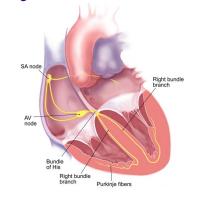


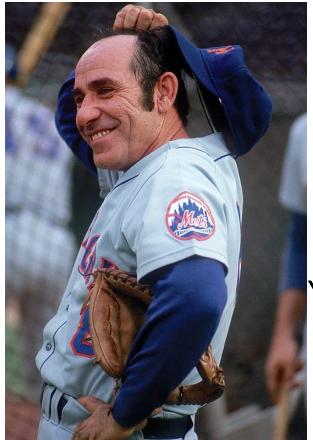
What causes my heart rhythm disorders?



Mark Drangsholt, D.D.S. Ph.D. Professor and Chair University of Washington Department of Oral Medicine <u>drangs@uw.edu</u>



"You can observe a lot by just watching"



Yogi Berra

Some of my self-tracking over past 12 years

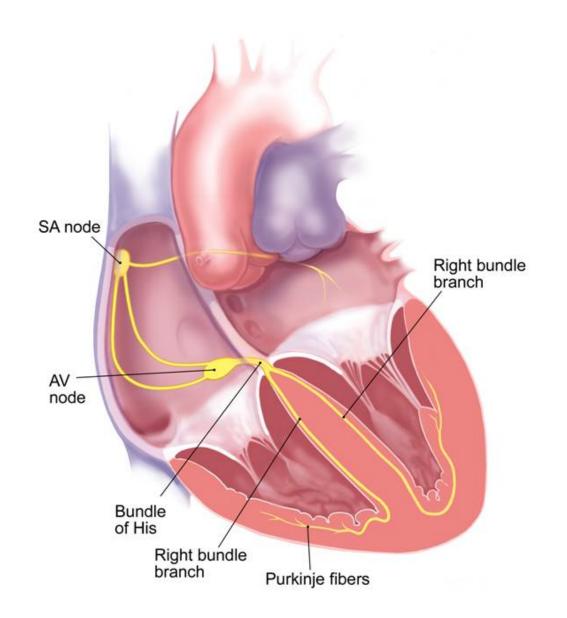
- Blood pressure vs. weight loss, exercise, medications
- Atrial fibrillation (cardiac rhythm disorder) vs. plausible transient daily triggers
- Deep sleep (ZEO) vs. aerobic exercise, caffeine, sex, other
- MyFitnessPal dietary tracking, exercise vs. body weight/percent body fat (Tanita).

Self-tracking over past 12 years

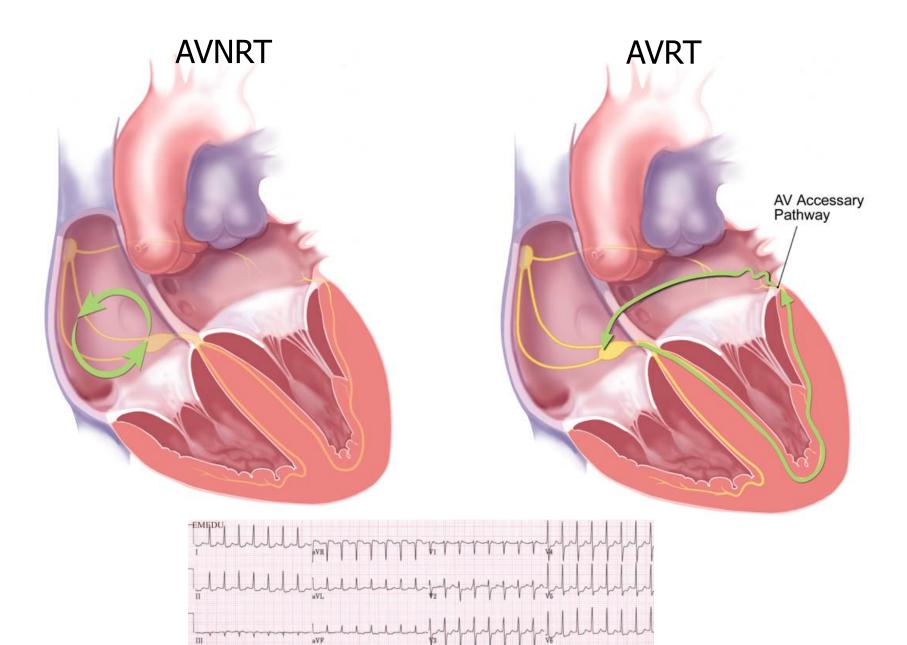
- Blood pressure vs. weight loss, exercise, medications
- Atrial fibrillation (cardiac rhythm disorder) vs. plausible transient daily triggers
- Deep sleep (ZEO) vs. aerobic exercise, caffeine, sex, other
- MyFitnessPal dietary tracking, exercise vs. body weight/percent body fat (Tanita).

Some Tachycardias (Heart Rate>100 beats per minute)

- Parosxymsal SupraVentricular Tachycardia (PSVT)- AV node Reentry (benign)
- PSVT AV Reentry (benign)
- Atrial flutter (relatively benign)
- Atrial fibrillation (stroke, heart failure)
- Wolff-Parkinson-White syndrome (sudden cardiac death)



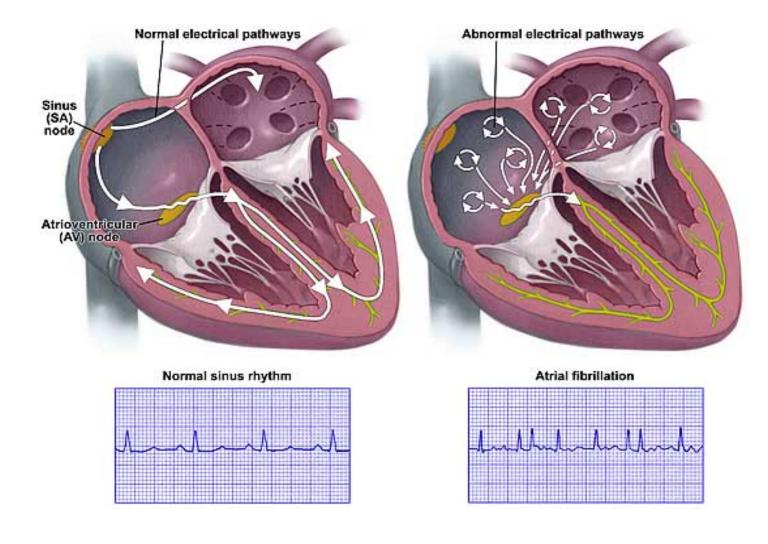
Normal electrical conduction through heart



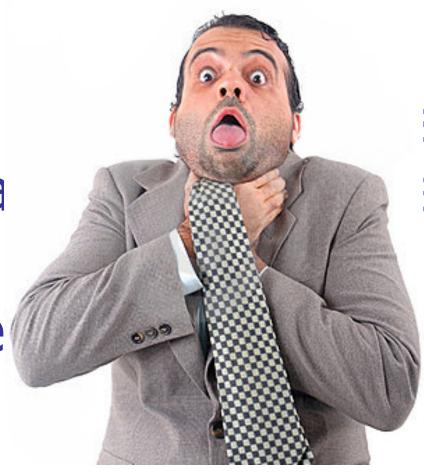
1-1-1-1-1-1-1-1

V1

7



Can I then a these dange



s, and ecrease

Created simple Excel table of all episodes for one year

Date	Time	Length	Where	Self dx	Onset	Offset	sympt oms

Est. Heart rate	Recorded with HRM?	Possible Triggers	comments				
	P ə lar.						
	LISTEN TO YOUR BODY						
			LAP 03-10- 04:10- * store				

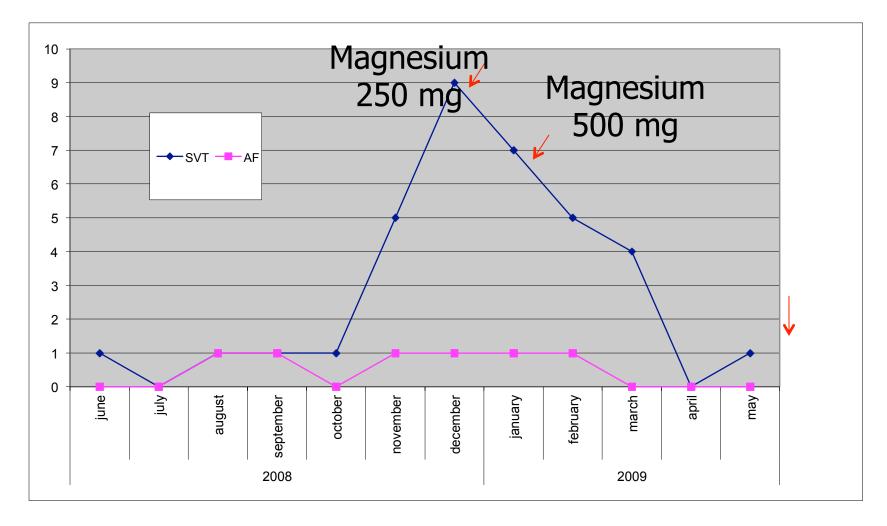
First, what type of arrhythmia?

- Used description of symptoms and signs to categorize as AF or PSVT
 - Choking feeling, irregular heart beat, impending doom, lightheaded - prodrome
 - Very rapid and regular heart beat, sometimes lightheaded

Attempted treatment trial

Magnesium supplement 250 mg per day

Episodes per month



Conclusion from trial

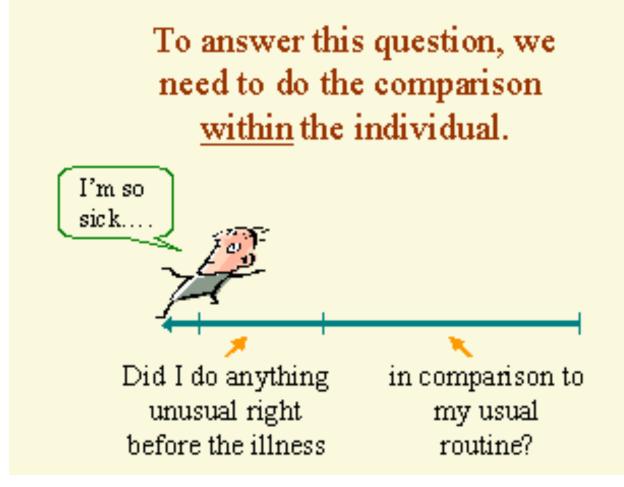
- 7 to 8 fold decrease in SVT
- No decrease in AF
- Other factors at play
 - Seasonality
 - Decreased exercise level
 - Compare to previous year?
 - No control

"The first principle is that you must not fool yourself, and you are the easiest person to fool"



Richard Feynman, PhD

What about the triggers? Can they be described? Can they be quantified? Case-Crossover Design is a scientific way to ask and answer the question that "Was the patient doing anything unusual just before the onset of the disease?"



It is a design that compares the exposure to certain agent during the interval when the event does not occur (control period), to the exposure during the interval when the event occurs (hazard period).



Control Data

The control data can be

- exposure information from a comparable time period; or
- (2). exposure information in the past according to the individual's usual frequency of exposure.

Analysis If the control data were from the past exposure

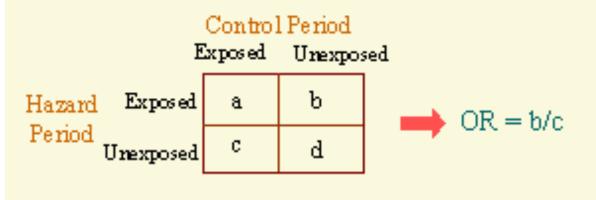
Step 1. Calculate the concurrence observed odds (a:b).

Odds that the exposure was during the hazard period right before the onset of disease.

(1:0) if there was exposure in hazard period.(0:1) if there was no exposure in hazard period.

Analysis If the control data were from a comparable control period

We will use the same method for a standard matched case-control study. But instead of case and control, we will have Hazard period and Control period.



SVT risk factors:

- Common events preceding attacks in the previous 1 hour:
 - High intensity exercise OR ~ 4.0
 - Afternoon caffeine OR ~ 3.0
 - Public speaking large groups OR ~ 3.0
- Common events in the previous 12 hours:
 - Inadequate sleep OR ~ 3.0

Atrial fibrillation risk factors

- Common events preceding attacks in the previous 2 hours:
 - Caffeine at any time
 - Air flight stress
 - >1 glass of wine
 - Public speaking large groups
- Common events preceding attacks in the previous 12+ hours:
 - Inadequate sleep

Take home message from tracking

- I used this information to minimize precipitants, and decreased the number of both types of episodes
- The additional information aided my cardiologists to make better decisions about which type of procedure to cure which rhythm problem

Where do self-tracking studies go in science, the levels of evidence?



Levels of Evidence: Treatment

Less bias 1 - 1 or more <u>randomized controlled</u> <u>trials</u>

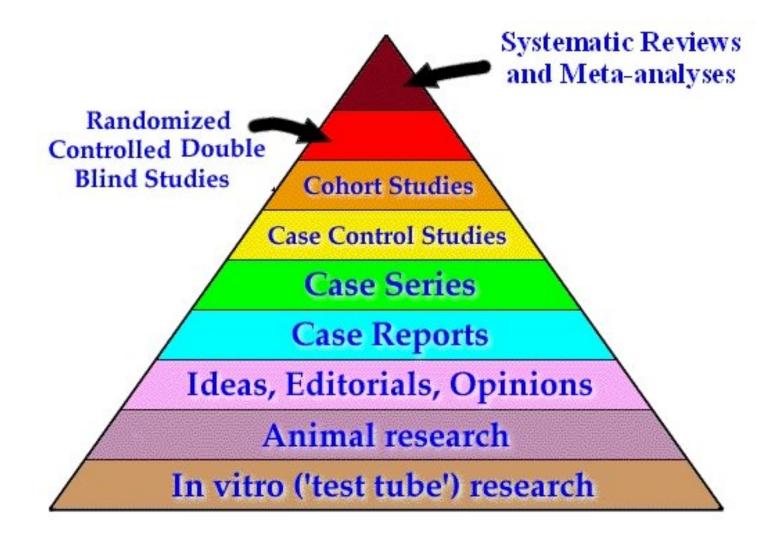
- **2** 1 or more <u>cohort studies</u>
- **3** 1 or more <u>case-control studies</u>

More bias 4 - 1 or more <u>case-series</u>

5 - <u>expert opinion</u> without above evidence

Bias = systematic error

The Evidence Pyramid: Types of Studies



Levels of Evidence: Groups of people

Less bias 1 - 1 or more <u>randomized controlled</u> <u>trials</u>

- **2** 1 or more <u>cohort studies</u>
- **3** 1 or more <u>case-control studies</u>

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5 - <u>expert opinion</u> without above evidence

Bias = systematic error

What type of study design is this information?

What is the scientific level of evidence?

Levels of Evidence: Within one person

Less bias **1** - 1 or more randomized controlled trials – "n of 1" trial

- 2 1 or more <u>cohort studies</u> -
- **3** 1 or more <u>case-crossover studies</u>

4 - 1 or more <u>case-series</u> -

- **5** <u>informed opinion</u> without above evidence
- *Bias = systematic error*

More bias

Summary

- Self tracking can aid in the identification of precipitating factors in a potentially life-threatening heart condition.
- Single subject designs as used by QS proponents deserve further study, validation and publication
- Single subject designs have their place in science and evidence-based medicine and

Thank you for your kind attention!

Mark Drangsholt drangs@uw.edu

Extra slides

AVRT risk factors

- ObesityMales
- Alcohol consumption
 Older than 40
- Electrolyte imbalance

Atrial flutter risk factors

- ObesityMales
- Alcohol consumption
 Older than 40
- Electrolyte imbalance

Known Atrial Fib risk factors

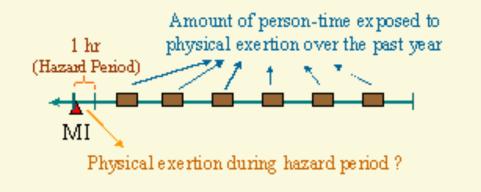
- Coronary heart disease
- Valve disease
- Inflamed heart muscle or lining
- Diabetes
- High blood pressure

AF risk factors in healthy people...

- Stressed or fatigued
- Too much caffeine or alcohol
- High intensity exercise
- Too little magnesium, potassium
- "big meal"

Example MI and physical exertion Control data 2:

The usual frequency of heavy physical exertion over the past year.



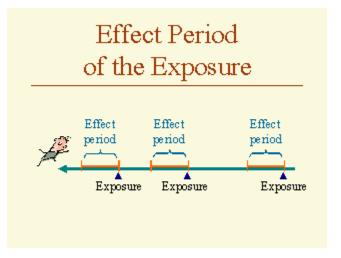
Case Crossover Design

- It is related to prospective crossover design.
- It is a matched <u>case-control</u> study but involves cases only and each individual serves as his/her own control.
- The data from a case crossover design can also resemble <u>cohort data</u> if the control data are units of person-time.

Effect Period of the Exposure

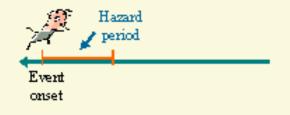
"... the period of altered risk in a population, to be the difference between the minimum delay before impact and the maximum carry-over time."

By Maclure M (AJE 1991;133:144-53)



Hazard Period

The period of time right before the onset of the event. Usually the length of haz ard period is the same as the length of the exposure effect period.



Control Data

The control data can be

- exposure information from a comparable time period; or
- (2). exposure information in the past according to the individual's usual frequency of exposure.

Example

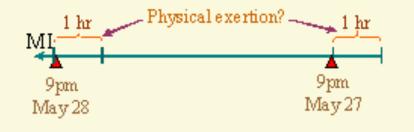
Triggering of MI by physical exertion (NEJM 1993;329:1677-83)

- Event: Myocardial Infarction (MI)
- Exposure: Heavy physical exertion
- Length of the exposure effect: 1 hour
- Hazard period: 1 hour before MI onset

2 sets of control data were used.

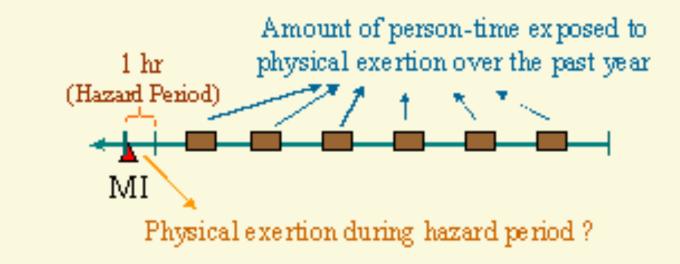
Example MI and physical exertion Control data 1:

The physical exertion information from a 1 hour period at the same time on the day before the onset of MI.



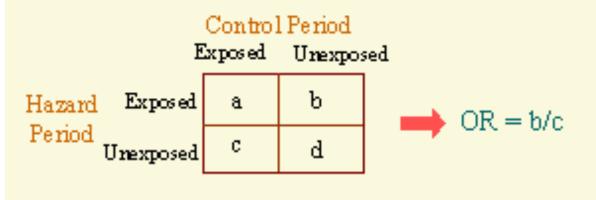
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Step 1. Calculate the concurrence observed odds (a:b).

Odds that the exposure was during the hazard period right before the onset of disease.

(1:0) if there was exposure in hazard period.(0:1) if there was no exposure in hazard period.

Analysis If the control data were from the past exposure

Step 2. Calculate the concurrence <u>expected odds</u> (x:y). Odds that a random event (disease) would have fallen in the effect period after

an episode of exposure.

Discussion Case Crossover Design

- The design can only be applied when the time lag between exposure and outcome is brief and the exposure must have little carryover effect.
- The results of this analysis are shortterm risks rather than cumulative risks.

Discussion Case Crossover Design

- Recall bias may occur during data collection.
- If there is within-individual confounding, stratification of the data may resolve the problem. To control several confounders simultaneously, conditional logistic regression can be used.

Bibliography of Case Crossover Design Application

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Petridou E, Mittleman MA, Trohanis D, et al. Transient exposures and the risk of childhood injury: a case-crossover study in Greece. Epidemiology 1998;9(6):622-5.

Meier CR, Jick SS, Derby, et al. Acute respiratory-tract infections and risk of first-time acute myocardial infarction. Lancet 1998;351(9114):1467-71.

Redelmeier DA, Tibshirani RJ. Interpretation and bias in case-crossover studies. Journal of Clinical Epidemiology 1997;50(11):1281-7.

Mittleman MA, Maldonado G, Gerberich SG, et al. Alternative approaches to analytical designs in occupational injury epidemiology. American Journal of Industrial medicine 1997;32(2):129-41.

Dixon KE. A comparison of case-crossover and case-control designs in a study of risk factors for hemorrhagic fever with renal syndrome. Epidemiology 1997;8(3):243-6.

Gullette EC, Blumenthal JA, Babyak M, et al. Effects of mental stress on myocardial ischemia during daily life. JAMA 1997;277(19):1521-6.

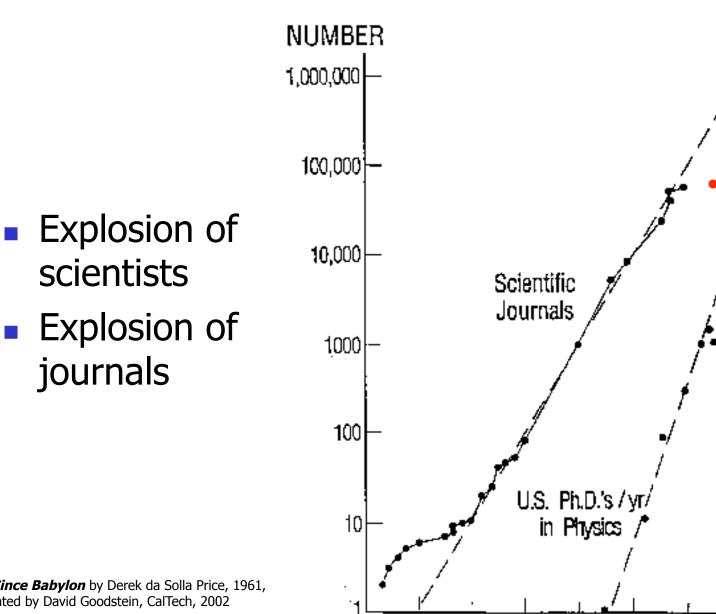
Mittleman MA, Maclure M, nachnani M, et al. Educational attainment, anger, and the risk of triggering myocardial infarction onset. The Department of Myocardial Infarction Onset Study Investigators. Archives of Internal Medicine 1997;157(7): 769-75.

Redelmeier DA, Tibshirani RJ. Association between cellular-telephone calls and motor vehicle collisions. New England Journal of Medicine 1997;336(7):453-8.

Muller JE, Mittleman A, Maclure M, et al. Triggering myocardial infarction by sexual activity. Low absolute risk and prevention by regular physical exertion. Determinants of myocardial infarction onset study investigators. JAMA 1996;275(18):1405-9.

Mittleman MA, Maclure M, Sherwood JB, et al. Triggering of acute myocardial infarction onset by episodes of anger. Circulation 1995;92(7):1720-5.

Vinson DC, Mabe N, Leonard LL, et al. Alcohol and injury. A case-crossover study. Archives of Family Medicine 1995;4(6): 505-11.



1700

1800

YEAR

1900

2000

Science Since Babylon by Derek da Solla Price, 1961, updated by David Goodstein, CalTech, 2002

Science or Evidence-Based Medicine

- Aim is to apply the best available evidence gained from the scientific method to clinical decision-making
- Method assesses the strength of evidence
- Randomized controlled trials or a summary of such trials (Systematic Review) is the highest level of evidence
- Expert opinion, or uncontrolled studies, are lowest

Levels of Evidence: Treatment

Less bias **1** - 1 or more *randomized controlled trials*

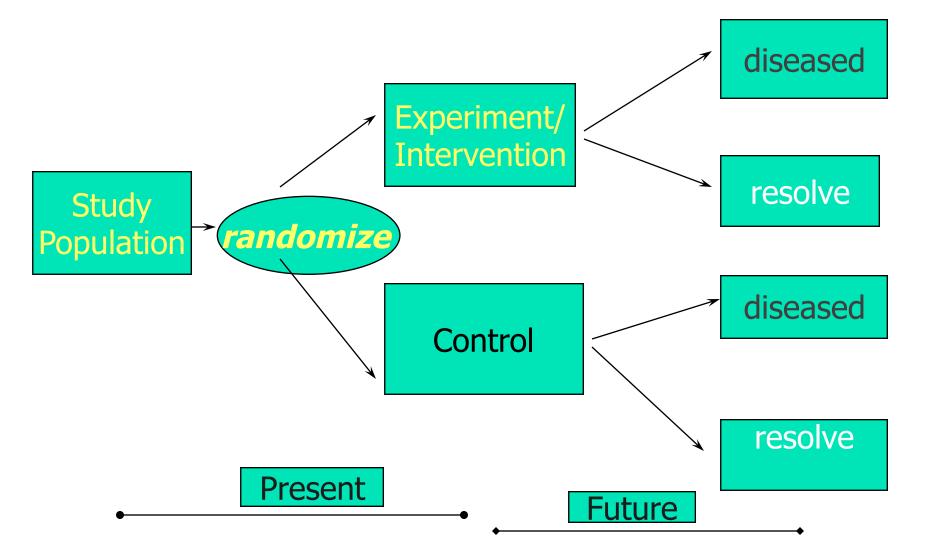
- **2** 1 or more *cohort studies*
- **3** 1 or more *case-control studies*

More bias **4** - 1 or more *case-series*

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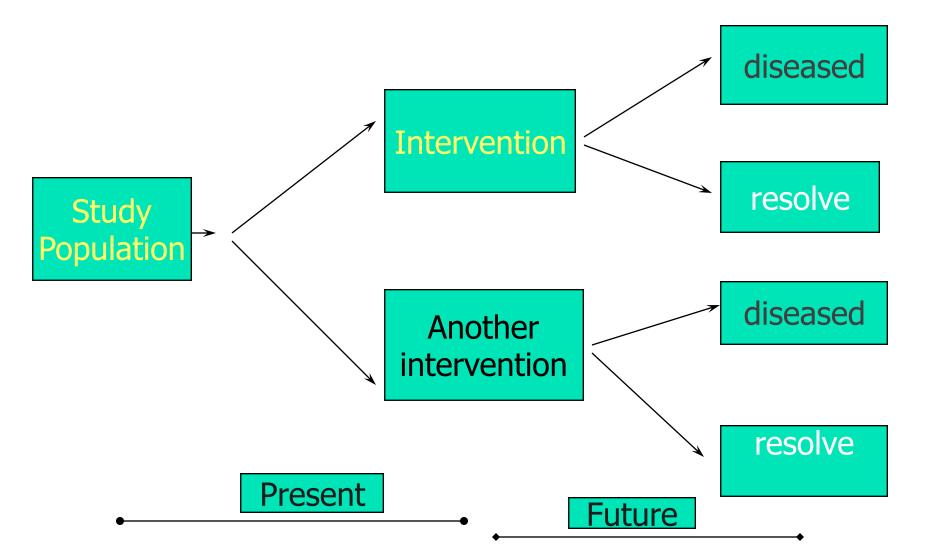
Experimental or Controlled Trial Design



Randomized Controlled Trials

- Resembles a true lab experiment
- Has a control group with an active or inactive intervention (placebo)
- Randomization creates equal groups on known AND unknown risk factors
- Neither patients nor observers commonly know which group they are in - blinding

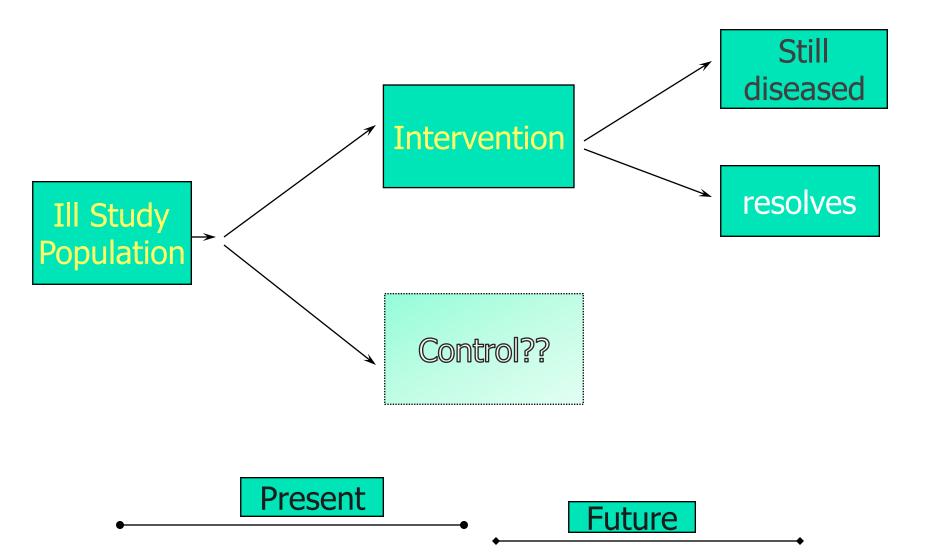
Controlled Trial or Cohort Design



Controlled Trials or cohort studies

- Some patients are given one treatment, some another
- No randomization groups are not necessarily equal – statistical analyses may be done to equalize groups
- Blinding not commonly done
- Time sequence is clear
- Second best design

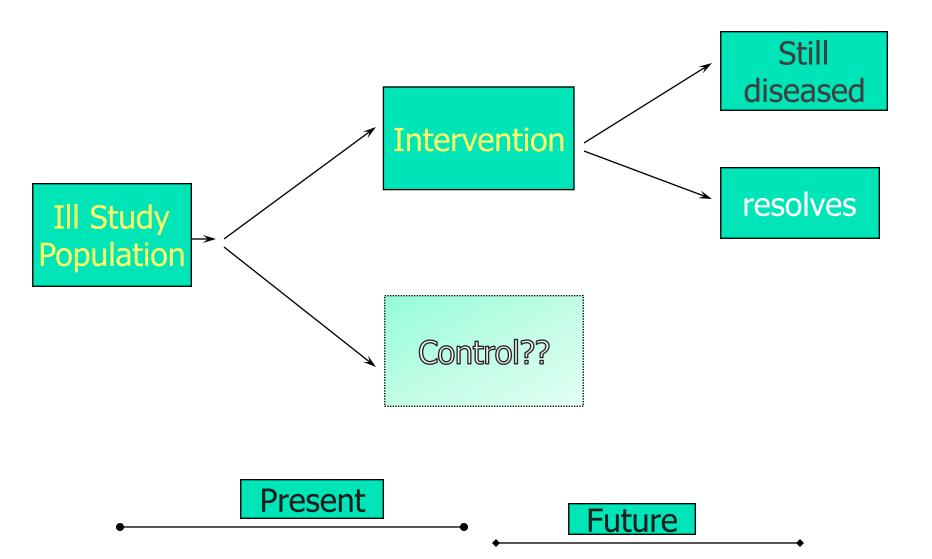
Case-series or follow-up of treated cases



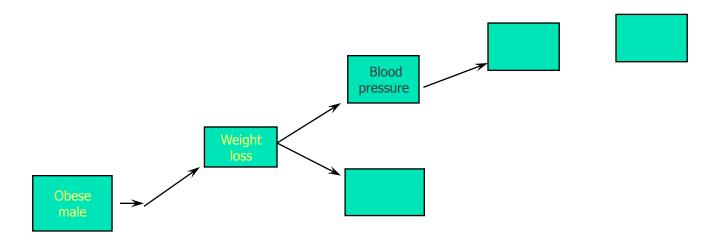
Case-series or follow-up of treated cases

- Common study design in clinical practice
- Lack of control group problematic esp. if clinical course of the condition is variable
- What would have happened in absence of treatment is not known
- Usually shows *inflated* estimates of true efficacy of therapy
- One of the weakest designs

Follow-up of treated case (self)



Follow-up of treated cases (self)





Other tests

- Shoe weight vs. running efficiency
- Heel strike/Forefoot vs. running efficiency
- Aero Helmet design, wheel design vs. bike speed.
- Gel use and cycling power output