589747. Weight, volume, duration, distance, etc. *Seriously* infinite number of possibilities.







### Scatter in a binary outcome:

1972: WLLWLLWWWWWWLWWWWLLLLWLLLWWWWWLLLWWWL

1974: WLWLLLWWWLWWLWWLWWWLLLWWLWLWLWLWLWWLWWWLWW









#### Scatter in a binary outcome:

1972: WLLWLLWWWWWWLLLLWLLLWLLLWWWWLLLLWWWL

1974: WLWLLLWWWLWWLWWLWWWLLLWWLWLWLWLWLWWLWWWLWW

#### Types of questions you could ask:

What variables predict wins and losses? Who you're playing against? Time of game? Temperature? Miles traveled? How many runs your offense scored? How many errors your defense made.

How do wins and losses affect salary?

1972: \$27,000 salary

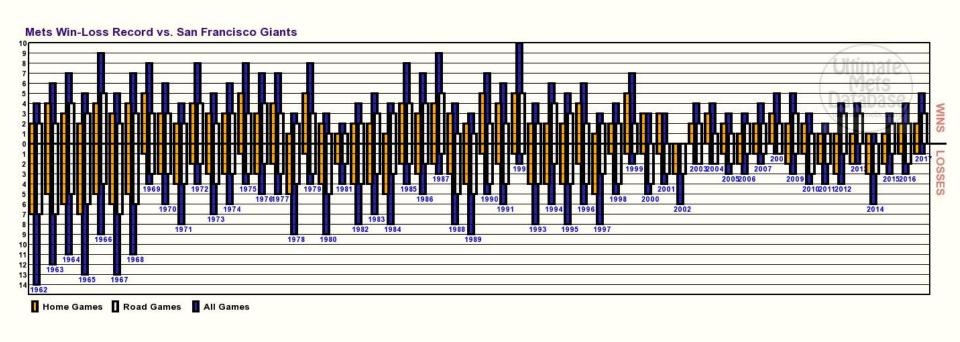
1973: \$47,000 salary

1974: \$100,000 salary





Binary nominal data (win/loss) converted into ratio data:









ш																		
	R	E	C	A	P			S	E	Q	U	E	N	C	E			
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The goal in statistics is to account for and explain that variance. Explain how an independent variable makes a dependent vary.

# Statistical inference.

Your goal is to infer things.
Take a *sample* (e.g., 100 people)
from the population (7.5 billion).
Analyze that sample of 100 people.
And then make some inferences
from your analyses about the
larger population.



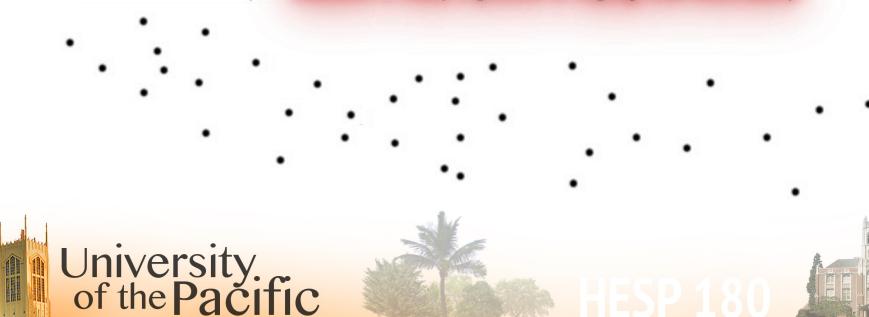




# CHAPTER 6 Understanding Research

#### FROM BENCH TO BEDSIDE

Although each of these examples occupies a different stage in the bench-to-bedside continuum, the respective researchers have a similar set of challenges to overcome. First, they must identify the purpose of their research (as well as the end user it proposes to affect). Only then can one properly decide how the data are to be collected. Once the data are collected, the nature of those data must guide the choice of statistical analysis. **Statistics is really a process of coping with uncertainty.** 



#### CHAPTER 6 Understanding Research

### **UNCERTAINTY AND CONSEQUENCES 106-107**

"In this world nothing can be said to be certain, except death and taxes" (a letter from Benjamin Franklin to Jean-Baptiste Leroy, 1789) (15). The passage of more than two centuries has not altered the truth in Franklin's words. Uncertainty is all around us, and it is the pursuit of "truth" or "proof" that drives research. Research yields data, but we are never sure that the data collected from a sample reflect the whole of the population from which the sample was drawn. This uncertainty led to the development of **inferential statistics** or analyses that permit researchers to estimate the probability that the data gathered are reflective of the population. To some extent, these processes manage the uncertainty inherent in research.







#### CHAPTER 6 Understanding Research

#### UNCERTAINTY AND CONSEQUENCES 107

Uncertainty, however, is a messy subject. Consider cigarette smoking. The link between smoking and lung cancer is well known and widely accepted. More than 85% of lung cancer cases are attributable to smoking (19). However, fewer than 20% of people who smoke cigarettes develop lung cancer. In southwest England, the risk of dying from lung cancer among all lifetime smokers has been estimated to be 15.9% for men and 9% for women. If those lifetime smokers have spent their lives smoking more than 25 cigarettes a day, the risks rise to 24.4% for men and 18.5% for women (14). In Canada, lifetime risks for developing lung cancer among current smokers were 17.2% for men and 11.6% for women (20). At the population level, the link between smoking and an increased risk of developing lung cancer is approaching the certainty of death and taxes, but at the individual level, the fact that someone smokes does not assure the development of the disease.









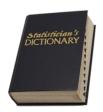
CHAPTER 6 Understanding Research

#### **UNCERTAINTY AND CONSEQUENCES**

Some of this uncertainty can be managed by minimizing error in the study design, both systematic and random. What's the difference between the two types?







In conducting research, there are: Systematic Errors and Random Errors

### Measurement error:

- Instruments aren't working right (e.g., improperly calibrated)
- Human error (e.g., people collecting data are bad at data collection)

Generally manageable by better preparation, maintenance of tools, training of staff, supervision to ensure technician consistency (e.g., interrater reliability), appropriate instruments and software, etc.









In conducting research, there are: Systematic Errors and Random Errors

# *Unknown* deviations that disrupt conclusions:

- Undefined, unpredictable, or otherwise unknown effects
- Setting/environmental factors, lack of enforced controls

Generally manageable if the researcher has enough foresight to identify possible confounders and create proper inclusion/exclusion criteria.







After controlling for as much error as possible, we remain a bit uncertain about how variables interact and affect each another. What effect will a collection of independents have on a dependent variable? The world is packed with variables. How do we use them to test predictions?





HESP:

- Years of participation in an activity
- Presence of a specific allele
- Free throw percentage
- Race/ethnicity
- Hours of sleep per night
- Incidence of the flu
- Number of bowls of crispy hexagons
- Change in body fat percentage
- Which doctor performed the procedure
- Marital status
- Use of anti-inflammatory drugs
- Age of onset of Alzheimer's
- Severity of stress
- Sex/gender
- Mile time

Which are most likely to be independent variables and which ones are dependents?





- Years of participation in an activity
- Presence of a specific allele
- Free throw percentage
- Race/ethnicity
- Hours of sleep per night
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- Sex/gender
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How might you control for the influence of **these** on your outcome variable?







- Presence of a specific allele
- Race/ethnicity

- Which doctor performed the procedure
- Marital status

Sex/gender

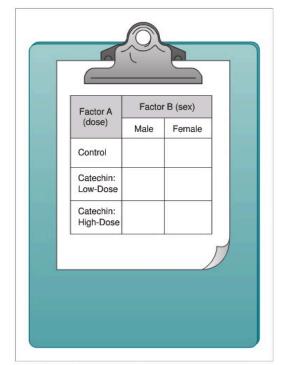
How might you control for the influence of **these** on your outcome variable?

Make them "constants" (only look at one gender, one race, etc.).





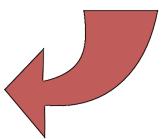
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**FIGURE 7-1:** Schematic of a two-factor design with three levels of factor A (dose of catechin) and two levels for the between factor B of sex (male or female).

How might you control for the influence of **these** on your outcome variable?

Or you can just evaluate them as independents.









# Nominal

Categorical, no intrinsic order

E.g.: gender, race, religion, political party

# Ordinal

Categorical, level or rank

E.g.: white belt, brown belt, black belt

# Interval

Categorical, level or rank, uniform spacing, no zero point

E.g.: Borg RPE scale (6–20), degrees Celsius

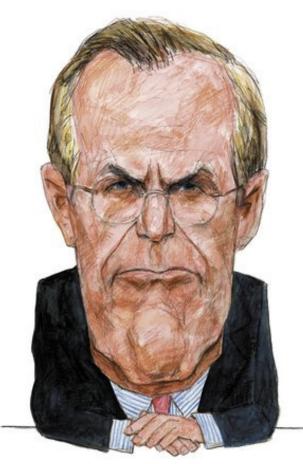
# Ratio

Categorical, level or rank, uniform spacing, zero as reference point

E.g.: height, speed, weight, area (acreage), distance

University.
of the Pacific

**HESP 180** 



"There are known knowns; there are things we know that we know. There are known unknowns; that is to say, there are things that we now know we don't know. But there are also unknown unknowns – there are things we do not know we don't know."

### Extraneous

Also called "confounding" or "intervening"

Not controlled or accounted for in study design

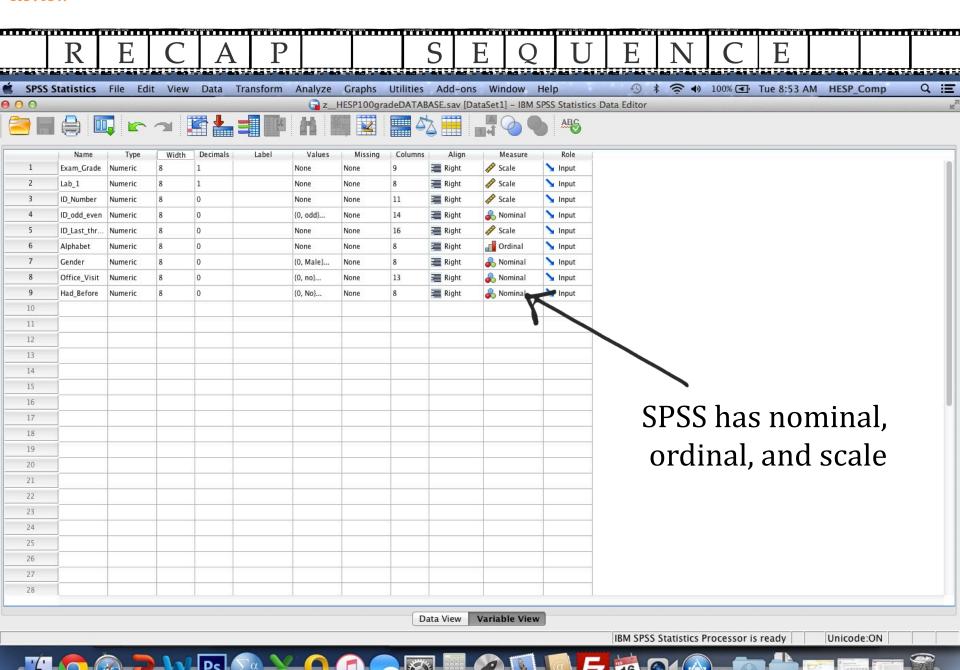
E.g.: nutrition status, level of fatigue











# There are different kinds of independent variables:

**Between-group variable:** You're looking at different subjects in each group. Men vs. women. Obese vs. non-obese. Republicans vs. democrats. You can run these analyses with a single episode of data collection (one time point).

Within-group variable: You're looking at the same subjects in each group. People when they were obese and then when they're not obese are People when they're republicans, and then later when they're depople in a drug trial while they're on the drug and later on the You need at least two episodes of data collection to compare subjects under different conditions.

University of the Pacific

#### There are different kinds of research models:

In a statistical model, you can have any number of independent (explanatory) variables explaining a dependent (outcome) variable. The number of *dependent* variables affects the class of analysis:

**Univariate analysis:** One dependent variable is examined (with as many

independents as you want, sort of).

**Multivariate analysis:** Multiple dependent variables are simple evaluated (using a bunch of independents)

evaluated (using a bunch of independents).







#### Review Observational research Descriptive research **Ecological study** Meta-Analysis Correlational study Systematic Review Cross-sectional study Randomized Controlled Trial Case-control study Controlled Clinical Trial Cohort study Experimental research Cohort Studies Case-Control Studies Randomized controlled trial Cross-Sectional Studies

Case Studies

Animal Models, In-Vitro Studies, Mechanisms Research

Anecdotal Data and Expert Opinion



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Types of epidemiological investigations

	Ecological	Study
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Correlational Study

Cross-Sectional Study

Cohort Study

Case-Control Study

<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)







Types of epidemiological investigations



\_\_\_ Cross-Sectional Study

Cohort Study

Case-Control Study

**Ecological studies** evaluate relationships between exposures and outcomes. Those exposures can either be geographical or temporal (i.e., time-related).

Needs to include as many risk-modifying factors as possible to avoid the effect of unknown unknowns.







<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Types of epidemiological investigations

**1980:** Using the map as a predictor of cancer:

Ecological Study

Correlational Study

Cross-Sectional Study

Cohort Study

**Case-Control Study** 

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Int J Epidemiol. 1980 Sep;9(3):227-31.

Do sunlight and vitamin D reduce the likelihood of colon cancer? Garland CF, Garland FC.

#### **Abstract**

It is proposed that vitamin D is a protective factor against colon cancer. This hypothesis arose from inspection of the geographic distribution of colon cancer deaths in the U.S., which revealed that colon cancer mortality rates were highest in places where populations were exposed to the least amounts of natural light--major cities, and rural areas in high latitudes. The hypothesis is supported by a comparison of colon cancer mortality rates in areas that vary in mean daily solar radiation penetrating the atmosphere. A mechanism involving cholecalciferol (vitamin D3) is suggested. The possibility that an ecological fallacy or other indirect association explains the findings is explored.

Sure, sunlight increases skin cancer (just ask a Floridian); but it may also reduce colon cancer. Are vitamin D pills anti-carcinogenic then?







Types of epidemiological investigations

Ecological Study

Correlational Study

Cross-Sectional Study

Cohort Study

**Case-Control Study** 

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

**2006:** An update to the map's effect:

#### LETTER TO THE EDITOR

Do sunlight and vitamin D reduce the likelihood of colon cancer? Time for a paradigm shift?

From PIET HEIN JONGBLOET

19 September 2006 doi:10.1093/ije/dyl206

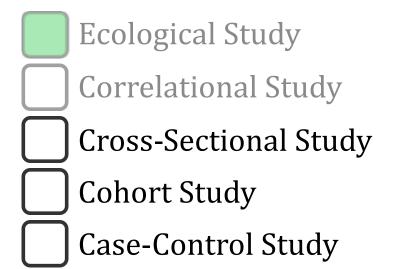
Correlation studies between higher colon cancer rates and higher geographic latitude and, thus, less vitamin D have been extended to other cancer sites, including breast, bladder, corpus uteri, oesophageal, kidney, lung, ovary, pancreas, prostate, rectum, stomach, multiple myeloma, and non-Hodgkin lymphoma. And, if that was not enough, also to other diseases—including hypertension, type 1 diabetes mellitus, multiple sclerosis and osteoporosis—that are linked to vitamin D insufficiency. This hypothesis, therefore, has found adherence and has been tested in different ways. However, it is broadly accepted that there was no vitamin D deficiency in the US.







Types of epidemiological investigations



<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

**2006:** An update to the map's effect:

Metropolitan vs. non-metropolitan? The sunlight exposure will be different.

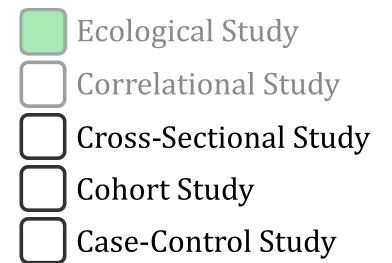
Seasonal Preovulatory Overripeness Ovopathy: During some seasons, the preovulatory phase of egg development is protracted, and that can increase the likelihood of congenital anomalies. Maybe. And maybe this increases the further away from the equator one is. And maybe the immune system anomalies are responsible for the elevated cancer rates.







Types of epidemiological investigations



\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

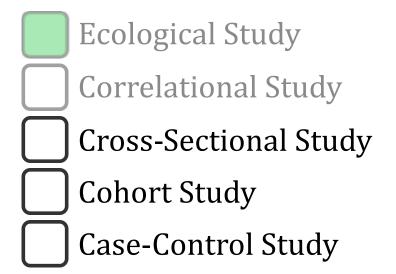
**2006:** An update to the map's effect:

This proposed causal relation of seasonally bound non-optimally matured oocytes to cancer, therefore, not only agrees with the geographical latitude effect under discussion but also with the disproportional month-of-birth deviations apparent in childhood leukaemia, 13–15 premenopausal breast cancer, 16–18 lung cancer, 19 and brain tumours in children 20,21 and adults. 22 These deviations in general correspond with the major total birth peak or major ovulatory season. 11,12 Further, it is in line with the enigmatic phenomenon of shorter life expectancy of people born in the first half of the year when compared with the second half, at least in Europe and upside down in Australia. 23





Types of epidemiological investigations



\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

**2006:** An update to the map's effect:

An analogous persistent paradigm regards the so-called Mediterranean diet allegedly protective for cardiovascular diseases. Olive oil in the diet has been argued to be a similarly confounding or indirect factor, which is associated with geographical latitude. Seasonally bound ovopathy and its pathophysiological consequences related to geographical latitude again explains the presumed association and needs a paradigm shift. 'Not the olive is important, but where the olive grows'. A similar paradigm shift related to vitamin D might be necessary. It would discard some inconsistencies encountered, such as fortification or addition of vitamin D in the nutrition still under debate. The epidemiological findings do not prove that higher levels of vitamin D would lower risk of colorectal cancer, while overexposure to UVA is harmful.





Types of epidemiological investigations





Cross-Sectional Study

Cohort Study

Case-Control Study

**Correlational studies** measure simple statistical relationships between variables. Do variables move together? If you attend class more, does that associate with a better grade? How strong is that relationship?

Strength of relationship is calculated as a correlation coefficient between -1 and +1. A correlation of zero means there's no relationship; the variables don't move together. A strong positive correlation means the variables move in the same direction (a greater squat max associates with a better vertical jump). A strong negative correlation means the variables move in opposite directions: The older you are, the lower your vertical jump. In general, anything less than 0.3 can be thought of as a bit weak.



<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Types of epidemiological investigations



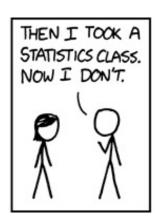


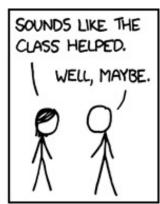
Cross-Sectional Study

Cohort Study

Case-Control Study

I USED TO THINK CORRELATION IMPLIED CAUSATION.





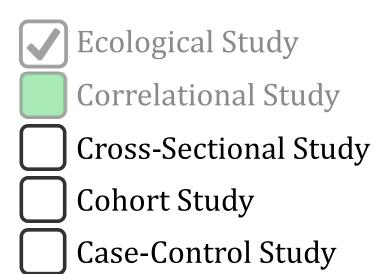
<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)







Types of epidemiological investigations



\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)





# Mortality Attributable to Low Levels of Education in the United States

Published: July 8, 2015

Patrick M. Krueger<sup>1,2</sup>\*, Melanie K. Tran<sup>1</sup>, Robert A. Hummer<sup>3</sup>, Virginia W. Chang<sup>4</sup>

#### **Variables**

Our outcomes include all-cause mortality, cardiovascular disease mortality (ICD-10 codes I00 through I99) and cancer mortality (ICD-10 codes C00 through C97). We use broad cause-of-death categories to limit the impact of misclassification of underlying causes of death on death certificates. Age at baseline is measured in quarter-years (range 25.0 to 84.75). Birth cohort is also measured in quarter-years (range 1901.25 to 1980.5). Because we use 19 cross-sectional waves of the NHIS, follow vital status for up to 21 years, and allow age to increase throughout the follow-up period, the age and birth cohort variables are not perfectly collinear.

Educational attainment is coded categorically as less than a high school degree, high school degree or GED, some college but no baccalaureate degree, baccalaureate degree, and any post-baccalaureate education. Our variable captures key educational milestones. The NHIS ascertains educational attainment as years completed prior to 1997 and as highest degree completed in 1997 and later; we recode years of education completed in the 1986–1996 waves of data to be consistent with the highest degree measure used in later waves. This coding strategy has been used previously to estimate educational disparities among U.S. adults [6, 7]. Race/ethnicity is coded categorically as non-Hispanic white, non-Hispanic black, non-Hispanic Asian/Pacific Islander, non-Hispanic Native American, Mexican American, Puerto Rican, Cuban, and all others. Sex is measured dichotomously.

HESP 180

Types of epidemiological investigations

**Ecological Study** 

Correlational Study

Cross-Sectional Study

Cohort Study

Case-Control Study

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

These three are collectively referred to as **observational studies** and fall within the scope of **epidemiology**.









Types of epidemiological investigations







Cohort Study

Case-Control Study

**Cross-sectional studies** evaluate prevalence, i.e., the number of cases of some condition at a given point in time. Some specific population (that whole population) at one point in time.

For example, look at people with a particular exposure (e.g., those who smoke) and calculate the proportion of that population who have a condition (e.g., COPD).

One point in time. Were you exposed? And do you have the condition?







<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Types of epidemiological investigations







\_\_\_\_ Cohort Study

Case-Control Study

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)





BMJ 1998;316:1643-6

Prevalence of serious eye disease and visual impairment in a north London population: population based, cross sectional study

A Reidy, D C Minassian, G Vafidis, J Joseph, S Farrow, J Wu, P Desai, A Connolly

Objective: To estimate the magnitude of serious eye disorders and of visual impairment in a defined elderly population of a typical metropolitan area in England, and to assess the frequency they were in touch with, or known to, the eye care services.

Design: Cross sectional survey using two stage cluster random sampling.

Setting: General practices in north London.
Subjects: Random sample of people aged 65 and older, drawn from a defined population of elderly people registered with 17 general practice groups.

Main outcome measures: Proportions and population prevalence estimates were determined for visual acuity, assessed with the person's own spectacles (if any), classified into four categories: prevalence of cataract, age related macular degeneration, and refractive error causing visual impairment and of definite primary open angle glaucoma; and status of

contact with eye services.

Results: 1547 of 1840 (84%) eligible people were examined. The population prevalence of bilateral visual impairment (visual acuity <6/12) was 30%, of which 72% was potentially remediable. 92 of these 448 cases (21%) had visual acuity <6/60 ("blindness") in one or both eyes. Prevalence of cataract causing

Types of epidemiological investigations







Cohort Study

Case-Control Study

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

#### **Cross-sectional study:**

BMJ 1998;316:1643-6

Prevalence of serious eye disease and visual impairment in a north London population: population based, cross sectional study

A Reidy, D C Minassian, G Vafidis, J Joseph, S Farrow, J Wu, P Desai, A Connolly

**Conclusions:** Untreated visual impairment and eye disorders affect a substantial proportion of people aged 65 years and older. These findings should contribute to the setting up of future strategies for preservation of sight and eye health services in general.







Types of epidemiological investigations







Cohort Study

Case-Control Study

#### **Cross-sectional study you can conduct:**

A cross-sectional study of your Pacific peers:

What percentage of students exercise at Baun?

What percentage of students have scholarships?

What percentage of students use tutoring?

Then go one step further: what predicts the use of tutoring? Is it GPA? Age? Gender? Language barriers? Declared major? What are the independent variables here?







<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Types of epidemiological investigations







Cohort Study



<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

Cohort studies are generally prospective, but if you've already collected the data for one purpose, and then you come up with a new purpose (a different research question that can be answered using the same database), you can obviously analyze that same database retrospectively for your new question.







Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

#### **Cohort studies:**

<u>Step 1:</u> Identify a population that does *not* have a disease (or whatever condition of interest).

Step 2: Identify two different samples within that population. Half of them are being exposed to something you think will cause the condition/disease and the other half of them do not have that exposure.

<u>Step 3:</u> Follow those samples and see if who gets the condition/disease. Is there a difference in prevalence between the two samples?







Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

#### **Cohort studies:**

*Step 4:* Calculate relative risk of getting the disease/condition.

<u>Step 5:</u> Report your findings, but be careful in how cause-and-effecty you are when phrasing those findings.







Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)



#### Prospective cohort study:

BMJ 1994;309:901-11

Mortality in relation to smoking: 40 years' observations on male British doctors

Richard Doll, Richard Peto, Keith Wheatley, Richard Gray, Isabelle Sutherland

Objective—To assess the hazards associated with long term use of tobacco.

Design—Prospective study of mortality in relation to smoking habits assessed in 1951 and again from time to time thereafter, with causes sought of deaths over 40 years (to 1991). Continuation of a study that was last reported after 20 years' follow up (1951-71).

Subjects—34 439 British male doctors who replied to a postal questionnaire in 1951, of whom 10 000 had died during the first 20 years and another 10 000 have died during the second 20 years.

Results—Excess mortality associated with smoking was about twice as extreme during the second half of the study as it had been during the first half. The death rate ratios during 1971-91 (comparing continuing cigarette smokers with lifelong non-smokers) were approximately threefold at ages 45-64 and twofold at ages 65-84. The excess mortality was chiefly from diseases that can be

Types of epidemiological investigations



**Ecological Study** 



**Correlational Study** 



**Cross-Sectional Study** 



**Cohort Study** 



**Case-Control Study** 

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)



#### Prospective cohort study:

BMJ 1998;316:1631-5

BMJ VOLUME 316 30 MAY 1998

Adverse socioeconomic conditions in childhood and cause specific adult mortality: prospective observational study

George Davey Smith, Carole Hart, David Blane, David Hole

**Objective:** To investigate the association between social circumstances in childhood and mortality from various causes of death in adulthood.

Design: Prospective observational study.

**Setting:** 27 workplaces in the west of Scotland.

Subjects: 5645 men aged 35-64 years at the time of

examination.

Main outcome measures: Death from various causes. Results: Men whose fathers had manual occupations when they were children were more likely as adults to have manual jobs and be living in deprived areas. Gradients in mortality from coronary heart disease, stroke, lung cancer, stomach cancer, and respiratory disease were seen (all P < 0.05), generally increasing from men whose fathers had professional and

managerial occupations (social class I and II) to those

Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)



#### **Retrospective cohort study:**

J Accid Emerg Med 1997;14:371-374

How do individuals with diabetes use the accident and emergency department?

Elizabeth C Goyder, Stephen W Goodacre, Johannes L Botha, Gautam G Bodiwala

Objective—To determine whether the frequency and pattern of use of the accident and emergency (A&E) department by individuals with diabetes is different from that of the general population.

Methods—A historical cohort of 696 individuals with diabetes from six randomly selected general practices and a non-diabetic comparison cohort matched on age, sex, and general practice were identified. The use of an urban A&E department by the two cohorts was compared for number of visits between 1984 and 1996 for injuries, diabetes related and non-diabetes related illness, proportion referred by a general practitioner, proportion arriving by ambulance, and proportion admitted.

Results—More visits were made by the diabetic cohort (1002 v 706, P = 0.0001); 121 visits were directly related to diabetes, including 52 for hypoglycaemia. The diabetic cohort also had more visits for medical illness unrelated to diabetes (357)

Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

#### **Cohort study you can conduct:**

<u>Dependent variable:</u> Admission to graduate school (odds ratio; how likely is it?).

**Population:** Students at Pacific.

<u>Primary independent variable:</u> (main exposure that you think will affect the odds of admission): Whether students have athletic scholarships.

<u>Other confounding variables to include:</u> Age, gender, parental education, participation in extracurricular activities, declared major, etc.







Types of epidemiological investigations



**Ecological Study** 



**Correlational Study** 



**Cross-Sectional Study** 



**Cohort Study** 



**Case-Control Study** 

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

**Case-control studies** are always retrospective. You select a sample based on the presence or absence of a disease. Then look at a specific exposure as a predictor of the current state of the samples.

You'll evaluate a minority of the population (in a cross-sectional study, it's one point in time, looking at the whole population).

Then you describe absolute and relative risk via prevalences of stuff. For example, what's the prevalence of brain damage toady based on sport participation a while ago?







Types of epidemiological investigations



**Ecological Study** 



**Correlational Study** 



**Cross-Sectional Study** 



**Cohort Study** 



**Case-Control Study** 

\* Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)



#### **Case-control study:**

Lancet 2006; 368: 647-58

Tobacco use and risk of myocardial infarction in 52 countries in the INTERHEART study: a case-control study

Koon K Teo, Stephanie Ounpuu, Steven Hawken, MR Pandey, Vicent Valentin, David Hunt, Rafael Diaz, Wafa Rashed, Rosario Freeman, Lixin Jiang, Xiaofei Zhang, Salim Yusuf, on behalf of the INTERHEART Study Investigators

Background Tobacco use is one of the major avoidable causes of cardiovascular diseases. We aimed to assess the risks associated with tobacco use (both smoking and non-smoking) and second hand tobacco smoke (SHS) worldwide.

Methods We did a standardised case-control study of acute myocardial infarction (AMI) with 27 089 participants in 52 countries (12 461 cases, 14 637 controls). We assessed relation between risk of AMI and current or former smoking, type of tobacco, amount smoked, effect of smokeless tobacco, and exposure to SHS. We controlled for confounders such as differences in lifestyles between smokers and non-smokers.

Findings Current smoking was associated with a greater risk of non-fatal AMI (odds ratio [OR] 2.95, 95% CI 2.77–3.14, p<0.0001) compared with never smoking; risk increased by 5.6% for every additional cigarette smoked. The OR associated with former smoking fell to 1.87 (95% CI 1.55–2.24) within 3 years of quitting. A residual excess risk remained 20 or more years after quitting (1.22, 1.09–1.37). Exclusion of individuals exposed to SHS in the never smoker reference group raised the risk in former smokers by about 10%. Smoking beedies alone (indigenous to South Asia) was associated with increased risk (2.89, 2.11–3.96) similar to that associated with cigarette smoking. Chewing tobacco alone was associated with OR 2.23 (1.41–3.52), and smokers who also chewed tobacco had the highest increase in risk (4.09, 2.98–5.61). SHS was associated with a graded increase in risk related to exposure; OR was 1.24 (1.17–1.32) in individuals who were least exposed (1–7 h per week) and 1.62 (1.45–1.81) in people who were most exposed (>21 h per week). Young male current smokers had the highest population attributable risk (58.3%; 95% CI 55.0–61.6) and older women the lowest (6.2%, 4.1–9.2). Population attributable risk for exposure to SHS for more than 1 h per week in never smokers was 15.4% (12.1–19.3).

Conclusion Tobacco use is one of the most important causes of AMI globally, especially in men. All forms of tobacco use, including different types of smoking and chewing tobacco and inhalation of SHS, should be discouraged to prevent cardiovascular diseases.

**HESP 180** 

Types of epidemiological investigations











<sup>\*</sup> Non-epidemiological work: randomized, controlled trials (and their less-controlled, less random companions)

#### **Case-control study you can conduct:**

<u>Dependent variable:</u> Graduation.

<u>Population:</u> Pacific students after graduation (some graduated; some failed to do so).

<u>Primary independent variable:</u> (main exposure that you think will affect graduation rate): Whether students live on campus.

Other confounding variables to include: Age, gender, parental education, participation in athletics, declared major, etc.







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#### Difference between a cohort study and a case-control study

Retrospective study that selects subjects based on their disease status.

	Cases	Controls
Exposed	A	В
Unexposed	С	D

Sick people ("disease positive"): case group.

Not sick people ("disease negative"): control group.

They should all (ideally) come from the same population. And the control group should have a similar distribution of exposure as the case group.







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#### Difference between a cohort study and a case-control study

Retrospective study that selects subjects based on their disease status.

	Cases	Controls
Exposed	A	В
Unexposed	С	D

Sick people ("disease positive"): case group.

Not sick people ("disease negative"): control group.

Compare A, B, C, and D (cases vs. controls; exposed vs. not exposed).

#### Generate an odds ratio:

Ratio of odds of exposure in cases  $(A \div C)$  to odds of exposure in controls  $(B \div D)$ , i.e.,  $(A \times D) \div (B \times C)$ .







#### Difference between a cohort study and a case-control study

#### If the odds ratio is > 1:

Then the infected people are more likely to have been exposed.

#### If the odds ratio is roughly 1ish:

Then the disease (condition/infection/whatever) isn't correlated with the exposure.

#### If the odds ratio is < 1:

Then the exposure seems to have a protective effect.







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### Difference between a cohort study and a case-control study

#### Odds ratio of 1.1:

	Cases	Controls
Exposed	1732	1652
Unexposed	1652	1732

#### Odds ratio of 1.5:

	Cases	Controls
Exposed	103	84
Unexposed	84	103





#### Difference between a cohort study and a case-control study

Subjects are selected based on exposure status.

At the beginning of the study, all subjects *should* be disease free and at risk of acquiring whatever disease.

Follow those subjects over time and reassess them later.







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#### Difference between a cohort study and a case-control study

**For example:** follow smokers and non-smokers over time.

Test lung cancer i	ncidence af	ter that period	of time.
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	Case	Non-case	Total
Exposed	A	В	(A + B)
Unexposed	C	D	(C + D)



Similar 2 × 2 table is constructed







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### Difference between a cohort study and a case-control study

But you don't calculate an odds ratio; you estimate relative risk.

*Relative risk*: probability of disease in the exposed group  $(A \div (A + B))$  over the disease over the probability of disease in the unexposed group  $(C \div (C + D))$ .

Or this: Probability of disease in exposed group

Probability of disease in unexposed group









EPIDEMIOLOGY: THE STUDY OF WHAT IS UPON PEOPLE.

#### Primary purpose of epidemiological research:

Collect information about populations that can be used to prevent, control, or treat various health problems.





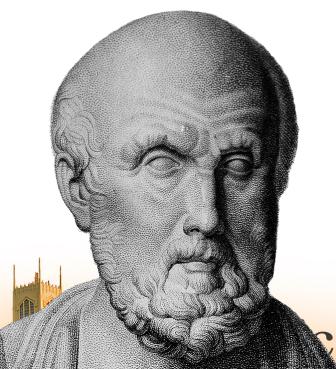


HIPPOCRATES Hippocrates' understanding of medicine was given up, but his principles of understanding remained.

#### **Epidemiological terms coined by Hippocrates:**

**Endemic:** diseases generally found in certain locations, but not others.

**Epidemic:** Diseases generally seen at certain times, but not others.



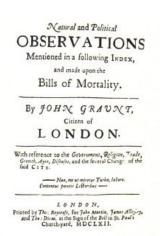
Hippocrates was the first to distinguish between the diseases that *visit* populations (epidemic) and those that reside within populations (endemic).

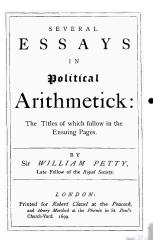


Eventually, *political arithmetic* becomes a thing.

Late-Renaissance effort by fancy wig-wearing Europeans to characterize demographic and economic data. Mostly a tool to estimate the size and growth of their populations (as well as the life expectancies of those being summed).

The increasingly complex analyses of these data, aimed at the creation of a healthier state, were coined "statistics"







	R	E	C	A	P		S	E	Q	U	E	N	C	E		
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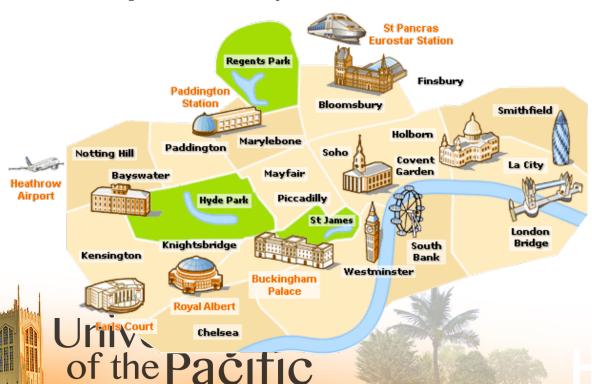
John Graunt (1620 – 1674) – British Haberdasher. Also one of the first demographers.

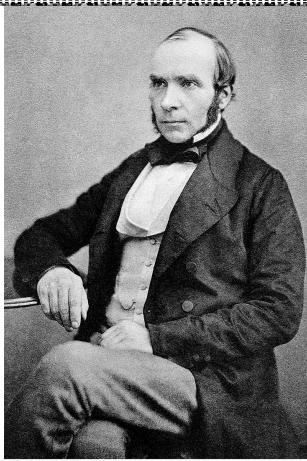
Graunt made the first life table (likelihood that you'll die based on your age). Probability of survival if you're 70, probability of survival if you're 76, etc.

N 100 100 100 100 100 100 100 100 100 10	Table 1. Graunt		
Age Interval	Prop. Deaths in Interval	Prop. Surviving til start of Interval	
0-6	0.36	1.00	
7-16	0.24	0.64	
17-26	0.15	0.40	R
27-36	0.09	0.25	
37-46	0.06	0.16	
47-56	0.04	0.10	
57-66	0.03	0.06	
37-76	0.02	0.03	
77-86	0.01	0.01	
<b>P</b> arsity			
rsity	ific	HESP1	

On August 31, 1854, lots of Londoners began to get cholera: the "1854 Broad Street Cholera Outbreak".

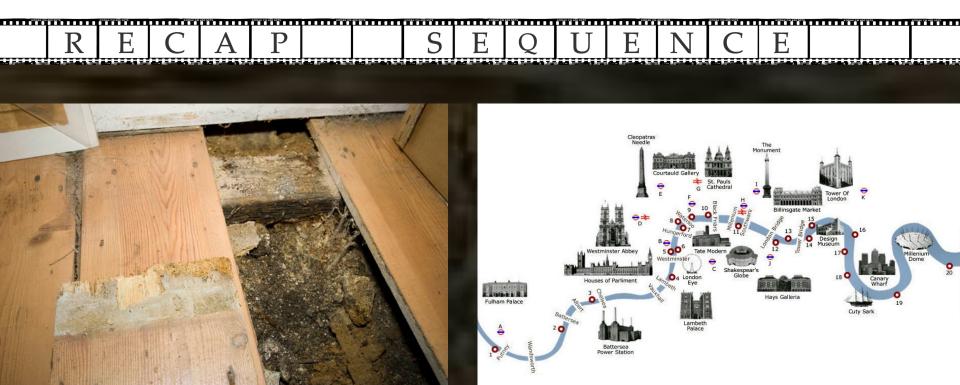
During the first three days, 127 people on or near Broad Street died. By September 10<sup>th</sup>, there were 500 deaths (after ¾ of residents fled the area). Mortality rate was >12% in parts of the city. Final death count: 616–668.





John Inow





Cesspools beneath people's basement floorboards were overflowing.

So they dumped the overflow into the River Thames.









People started dying.







"From inhaling the odour of beef the butcher's wife obtains her obesity."

Professor H Booth, writing in the Builder, July 1844

This assertion is perhaps the most extravagant manifestation of a belief that prevailed in the medical profession for much of the 19th century and survived in some quarters into the 20th century. This belief held that most, if not all, disease was caused by inhaling air that was infected through exposure to corrupting matter. Such matter might be rotting corpses, the exhalations of other people already infected, sewage, or even rotting vegetation. The "miasmatic" explanation of the cause of disease figured prominently in the long debates among the people who were responsible for combating the cholera epidemics that afflicted Britain, and particularly London, between 1831 and 1866.

BMJ VOLUME 323 22-29 DECEMBER 2001 Death and miasma in Victorian London: an obstinate belief Stephen Halliday



### It must be the stench that's killing them...







All smell is, if it be intense, immediate acute disease; and eventually we may say that, by depressing the system and rendering it susceptible to the action of other causes, all smell is disease.

Metropolitan Sewage Committee proceedings. *Parliamentary Papers*. (1846). 10: 651.



It must be the stench that's killing them...

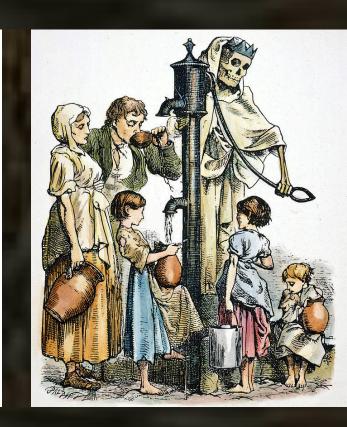








John Snow observed that cholera struck especially hard around a water pump on Broad Street. He persuaded the local parish to remove the pump handle. And he suggested that drinking water should be piped in from regions waterways that were free from pollution.











Snow J. (1857). Cholera and the water supply in the southern districts of London.

**1857:** John Snow published the first modern epidemiological paper. He compared cholera-related mortality rates between customers of two water companies:

- **1. Southwark Water Company** (water came from the most polluted region of the River Thames).
- **2. Lambeth Water Company** (water came from Thames Ditton, where there was no sewage in the tideway).

Cholera-related mortality among Southwark customers was six times higher.

He was correct. So what obviously happened?













Epidemiology after Snow

People see cause and effect in everything.





RECAP SEQUENCE

Epidemiology after Snow

People see cause and effect in everything.



We're great at pattern reception and we apply it to everything.



#### **Michael Shermer:**

"Humans are pattern-seeking, story-telling animals, and we are quite adept at telling stories about patterns, whether they exist or not."















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Correlation just finds *that* there's a relationship, not the nature of it. Computers are bad at this. Remember the captcha.



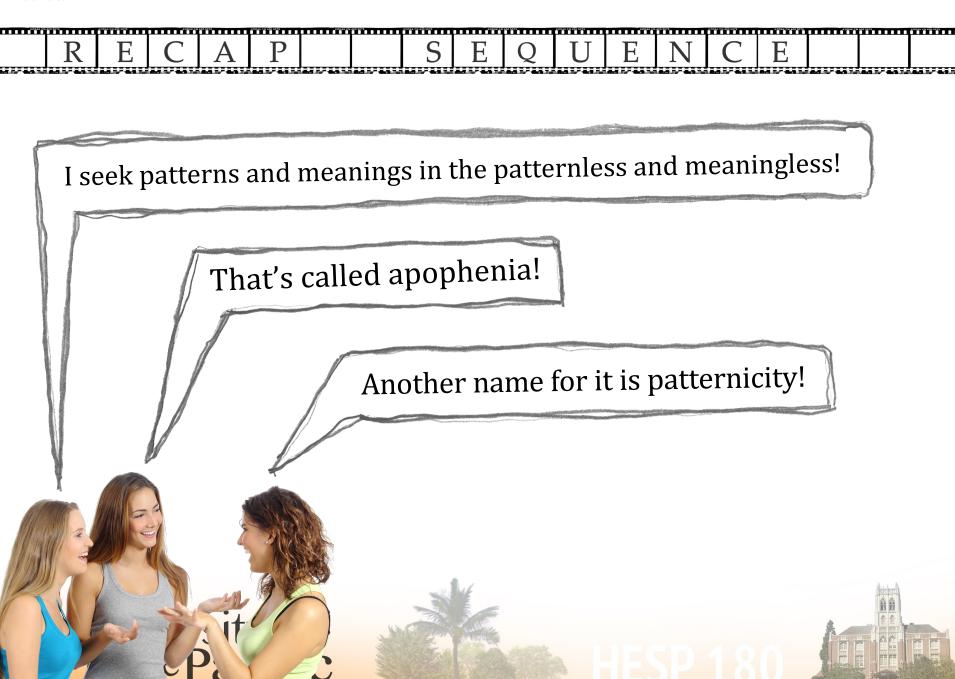




People are bad too (in terms of accuracy), but we do it anyway.

My sister said she'd pick me up at noon. She just got here. The movie starts in about three minutes. I'm furious.

It's easy for us to figure out why she's pissed. Not easy for a computer (pissed because the movie starts in three minutes?).



A human brain seeks causal relationships.

We seek (and find, however falsely) a cause where none exists.

- 1. You went out to eat.
- 2. You got sick the next day.
- 3. It must have been the food.
- 1. You went outside in the cold rain and didn't bundle up well enough.
- 2. You got the flu.

(That's not how the flu is transmitted, but people connect the two anyway.)

Incorrect causal intuitions. That's what you want to avoid.







- **1.** Take a population.
- 2. Observe them (their diets, habits, behaviors, etc.).
- 3. Notice patterns related to health and disease of that population.
- 4. Don't change anything; just document all the things you notice.
- **5.** Make conjectures / surmise causes.
- **6.** Publish wrong stuff.







# I DECASP) SEQUINCE 1010(C) SEQUINCE 1010(C) SEQUINCE

# Major epidemiological domains:

- Origin/outbreak of disease
- Disease transmission
- Screening and biomonitoring
- Comparisons of treatment effects





# Overarching purpose:

- Who?
- What?
- Where?
- When?

of health-related events.







## Traditional and Modern Epidemiology

... a shift in the level of analysis from the population to the individual. Most modern epidemiologists still do studies in populations, but they do so in order to study decontextualized individual risk factors, rather than to study population factors in their social and historical context. American Journal of Public Health May 1996, Vol. 86, No. 5

Traditional Epidemiology, Modern Epidemiology, and Public Health

Neil Pearce, PhD







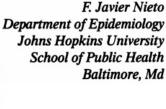
Post-Snow epidemiology was an attempt to apply science to the world around us:

American Journal of Public Health

March 1999, Vol. 89, No. 3

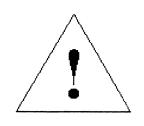
Cardiovascular Disease and Risk Factor Epidemiology: A Look Back at the Epidemic of the 20th Century







Post-Snow epidemiology was an attempt to apply science to the world around us:



GOVERNMENT WARNING: (1) According to the Surgeon General, women should not drink alcoholic beverages during pregnancy because of the risk of birth defects. (2) Consumption of alcoholic beverages impairs your ability to drive a car or operate machinery, and may cause health problems.





hunder bird

Epidemiology continues to have its place: It tries to make sense of epidemics.

**Proposed in California and New York:** 

STATE OF CALIFORNIA SAFETY WARNING: Drinking beverages with added sugar(s) contributes to obesity, diabetes, and tooth decay.



Is It Time For A Warning Label On Sugar-Loaded Drinks?

PONCIE RUTSCH Published April 9, 2015 · 3:52 PM ET







RECAPSEQUENCE

Breast Cancer Vol. 5 No. 4 October 1998

#### Risk Factors for Breast Cancer among Japanese Women: A Case-Control Study in Ibaraki, Japan

Masaru Ueji\*1, Ei Ueno\*2, Douglas Osei-Hyiaman\*1, Tomoko Saito\*1, Hideto Takahashi\*3, and Katsumi Kano\*3

Results: After adjustment for potential confounders, heavy weight and higher body mass index were associated with an increased risk of breast cancer among postmenopausal women (OR=1.76, 95% CI=0.69, 4.48; OR=1.57, 95% CI=0.61, 3.99, respectively). Current or ex-smokers were found to be at an increased risk for breast cancer (OR=3.33; 95% CI=1.63, 6.80). Women who take hot baths had a decreased risk for breast cancer (OR=0.67; 95% CI=0.43, 1.06). Recreational physical activity was associated with a reduced risk of breast cancer (PTrend=0.005). OR for breast cancer among physically active women was 0.36 (95% CI=0.19, 0.70), as compared with inactive women. Taller women had an increased risk of breast cancer relative to shorter women (OR=1.49; 95% CI=0.83, 2.70). No significant association between alcohol consumption and breast cancer risk was detected.

Conclusions: Our results suggest that several potentially modifiable lifestyle factors may be useful for the prevention of breast cancer.











An IRB is unlikely to approve a study in which participants are randomly assigned to one of two groups:

**Group A:** Smoke-free control

**Group B:** Two packs per day.

So epidemiological principles are used.





# Smoking and cancer.

The relationship between snuff and nasal cancer was identified as early as 1761:

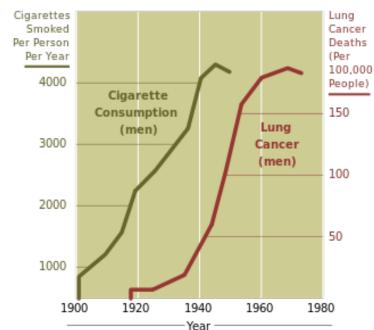
Redmond DE JR. Tobacco and cancer: the first clinic report, 1761.

N Engl J Medicine 1970;282:18-23.





#### 20-Year Lag Time Between Smoking and Lung Cancer



Fritz Lickint evaluated time-trend data, which implicated smoking as a cause of lung cancer.



Lickint F. Der Bronchialkrebs der Raucher. Munch Med Wschr 1935; 82:122-24.







RECAPISEQUENCE

#### Findings supported by case-control studies in the US and UK:

Doll R, Hill AB. Smoking and carcinoma of the lung. Preliminary report. *Br Med J* 1950;**2:**739–48.

Levin ML, Goldstein H, Gerhardt PR. Cancer and tobacco smoking. Journal of the American Medical Association 1950;143:336–38.

Mills CA, Porter MM. Tobacco smoking habits and cancer of the mouth and respiratory system. Cancer Research 1950;10:539-42.

Schrek R, Baker LA, Ballard GP, Dolgoff S. Tobacco smoking as an etiologic factor in disease. *Cancer Research* 1950;**10:**49–58.

Wynder EL, Graham EA. Tobacco smoking as a possible etiologic factor in bronchogenic carcinoma. *Journal of the American Medical Association* 1950;**143**:329–36.







This is where we wind up:

SURGEON GENERAL'S WARNING: Smoking Causes Lung Cancer, Heart Disease, Emphysema, And May Complicate Pregnancy.









But we have to be careful:



# Maternal Grandmother's Smoking Correlated With Autism Diagnosis

40.5K





Share on Twitter





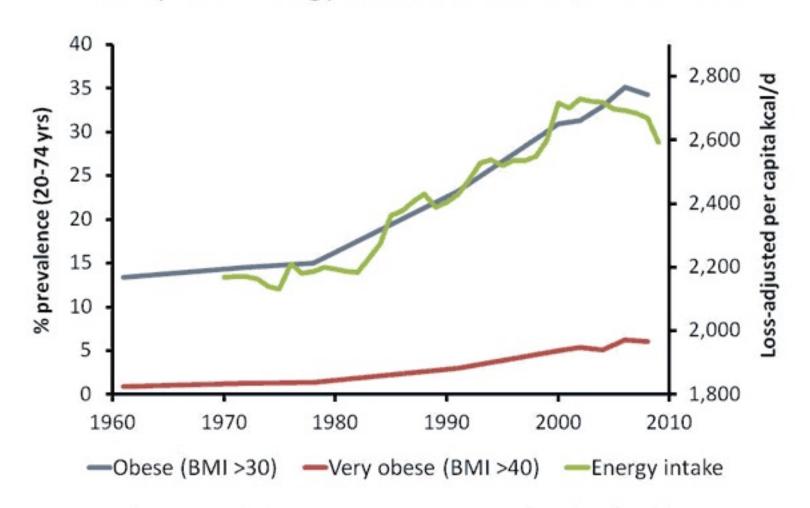


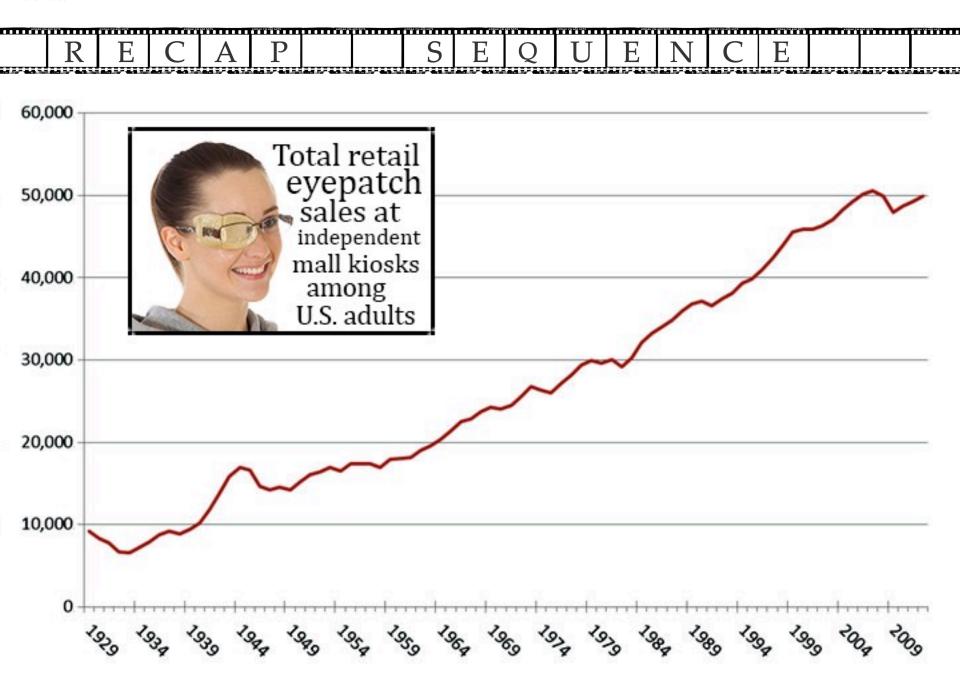




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### Obesity and Energy Intake in the US, 1961-2009





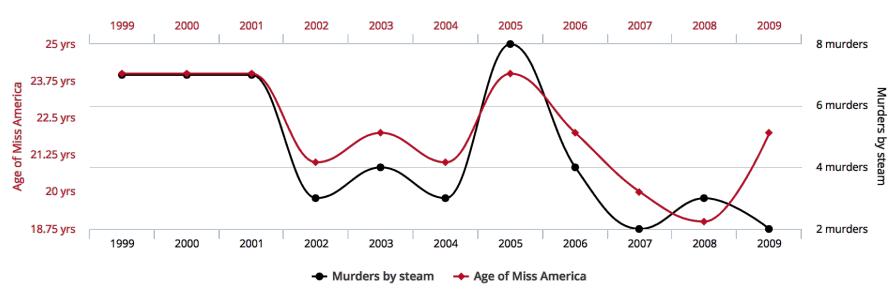
www.tylervigen.com/spurious-correlations



# Age of Miss America correlates with

#### Murders by steam, hot vapours and hot objects

Correlation: 87.01% (r=0.870127)



Data sources: Wikipedia and Centers for Disease Control & Prevention







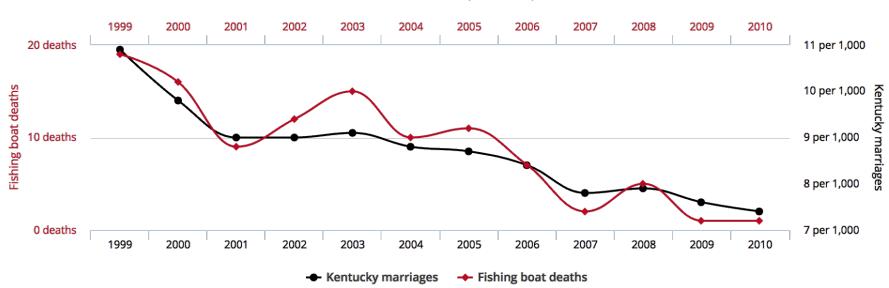
www.tylervigen.com/spurious-correlations



# People who drowned after falling out of a fishing boat correlates with

#### Marriage rate in Kentucky

Correlation: 95.24% (r=0.952407)



Data sources; Centers for Disease Control & Prevention and National Vital Statistics Reports





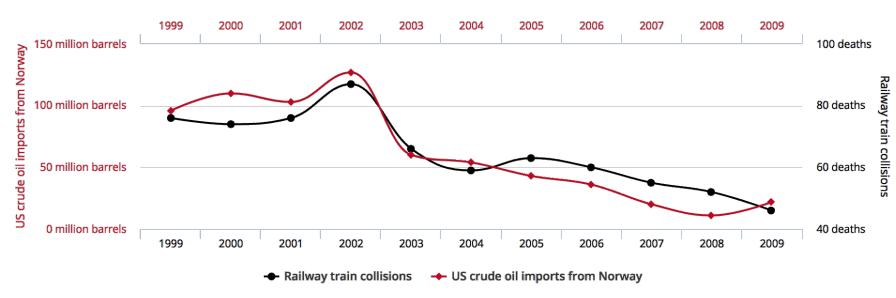
www.tylervigen.com/spurious-correlations



# US crude oil imports from Norway correlates with

#### Drivers killed in collision with railway train

Correlation: 95.45% (r=0.954509)



Data sources: Dept. of Energy and Centers for Disease Control & Prevention



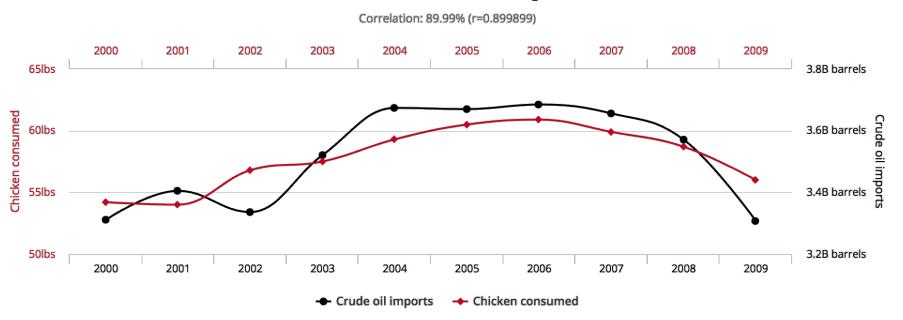


i www.tylervigen.com/spurious-correlations



## Per capita consumption of chicken correlates with

#### Total US crude oil imports



Data sources: U.S. Department of Agriculture and Dept. of Energy







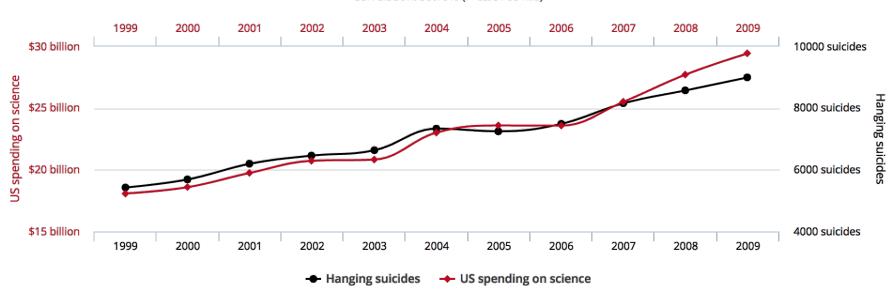
www.tylervigen.com/spurious-correlations



## US spending on science, space, and technology correlates with

#### Suicides by hanging, strangulation and suffocation

Correlation: 99.79% (r=0.99789126)



Data sources: U.S. Office of Management and Budget and Centers for Disease Control & Prevention





www.tylervigen.com/spurious-correlations

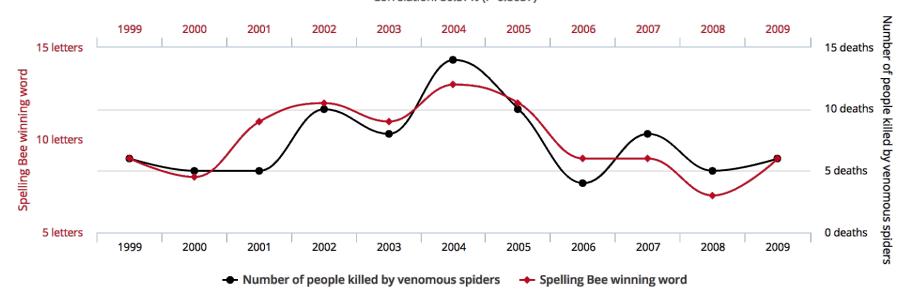


#### Letters in Winning Word of Scripps National Spelling Bee

correlates with

#### Number of people killed by venomous spiders

Correlation: 80.57% (r=0.8057)



Data sources: National Spelling Bee and Centers for Disease Control & Prevention





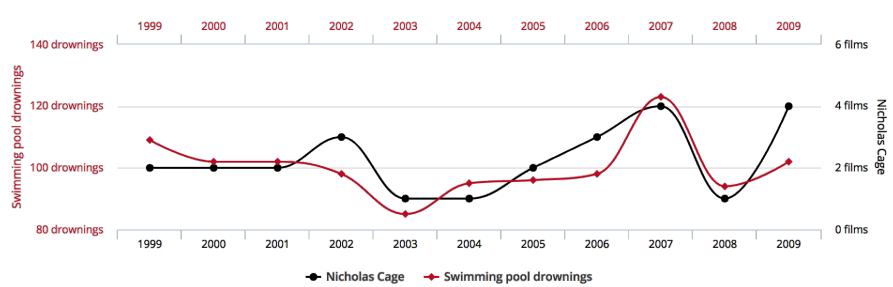
www.tylervigen.com/spurious-correlations



## Number of people who drowned by falling into a pool correlates with

#### Films Nicolas Cage appeared in

Correlation: 66.6% (r=0.666004)



Data sources; Centers for Disease Control & Prevention and Internet Movie Database





www.tylervigen.com/spurious-correlations

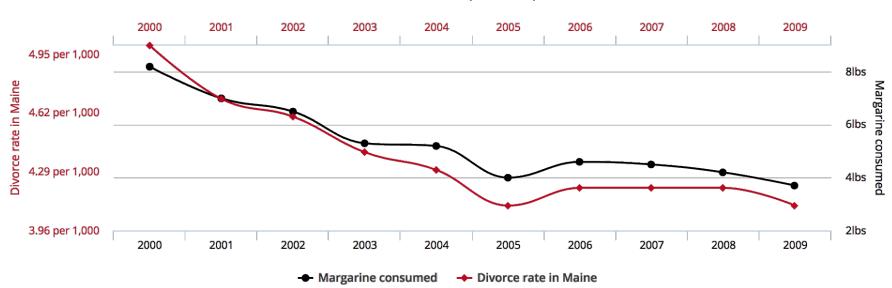


#### Divorce rate in Maine

correlates with

#### Per capita consumption of margarine

Correlation: 99.26% (r=0.992558)



Data sources: National Vital Statistics Reports and U.S. Department of Agriculture





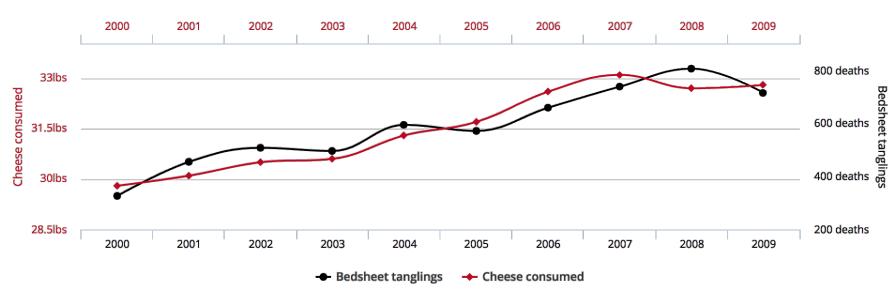
www.tylervigen.com/spurious-correlations



## Per capita cheese consumption correlates with

#### Number of people who died by becoming tangled in their bedsheets

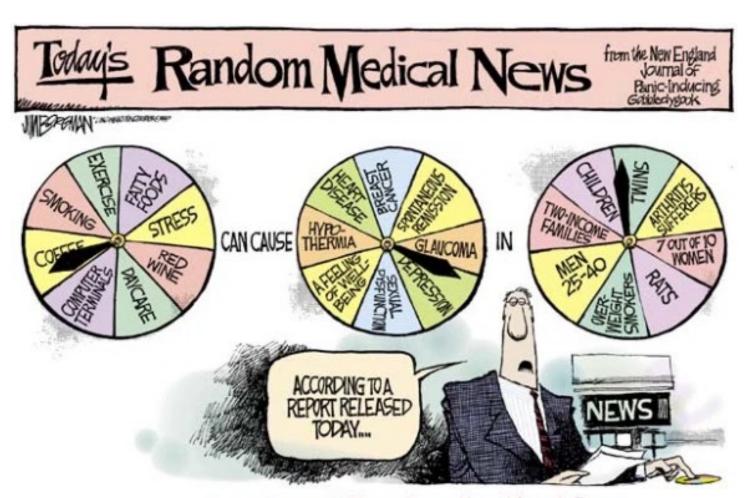
Correlation: 94.71% (r=0.947091)



Data sources: U.S. Department of Agriculture and Centers for Disease Control & Prevention







One view of the value of epidemiology

clofibrate magnésico chobet® clofibrate magnésico 400 mg PHRENOLOGY L.N. FOW LER PISCES Unive. of the Pacific SUINATTIDAS

#### A big clue:

Most comparisons in "meat studies" compare the bottom quintile (vegetarians) to the top quintile (serious carnivores).

As you move from the bottom to the top, what else do you see?



#### **Bad Box**

More smoking, drinking, more sedentary lifestyles, and pretty much every other bad behavior, in addition to high BMI, high BP, etc.

### **Good Box**

Vegetarians are more concerned with their health in a number of ways. More likely to visit the doctor, do yoga, stay hydrated, get more sleep; they probably brush their teeth more.



## RECAPSEQUENCE

Table 1. We have found 12 papers in which claims coming from observational studies were tested in randomised clinical trials. Many of the trials are quite large. In most of the observational studies multiple claims were tested, often in factorial designs, e.g. vitamin D and calcium individually and together along with a placebo group. Note that none of the claims replicated in the direction claimed in the observational studies and that there was statistical significance in the opposite direction five times

ID no.	Pos.	Neg.	No. of claims	Treatment(s)	Reference
1	0	1	3	Vit E, beta-carotene	NEJM 1994; <b>330</b> : 1029–1035
2	0	3	4	Hormone Replacement Ther.	JAMA 2003; 289: 2651-2662, 2663-2672, 2673-2684
3	0	1	2	Vit E, beta-carotene	JNCI 2005; <b>97</b> : 481-488
4	0	0	3	Vit E	JAMA 2005; <b>293</b> : 1338-1347
5	0	0	3	Low Fat	JAMA. 2006; 295: 655-666
6	0	0	3	Vit D, Calcium	NEJM 2006; <b>354</b> : 669-683
7	0	0	2	Folic acid, Vit B6, B12	NEJM 2006; <b>354</b> : 2764-2772
8	0	0	2	Low Fat	JAMA 2007; <b>298</b> : 289-298
9	0	0	12	Vit C, Vit E, beta-carotene	Arch Intern Med 2007; 167: 1610-1618
10	0	0	12	Vit C, Vit E	JAMA 2008; <b>300</b> : 2123-2133
11	0	0	3	Vit E, Selenium	JAMA 2009; <b>301</b> : 39-51
12	0	0	3	HRT + Vitamins	JAMA 2002; <b>288</b> : 2431–2440
Totals	0	5	52		

12 RCTs tested 52 observational claims: 0 findings upheld; 10% found an effect in the *opposite* direction.



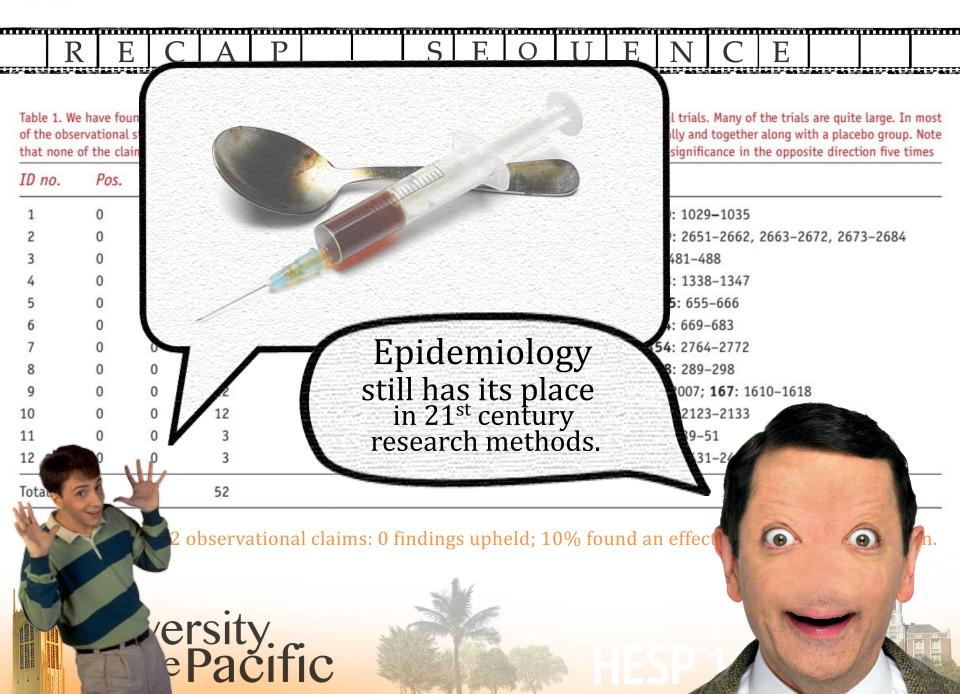




#### E DREDGING FOR GOLD - TYPICAL GOLD - DIVING OPERATION BUDDY DIVER , GASOLINE MOTOR (MAY OPERATE BOTH Table 1. We have foun l trials. Many of the trials are quite large. In most SUCTION AND AIR PUMPS ! of the observational st lly and together along with a placebo group. Note SLUICE that none of the clain significance in the opposite direction five times ID no. Pos. SUCTION DREDGE 0 : 1029-1035 : 2651-2662, 2663-2672, 2673-2684 0 481-488 0 GRAVEL OVERBURDEN : 1338-1347 GOLD DEEP IN CREVICES **5**: 655-666 0 DREDGE INTAKE : 669-683 0 BLACK SANDS 54: 2764-2772 Low Fat JAMA 2007; 298: 289-298 Vit C, Vit E, beta-carotene Arch Intern Med 2007; 167: 1610-1618 Vit C, Vit E 10 JAMA 2008; 300: 2123-2133 11 Vit E, Selenium JAMA 2009; 301: 39-51 HRT + Vitamins 12 JAMA 2002; 288: 2431-2440 52 Total 2 observational claims: 0 findings upheld; 10% found an effect in the *opposite* direction.

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Pacific

**HESP 180** 

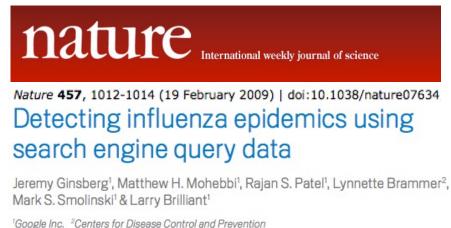


## 21st century epidemiology:



## RECAP SEQUENCE





Google receives 3 billion search queries every day and saves (and analyzes) them all. Analyses of those queries engenders a new form of public health surveillance.







Google

Vs.



H1N1 outbreak. How bad is it? Where is it spreading?

#### Traditional mode of reporting:

- 1. Patients would wait a while and then go to the doctor.
- 2. Doctors would wait a while and then report the cases.
- 3. The CDC would wait a while and then tabulate all cases.

All CDC reports were at least a week old, probably two.

.







Google

Vs.



H1N1 outbreak. How bad is it? Where is it spreading?

"Big data" mode of reporting:

Google evaluates search terms and analyzes for terms that have a strong correlation with the spread of disease. They're able to watch and report the spread in real time, not a week or two late.







### WIRED

DAVID LAZER AND RYAN KENNEDY SCIENCE 10.01.15 7:00 AM

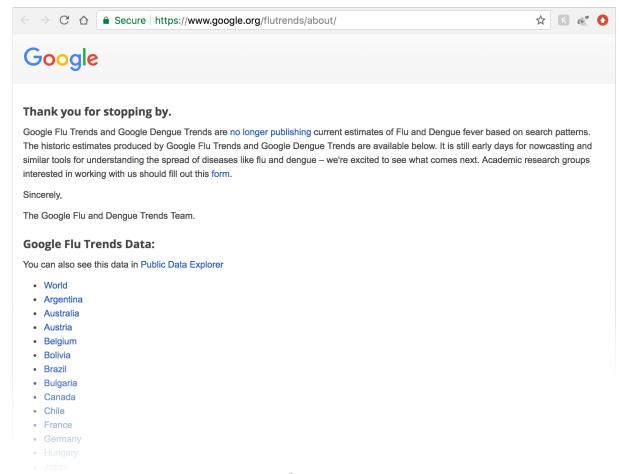
## WHAT WE CAN LEARN FROM THE EPIC FAILURE OF GOOGLE FLU TRENDS

And then, GFT failed—and failed spectacularly—missing at the peak of the 2013 flu season by 140 percent. When Google quietly euthanized the program, called Google Flu Trends (GFT), it turned the poster child of big data into the poster child of the foibles of big data. But GFT's failure doesn't erase the value of big data. What it does do is highlight a number of problematic practices in its use—what we like to call "big data hubris." The value of the data held by entities like Google is almost limitless, if used correctly. That means the corporate giants holding these data have a responsibility to use it in the public's best interest.















Digital information that can be "crunched" is relatively new.



**1986:** ~40% of the world's general purpose computing power was in pocket calculators. At the time, pocket calculators had more processing power than all personal computers combined.





#### Oren Etzioni

First undergrad to major in Computer Science at Harvard (graduated 1986).

Ph.D. at Carnegie Mellon University (1991)

Currently a professor at the University of Washington (director of the Turing Center) <a href="https://www.cs.washington.edu/people/faculty/fulltime">https://www.cs.washington.edu/people/faculty/fulltime</a> and <a href="https://turing.cs.washington.edu/">http://turing.cs.washington.edu/</a>

Current CEO of the Allen Institute for Artificial Intelligence (research institute funded by Microsoft Co-Founder Paul Allen). Here: <a href="http://oren-etzioni.people.allenai.org/">http://oren-etzioni.people.allenai.org/</a>



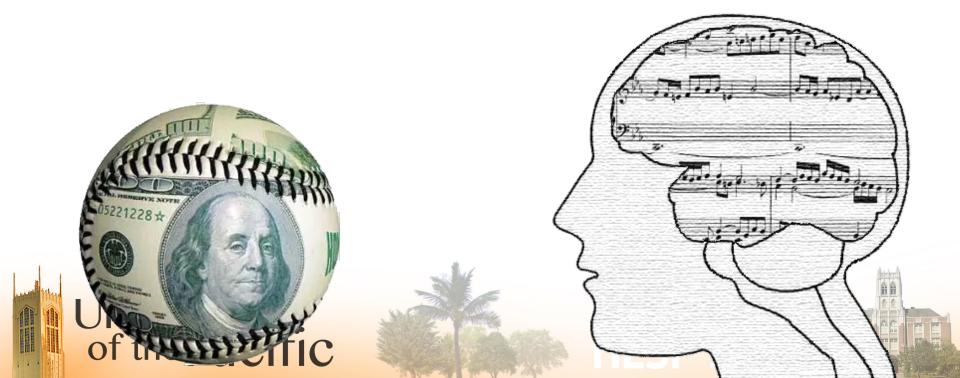




## Music, Sports, and the Entertainments...

Music is less art than science now.

Sports are almost completely data driven.



### The New York Times

**BUSINESS DAY** 

#### What Wal-Mart Knows About Customers' Habits

By CONSTANCE L. HAYS NOV. 14, 2004

By its own count, Wal-Mart has 460 terabytes of data stored on Teradata mainframes, made by NCR, at its Bentonville headquarters. To put that in perspective, the Internet has less than half as much data, according to experts.





R E C A P S E

# THE END OF THEORY: THE DATA DELUGE MAKES THE SCIENTIFIC METHOD OBSOLETE

CHRIS ANDERSON

06.23.08 12:00 PM

Chris Anderson (*Wired* Editor–in-Chief) argues "big data" (epidemiology) is all you need. In a data driven world, there's no need for theories.

The Petabyte Age is different because more is different. Kilobytes were stored on floppy disks. Megabytes were stored on hard disks. Terabytes were stored in disk arrays. Petabytes are stored in the cloud.

Scientific method replaced by statistical correlations, devoid of theory.







# WIRED

R E C A P S E

# THE END OF THEORY: THE DATA DELUGE MAKES THE SCIENTIFIC METHOD OBSOLETE

CHRIS ANDERSON

06.23.08 12:00 PM

This is a world where massive amounts of data and applied mathematics replace every other tool that might be brought to bear. Out with every theory of human behavior, from linguistics to sociology. Forget taxonomy, ontology, and psychology. Who knows why people do what they do? The point is they do it, and we can track and measure it with unprecedented fidelity. With enough data, the numbers speak for themselves.







# WIRED

R E C A P S E

# THE END OF THEORY: THE DATA DELUGE MAKES THE SCIENTIFIC METHOD OBSOLETE

CHRIS ANDERSON

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There is now a better way. Petabytes allow us to say:
"Correlation is enough." We can stop looking for models. We
can analyze the data without hypotheses about what it
might show. We can throw the numbers into the biggest
computing clusters the world has ever seen and let
statistical algorithms find patterns where science cannot.







Is this a good position? Is "big data" all we need?







## Epidemiology is very useful to answer some questions:

What conditions (sleeping position, etc.) maximize the odds of SIDS?

How much asbestos exposure is necessary to induce mesothelioma?

How many volts are necessary to ensure death by electrocution in well-behaved prepubescent children who love balloons?

What else? I might ask you to invent a question on a test (that we haven't previously discussed) that would be best investigated by epidemiology.





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You can only study what you can measure.

You can only correct for what you can measure.

You can only make claims about what was measured and controlled for.

Epidemiology isn't doing that.







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### INTRODUCTION: JOIN THE SCIENTIFIC CONVERSATION

When it is time for you to design, complete, and share the findings of your own research project, whether you are a graduate or advanced undergraduate student, you are inserting yourself into the conversation. A seamless transition requires that you first learn what has been discussed in the previous conversation. Fortunately, the historical transcripts are archived in the previous conversation. By taking the time to familiarize yourself with the existing lit to be asked in your research project.

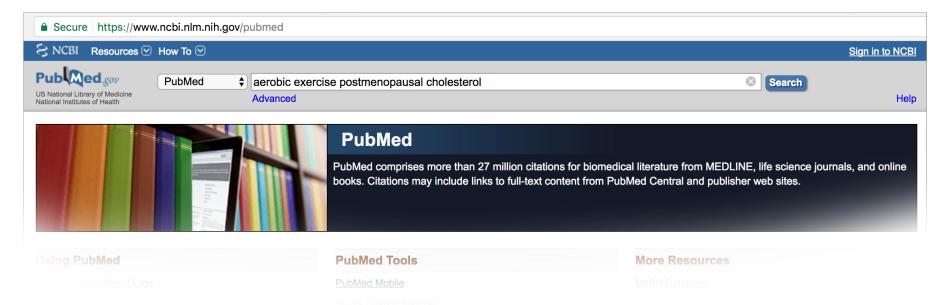
Where do I fit in to all of this

University of the Pacific



We already talked about that! You're being a bore. Either say something new or remain in life's audience.





Once your topic is sufficiently narrowed, and all of your variables are identified, start exploring the currently-existing literature. This will require a lot of work. It won't just be a few hours. Count on it taking nearly forever.

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of the Pacific



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The foundation of a good research project is a good **research question**, which originates from a strong understanding of the past and current literature in your field. When you have identified a field of study that interests you, it is then your duty as a researcher to become familiar with the findings of other researchers in your field. One of the benefits that comes with dedicating time to aggressively learning the relevant literature is that you will begin to detect **gaps in the literature**, and you will inevitably find yourself asking a question about a phenomenon in the field that has yet to be answered.





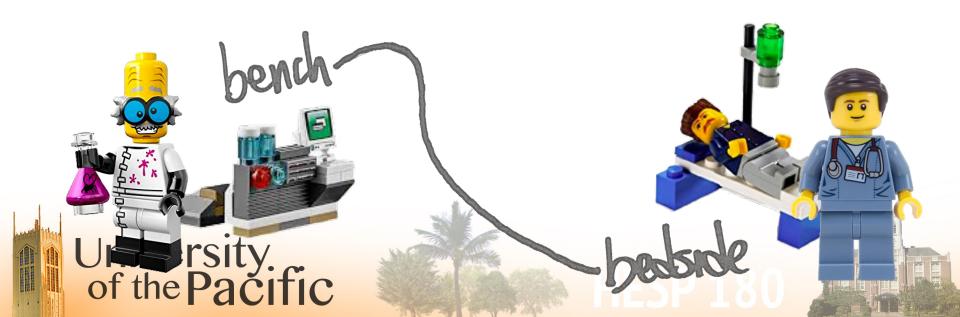




## Where in the continuum do your gaps reside?

#### CHAPTER 6 Understanding Research

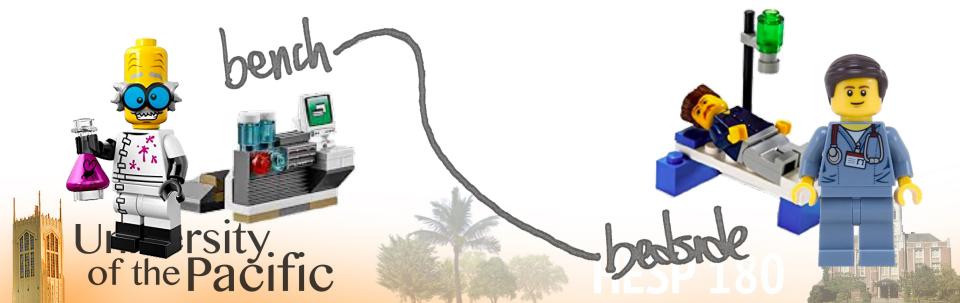
94 However, Hoover's 20th century dichotomy has expanded into a much broader gamut. It has evolved into a continuum that begins at bench science (animal models, human labs, etc.) and ends at the bedside (research aimed at informing clinical decisions).



## Where in the continuum do your gaps reside?

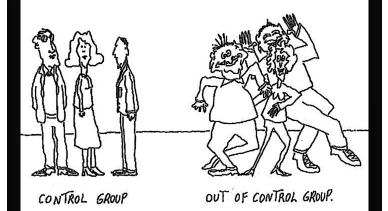
**Basic (or pure) research:** Bench science. You're a microbiologist working with Petri dishes. You're a chemist who uses Bunsen burners and thinks about petroleum. You're a geneticist studying adaptations in fruit flies. Originally limited to university settings.

**Applied research:** Research that translates into "best practices". Originally an "industry" thing.



Gaps are bound to exist where data collection is difficult to accomplish.

Well-controlled studies need control groups and few pro athletes (or Olympians) are willing to be in the control.



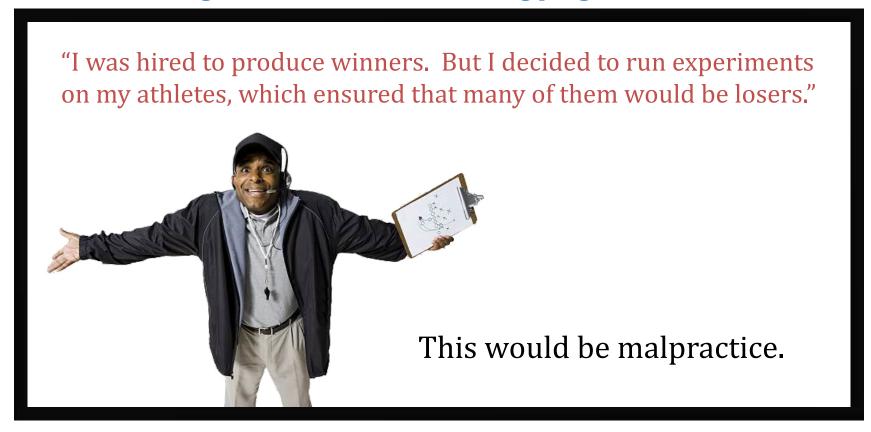
"We're testing a conditioning program that should be the most effective thing you can do to prepare for your sport. We need 60 world-class volunteers. We'll randomize 20 to the group that gets the program we think is best; we'll randomize another 20 to a comparison program, which we believe will be a bit less effective; and the remaining 20 will be a control group; none of them will be allowed to exercise for the duration of the experiment."

Why does this not happen?





There's a shortage of evidence-based training programs for elite athletes.







### Internal vs. external validity.

**Internal validity** is soundness of one's methods (controlling for extraneous variables that might confound findings). With perfect internal validity, your claim is very probably true, but a narrow question with a narrow answer.

**External validity** is the ability to generalize. "Zumba causes a drop in body fat" (or whatever). Broad question, broad answer.

#### **Consider blood pressure:**

On average, exercise reduces blood pressure by 4 to 9 mmHg.

But if you look at the individual people who make up that average, you'll see a different story.







Internal vs. external validity.



**Diseases and Conditions** 

## **High blood pressure (hypertension)**

Becoming more active can lower your systolic blood pressure — the top number in a blood pressure reading — by an average of 4 to 9 millimeters of mercury (mm Hg). That's as good as some blood pressure medications. For some people, getting some exercise is enough to reduce the need for blood pressure medication.



If you regularly perform aerobic exercise, will that lower your blood pressure?





723 subjects.

~400 responded to exercise with reductions in SBP.

Mean and median changes were reductions in SBP. Thus "exercise reduces blood pressure!"

> 300 subjects experienced an *increase* in SBP.

#### Current Hypertension Reviews, 2013, Vol. 9, No. 2

The Promises and Challenges of the Use of Genomics in the Prescription of Exercise for Hypertension: The 2013 Update

Garrett I. Ash<sup>a,\*</sup>, John D. Eicher<sup>b</sup> and Linda S. Pescatello<sup>a</sup>

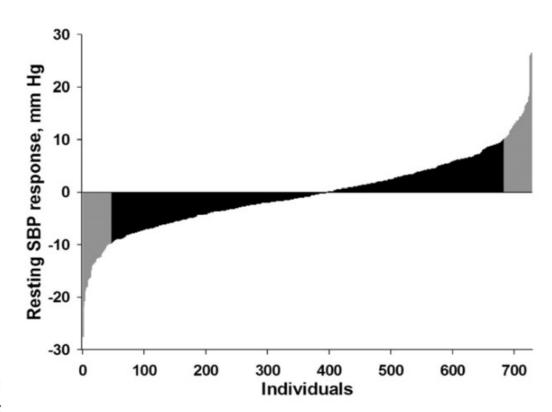


Fig. (6). Distribution of response to the HERITAGE exercise program with adverse and favorable responders highlighted in gray. SBP, systolic blood pressure. Adapted with permission from Bouchard C, Blair SN, Church TS, Earnest CP, Hagberg JM, Hakkinen K, et al. Adverse metabolic response to regular exercise: is it a rare or common occurrence? PLoS One 2012; 7(5): e37887.







12% of subjects experienced blood pressure increases of at least 10 mmHg.

Similar distributions were found to exist among these same subjects in insulin, cholesterol, and triglycerides.

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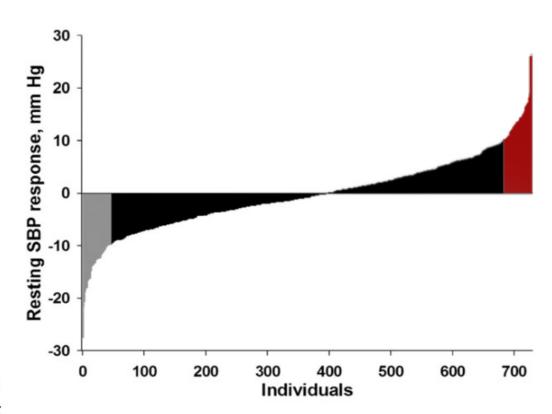
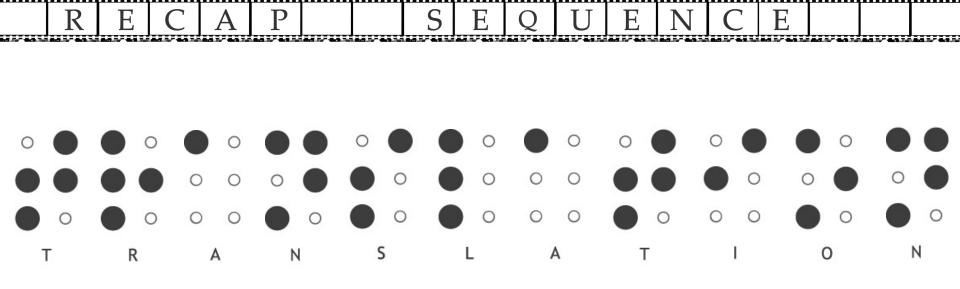


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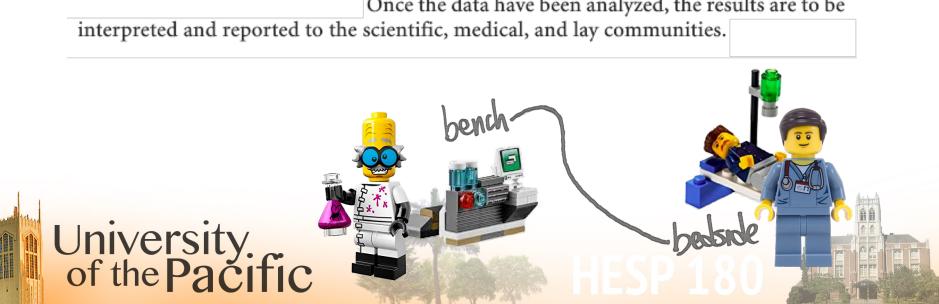




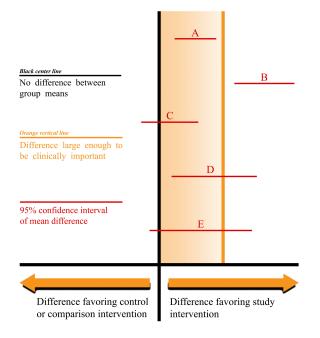


#### 94 Translation:

Once the data have been analyzed, the results are to be interpreted and reported to the scientific, medical, and lay communities.



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### **Magnitude of Effect**

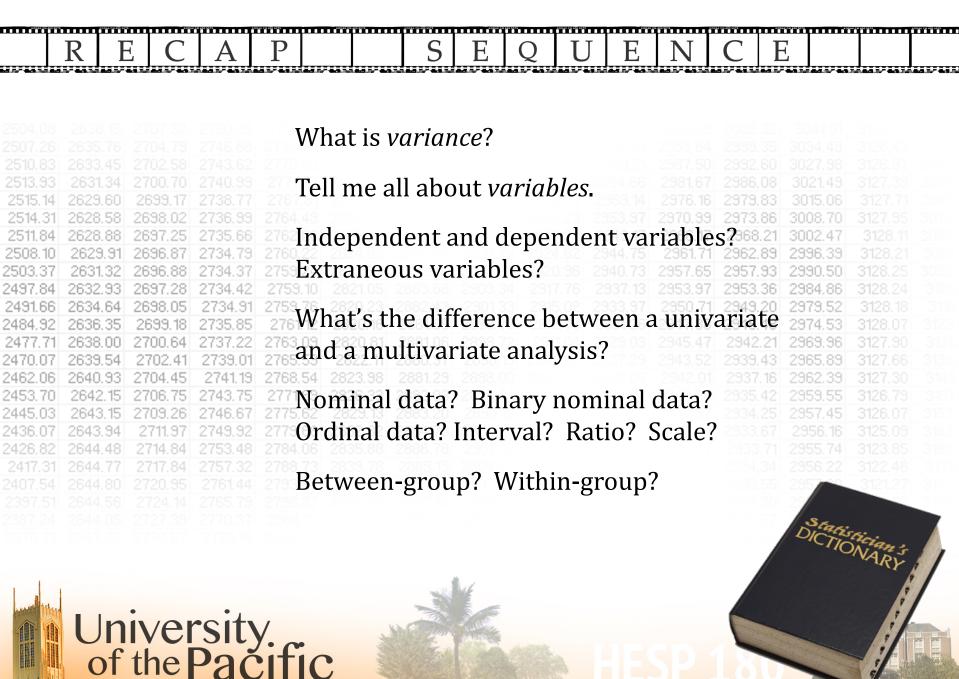
Not only is uncertainty not fully mitigated through inferential statistical analyses, it is often exacerbated. We attribute much of this problem to the use and misuse of the word "significant." It is not uncommon to read that investigators found "significant differences" between groups, suggesting a benefit of an intervention being studied or read about the "significant findings" of a study. The statistical meaning of significant is that the probability that a type I error has been committed is lower than the risk the investigators were willing to take. The word does not imply that the differences observed were of a magnitude worthy of attention.

Increasingly, the certainty or uncertainty of the magnitude of an effect is being conveyed through the reporting of **confidence intervals** (CIs). These statistics provide a range in which the population value is believed to lie at a given level of certainty. A 95% CI of mean differences provides knowledge (with 95% certainty) as to where a population mean difference may lie.





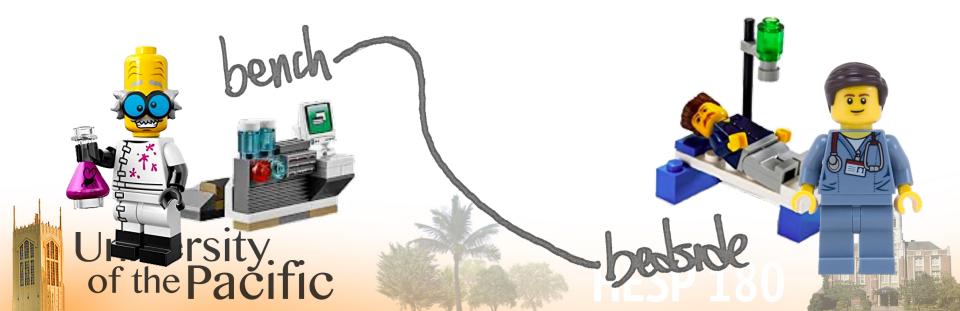




RECAPSEQUENCE

### CHAPTER 6 Understanding Research

- Evidence-based practice
- What are the differences between basic (pure) and applied research?
- What does "bench to bedside" mean?



## RECAPSEQUENCE

### GLOSSARY 118

**Biopsychosocial model:** a framework by which human health and disease is evaluated in terms of biologic, psychologic, and social factors (and the interplay between each)

**Confidence interval:** in statistics, a CI estimates a population parameter. After measuring a variable in a sample of that population and calculating the mean, one can create a range in which the true mean of the population is likely to fall. This range is the CI. Are you 99% confident the true population mean lies within a particular range? That's a 99% CI. Are you 90% confident? That's a 90% CI (which is likely to span a much broader range)

**Descriptive statistics:** a quantitative characterization of a particular set of information. Unlike inferential statistics, it does not seek to make inferences based on probability but merely to describe or summarize what exists (quantitatively)

**Frequentist:** the inferential alternative to Bayesian statistics; the framework that allows for the testing of a null hypothesis and the calculation of CIs

**Inferential statistics:** unlike descriptive statistics, which summarize and characterize information, inferential statistics make inferences about collections of data, testing predictions about the larger population based on the distribution of the collected sample

**Null hypothesis:** a statistical starting point for most scientific investigations, stating that there is no difference between two measurable phenomena. Researchers then collect and analyze data so that they can either confirm or reject this hypothesis







What's political arithmetic?

Remember all the stuff about variables...

What's epidemiology?

Describe epidemiology in the 19<sup>th</sup> century.

Describe epidemiology in the 20<sup>th</sup> century.

Today: epidemiology in the 20<sup>th</sup> and 21<sup>st</sup> centuries.

What are the three major types of epidemiological studies?







Every student has the exact same access to the exact same readings, slideshows, and course materials. Every student has the exact same access to the exact same lectures, office hours, and GAs/tutors. And every student takes the exact same test.

If you don't have a drink in your hand, it's your own fault.

Some students get As, some get Bs, some get Cs, others get Ds, and there might be an F or two. This distribution has everything to do with the student and nothing to do with the nature of instruction or examination. You are fully in control of your grade. And life.

University.
of the Pacific



I'm not an "easy A". And anyone who is doesn't care about you; they only care about themselves. Professors who assign As to students who didn't earn them do so for three reasons: 1) So you'll like them, 2) So you won't complain, and 3) So you'll give them friendlier student evaluations at the end of the semester. Issuing an easy A is a selfish and terrible thing to do.

Life is hard and unforgiving, you'll find. Much more difficult than school. It grades way harder than I do. Pretty much everyone in America has (or is getting) a college degree. No matter what you want to do with your life, you're going to have to outcompete huge crowds of educated, talented, motivated people. And it's nearly impossible. So if you spend your undergrad years comfortably coasting your way to As, there's little chance you'll find success after college. Life really is that hard. So anyone who gives you easy grades is not setting you up for success. My life would be easier (nobody would complain about anything) if I gave out As for free. But the future net worth of my students would suffer summin' fierce. So I won't do that. BUT: everything you need to get your A is in the slides and lectures, and I tell you which parts of the books are helpful. That doesn't make the A "easy", but it *definitely* makes it achievable. If you want it, it's yours. And it's useful to have.



