

Disclosure

Nothing to disclose

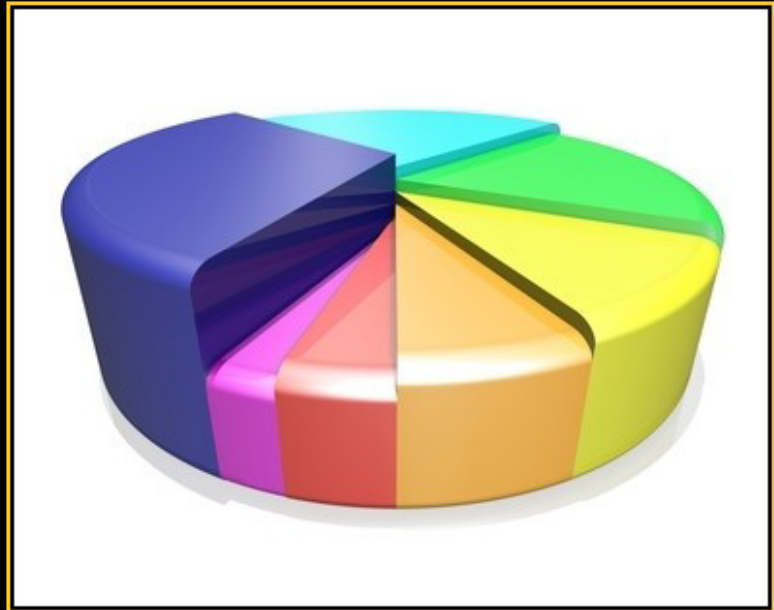


Why do I need to know statistics...?



STATISTICS

60 percent of the time it works every time..



DATA

PEOPLE CAN MAKE UP STATISTICS TO PROVE ANYTHING!

...14% OF ALL PEOPLE KNOW THAT



THE OHIO STATE UNIVERSITY

WEXNER MEDICAL CENTER

...and for any other reason...

**THEY WILL BE ON YOUR
SPECIALTY BOARD
EXAM!!!**



Learning Objectives

- Review basic study design & levels of evidence common to clinical research
- Review basic applications of hypothesis testing:
 - purpose of *p-value*
 - tests for determining difference b/w groups
 - (eg. *t- test & ANOVA, etc.*)
 - tests for determining relationships
 - (eg. *Correlation analysis, regression, etc.*)
- Understand difference between *prevalence & incidence*
- Review & understand results of basic statistical analysis commonly used in clinical diagnostic research:
 - *Sensitivity, Specificity*
 - *Positive Predictive Value, Negative Predictive Value*



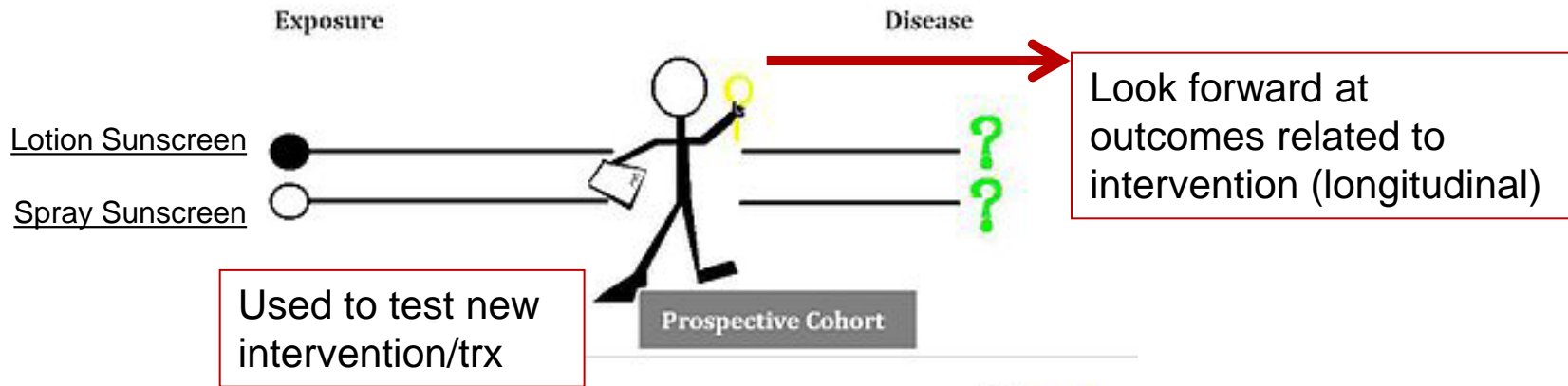
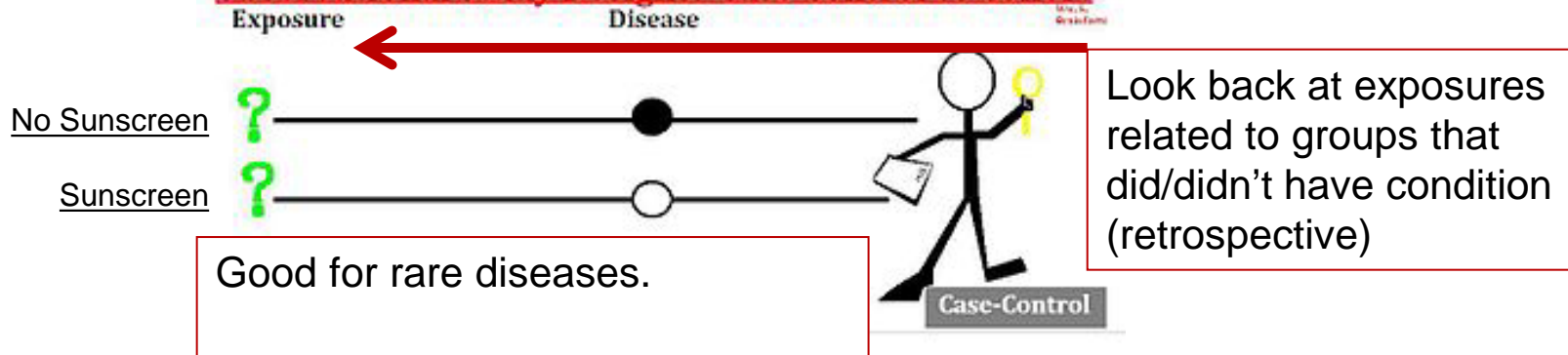
Common Clinical Research Design Types Study





“...how did dey do dat?”

- **Systematic Reviews/Meta-Analysis**
 - Focused review and synthesis of results from RCTs
- **Randomized Controlled Trial:**
 - Subject randomized into different groups
- **Cohort:**
 - Examine 2 or more groups over time
- **Case Control:**
 - Patients with condition are matched to a control group
- **Cross-Sectional:**
 - Data is collected at a single point in time (prevalence)
- **Case Reports/Case Series:**
 - Medical histories in one or more patients with condition or treatment



Observational Study Designs: Case Control vs Cohort



KEY	
	Investigator/Researcher begins their research. When the researcher enters the scene.
	Present
	Absent
	What we are seeking; the information we are trying to obtain; what we do not know; our question.

STATISTICS
THE DISCIPLINE THAT PROVES
THE AVERAGE HUMAN HAS
ONE TESTICLE



What are these?

- P – value
- T-test
- Analysis of Variance (ANOVA)
- Pearsons Correlation (r)
- Regression (r^2)



Error?

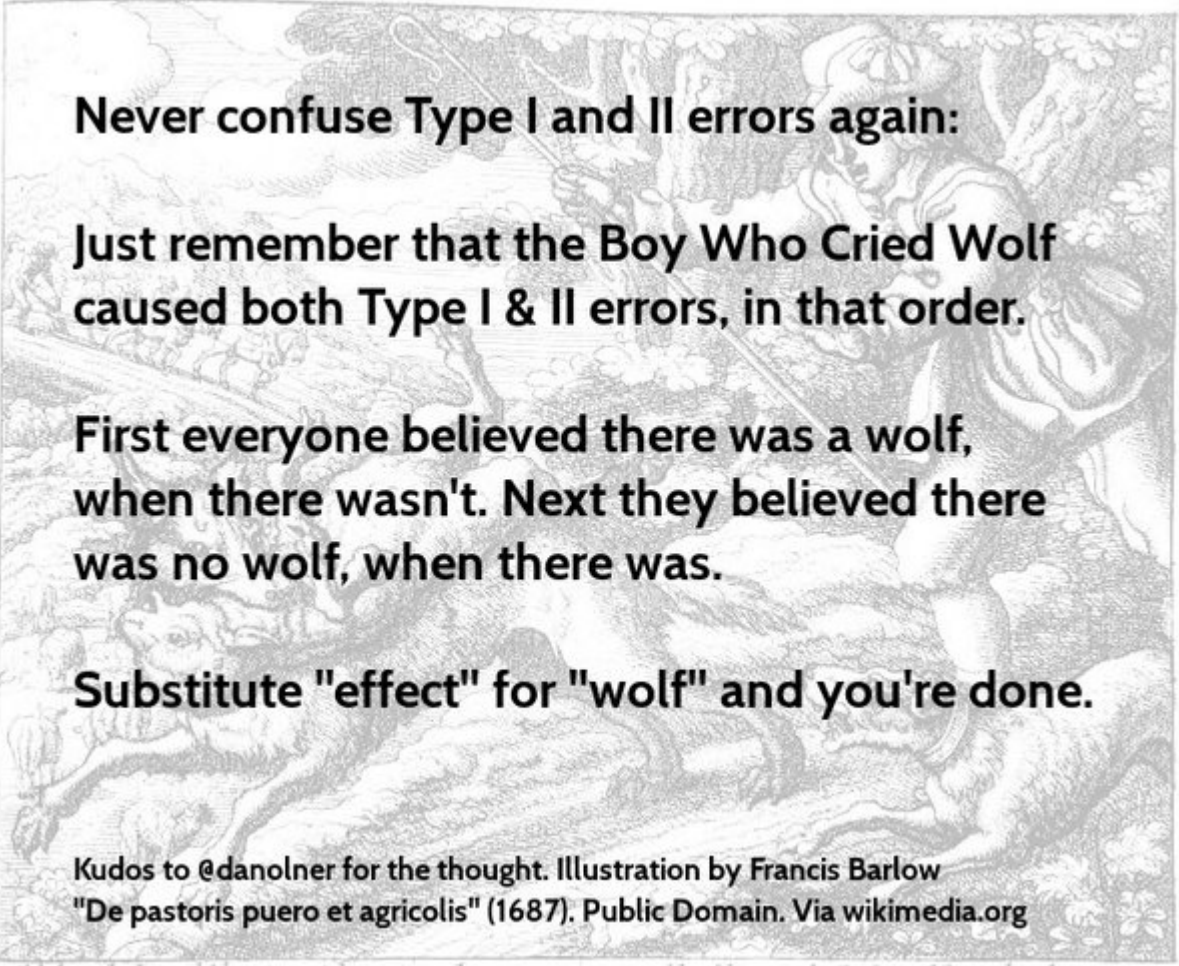
- **Type I:** (false '+')
Concluding there **IS** a difference between groups when there really isn't...
- **Type II:** (false '-')
Concluding there is **NO** difference between groups when there actually is...

α

		Reality	
		True	False
Measured/ Perceived	True	Correct 😊	Type I False Positive
	False	Type II False Negative	Correct 😊

β





Never confuse Type I and II errors again:

Just remember that the Boy Who Cried Wolf caused both Type I & II errors, in that order.

First everyone believed there was a wolf, when there wasn't. Next they believed there was no wolf, when there was.

Substitute "effect" for "wolf" and you're done.

Kudos to @danolner for the thought. Illustration by Francis Barlow
"De pastoris puero et agricolis" (1687). Public Domain. Via [wikimedia.org](https://commons.wikimedia.org/wiki/File:De_pastoris_puero_et_agricolis.jpg)



Significance...?

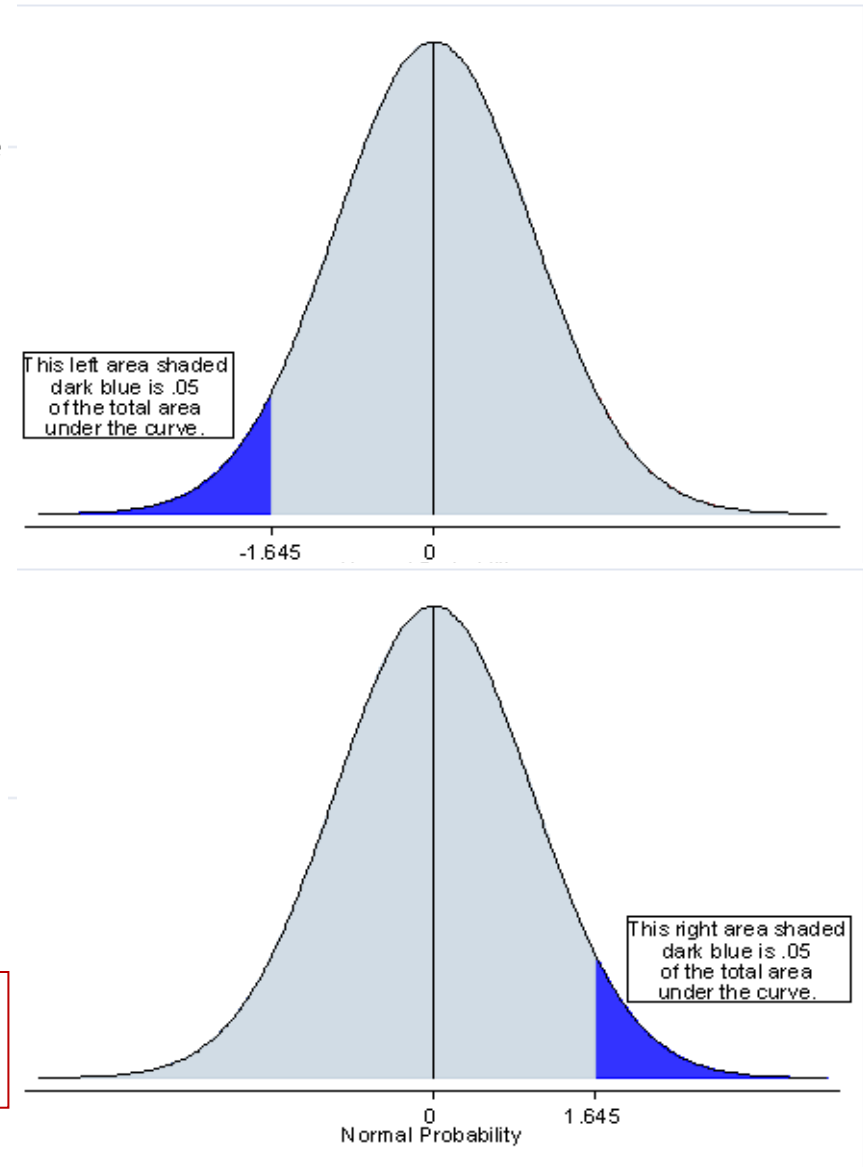
It's all about P for "percentage"

p-value:

- Probability of committing a type I error
- $p=.05$
 - 5% probability that the difference b/w means/groups occurred by chance
 - 5% chance of type I error



Roll a 20-sided die and you'll notice that any given number comes up pretty often!



How do I know if there is a difference?

Parametric

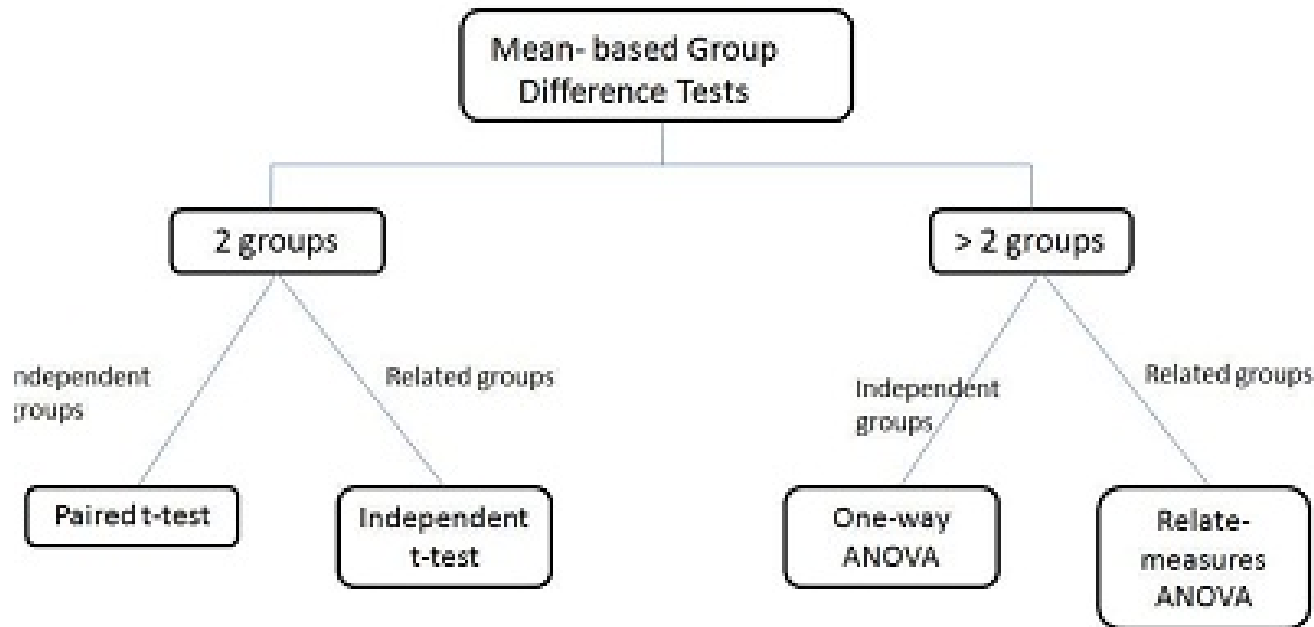
- **T-Test:** means of **2** groups

- **Analysis of Variance (ANOVA):** means of **>2** groups

Non-Parametric

- **Mann-Whitney U-Test:** means of **2** groups

- **Kruskal-Wallis ANOVA by ranks (H or χ^2):** means of **>2** groups



Scenario

- Aim 1: To determine the optimal exercise intervention (volitional quad set or electrical stimulation) in improving quad strength 1 week following ACLR.
 - H_1 :
- Aim 2: To characterize the relationship between quad strength and knee effusion following ACLR.
 - H_2 :



Which statistical tests should be used?

Patients s/p

T-Test

- Dependent Variable: Quad strength
- Intervention:
 - **Group A** → Quad sets
 - **Group B** → Electrical Stimulation
- What test would you use to determine if quad strength was different between the groups A & B following the interventions?



Which statistical tests should be used?

Patients s/ ACLR...

Group A
(Females)

ANOVA

- Dependent Variable: **Quad Strength**
- Intervention:
 - **Group A** → Quad sets
 - **Group B** → Electrical Stimulation
- What test would you use to determine if quad strength was different between groups A, B, males, & females following the interventions?



Which statistical tests should be used?

Patients s/

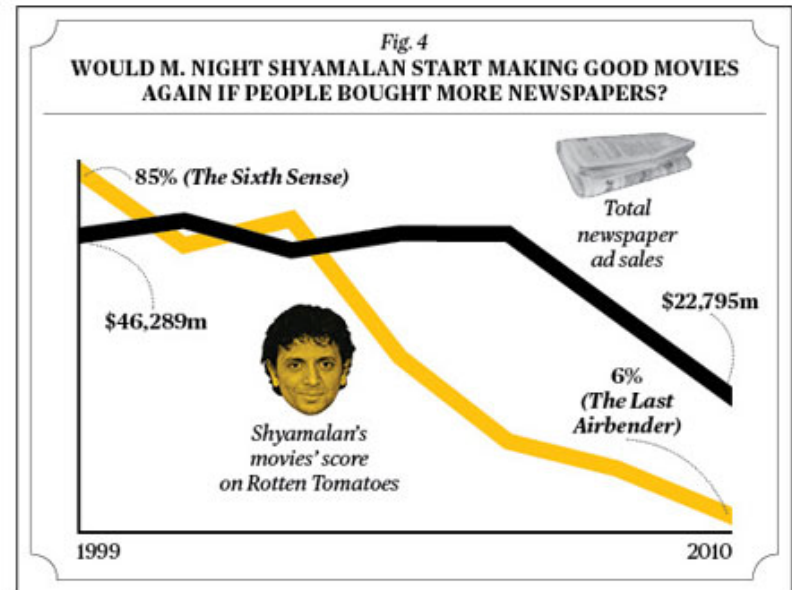
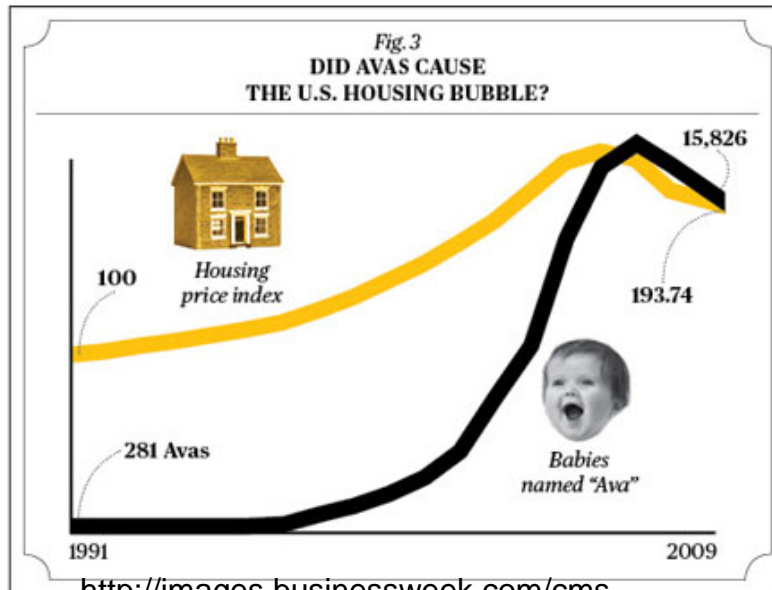
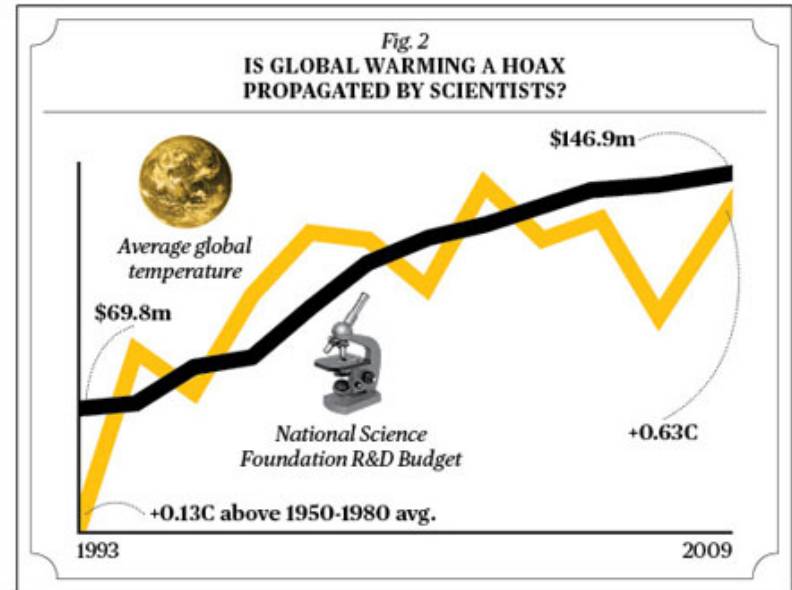
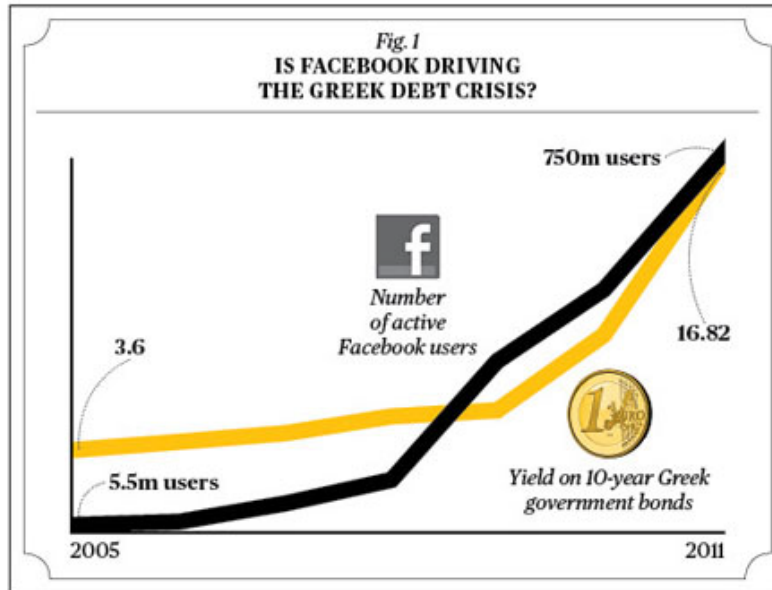
- De
- Intervent
 - **Group A** →
 - **Group B** → Electri

Pearson's
Correlation
(r)

- What test would you use to determine association between quad strength and knee effusion p



Pearson's Correlation (r)



Pearson's Correlation (r)

- Direction & strength of linear relationships
- Not causative

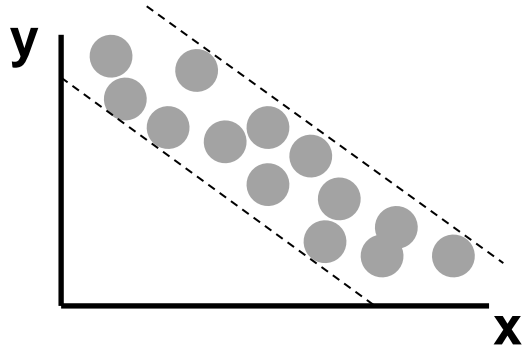


Negative					Positive				
< .70	.40-.69	.30-.39	.20-.29	.01-.19	.01-.19	.20-.29	30-.39	.40-.69	≥ .70
Very Strong	Strong	Moderate	Weak	None	None	Weak	Moderate	Strong	Very Strong

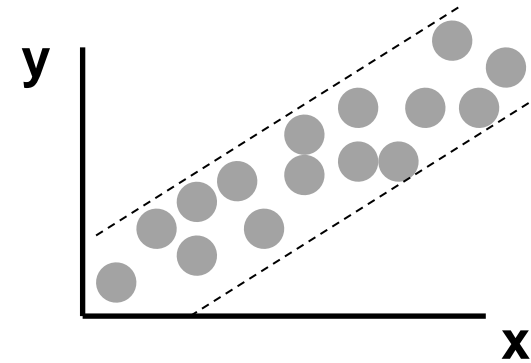


Pearson's Correlation (r)

Strong '-' relationships



Strong '+' relationships



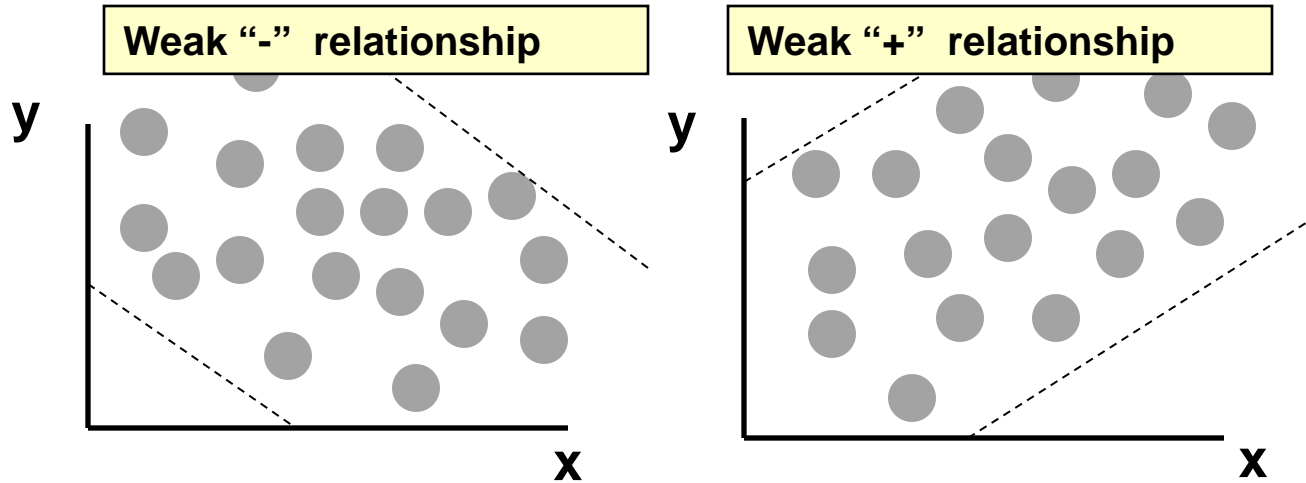
Negative

Positive

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Very Strong	Strong	Moderate	Weak	None	None	Weak	Moderate	Strong	Very Strong



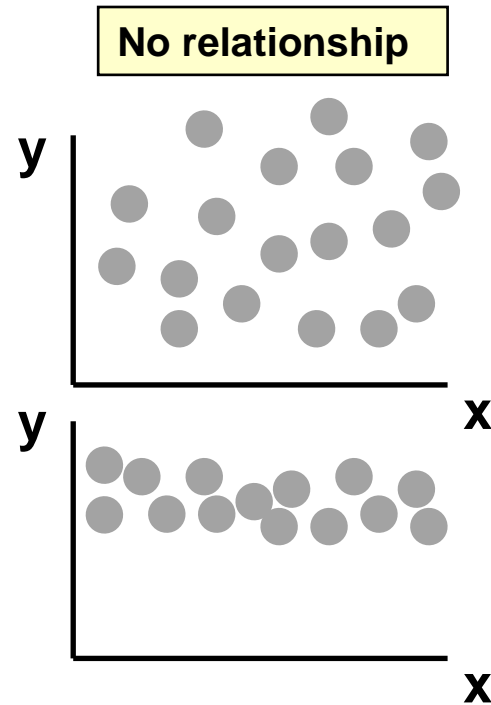
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Pearson's Correlation (r)



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Scenario

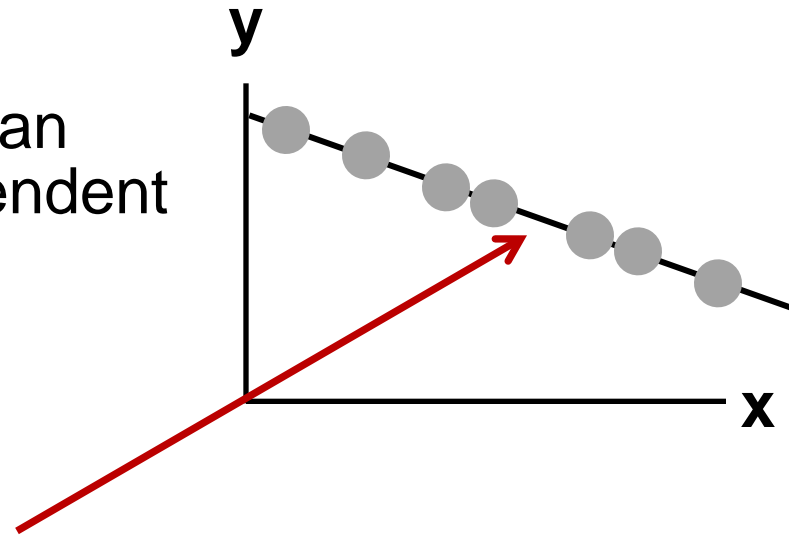
- Aim 3: To predict the contribution of quad strength to IKDC score following ACLR.
 - H_3 :



Linear Regression (r^2)

Predict the value of a dependent variable (*outcome* → *IKDC Score*) based on the value of at least one independent variable (*predictor* → *Quad Strength*)

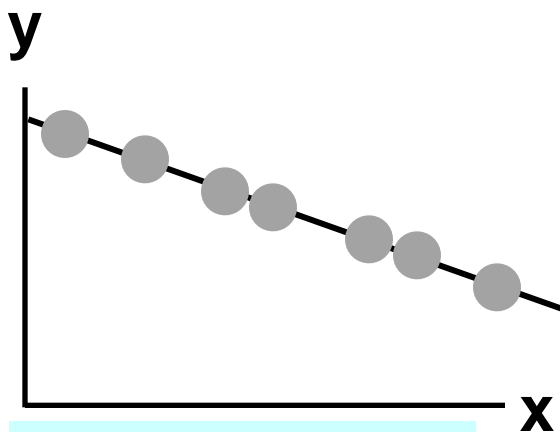
- Explain the impact of changes in an independent variable on the dependent variable



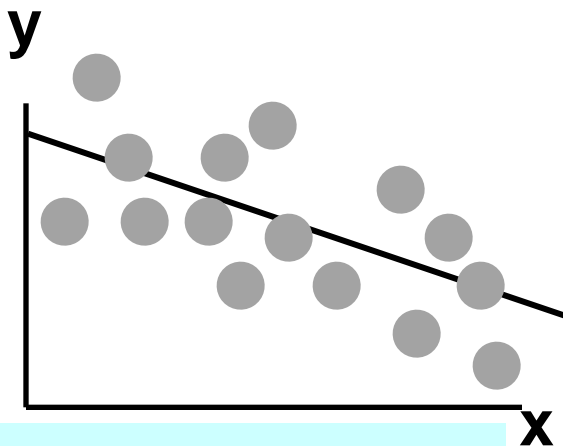
Regression line summarizes relationship between explanatory, x , & response variable, y
predict value of y for a given value of x



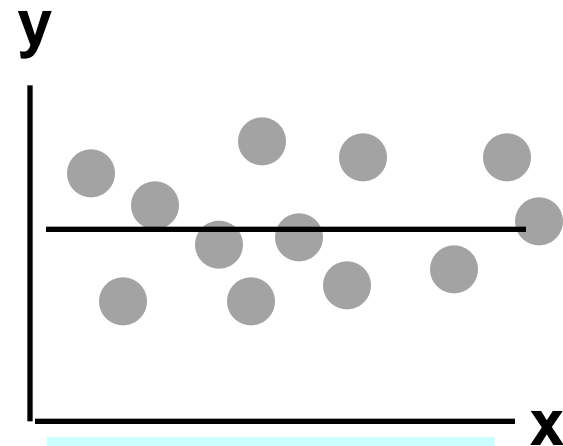
r & r² (How much explanation of variance?)



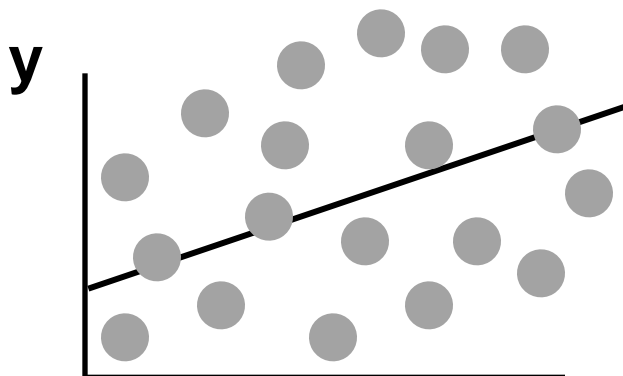
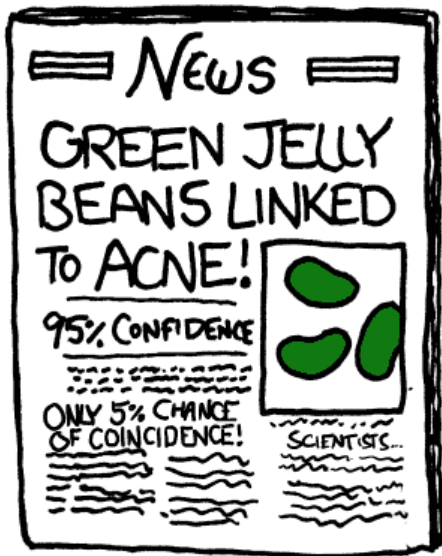
$r = -1; r^2 = 1$



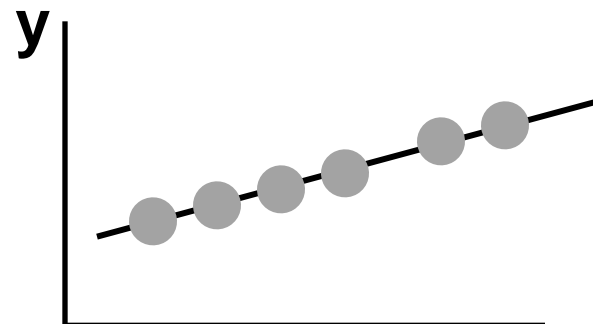
$r = -.6; r^2 = .36$



$r = 0; r^2 = 0$



$r = +.3; r^2 = .09$



$r = +1; r^2 = 1$

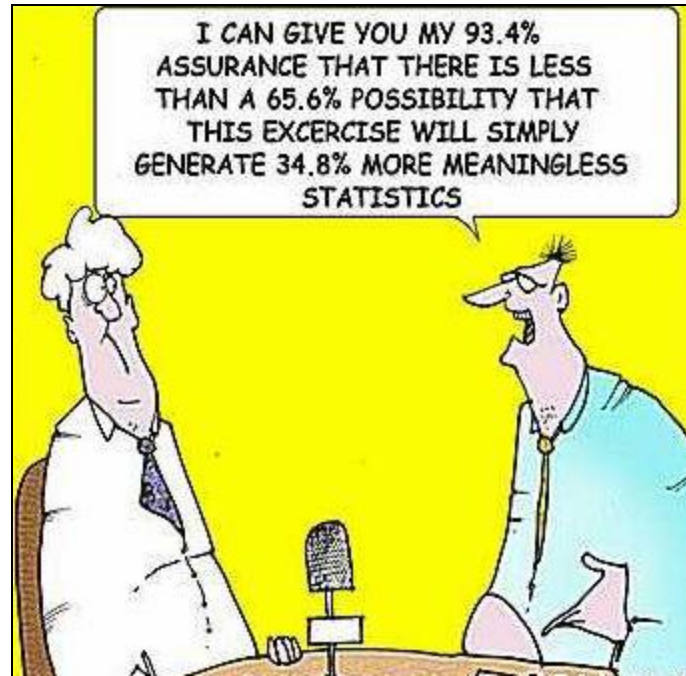


What are these?

- P – value
- T-test
- ANOVA
- Pearsons Correlation
- Regression



Have you had enough yet...?



Diagnostic Testing....oh boy...

	Reference Standard Positive	Reference Standard Negative
Diagnostic Test Positive	True + Results a	False + Results b
Diagnostic Test Negative	False - Results c	True - Results d



Statistic	Formula	Description
Overall Accuracy	$(a+d)/(a+b+c+d)$	Percentage of patients who are correctly diagnosed
Sensitivity	$a/(a+c)$	Proportion of patients with the condition who have a + test result
Specificity	$d/(b+d)$	Proportion of patients without the condition who have a - test result
Positive Predictive Value	$a/(a+b)$	Proportion of patients with a + test result who have the condition
Negative Predictive Value	$d/(c+d)$	Proportion of patients with a - test result who don't have the condition
Positive Likelihood Ratio	$\text{Sensitivity}/(1 - \text{Specificity})$	If the test is +, the increase in odds favoring the condition
Negative Likelihood Ratio	$(1 - \text{Sensitivity})/\text{Specificity}$	If the test is -, the decrease in odds favoring the condition



SnNOuT: High Sensitivity, Negative test, Rule out Condition

SpPIn: High specificity, Positive test, Rule In condition

		Condition		
		Positive	Negative	
Test Outcome	Positive	True Positive	False Positive	PPV TP/(TP+FP)
	Negative	False Negative	True Negative	NPV TN/(FN+TN)
		Sensitivity TP/(TP+FN)	Specificity TN/(FP+TN)	



SnNOuT: High Sensitivity, Negative test, Rule out Condition

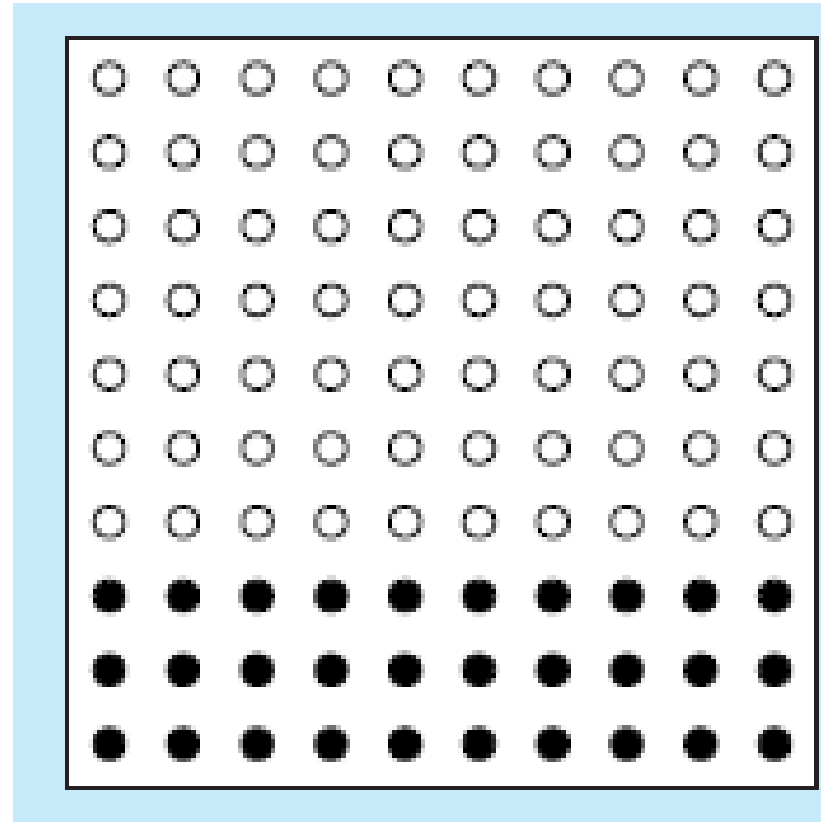
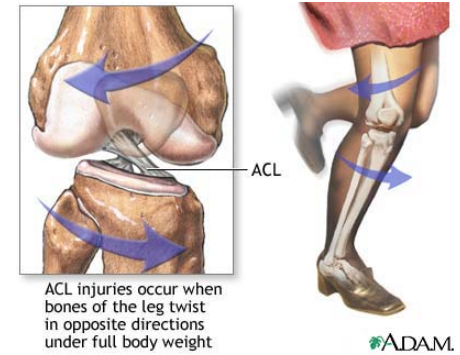
SpPIn: High specificity, Positive test, Rule In condition

		ACL Tear		
		Positive	Negative	
Lachman	Positive	24	14	PPV $24/(24+14)$
	Negative	6	56	NPV $56/(6+56)$
		Sensitivity $24/(24+6)$	Specificity $56/(14+56)$	Total = 100



Example

- Population/Sample: 100
- Torn ACL: 30
- **Prevalence: $30/100 = 30\%$**



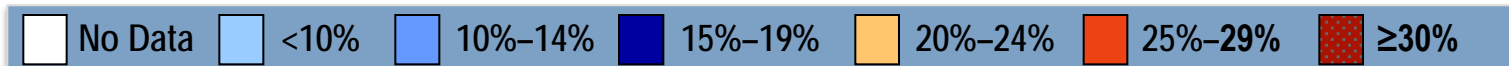
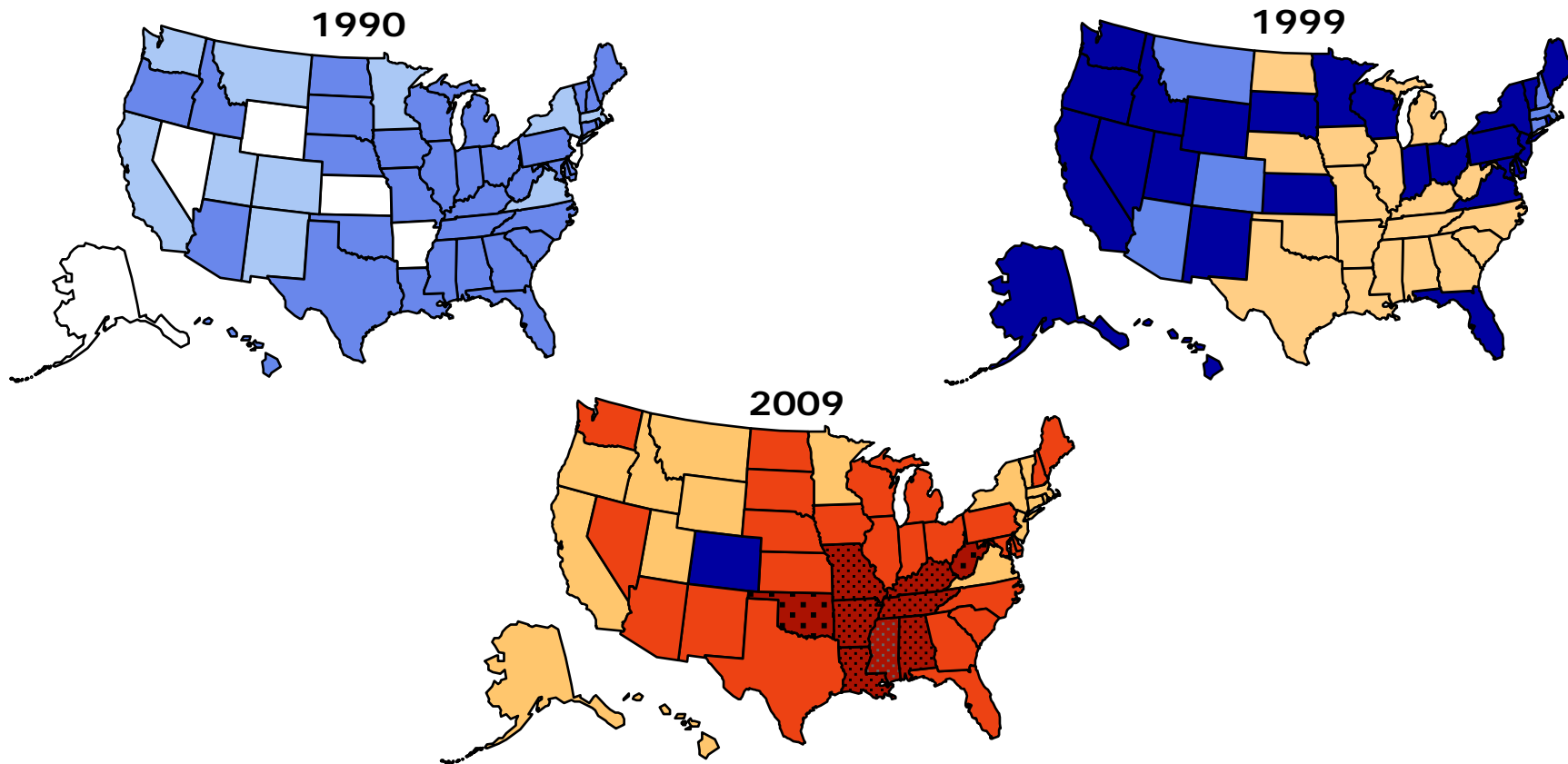
How much is what...?

- **Prevalence:**
 - how much of condition is in population at a **particular point in time**
 - 30 case in a sample of 100
 - $30/100 = 0.30$
 - $0.30 \times 100 = 30\%$
 - **% or # cases per 100,000**



% of Obesity* Among U.S. Adults BRFSS, 1990, 1999, 2009

(*BMI ≥ 30 , or about 30 lbs. overweight for 5'4" person)



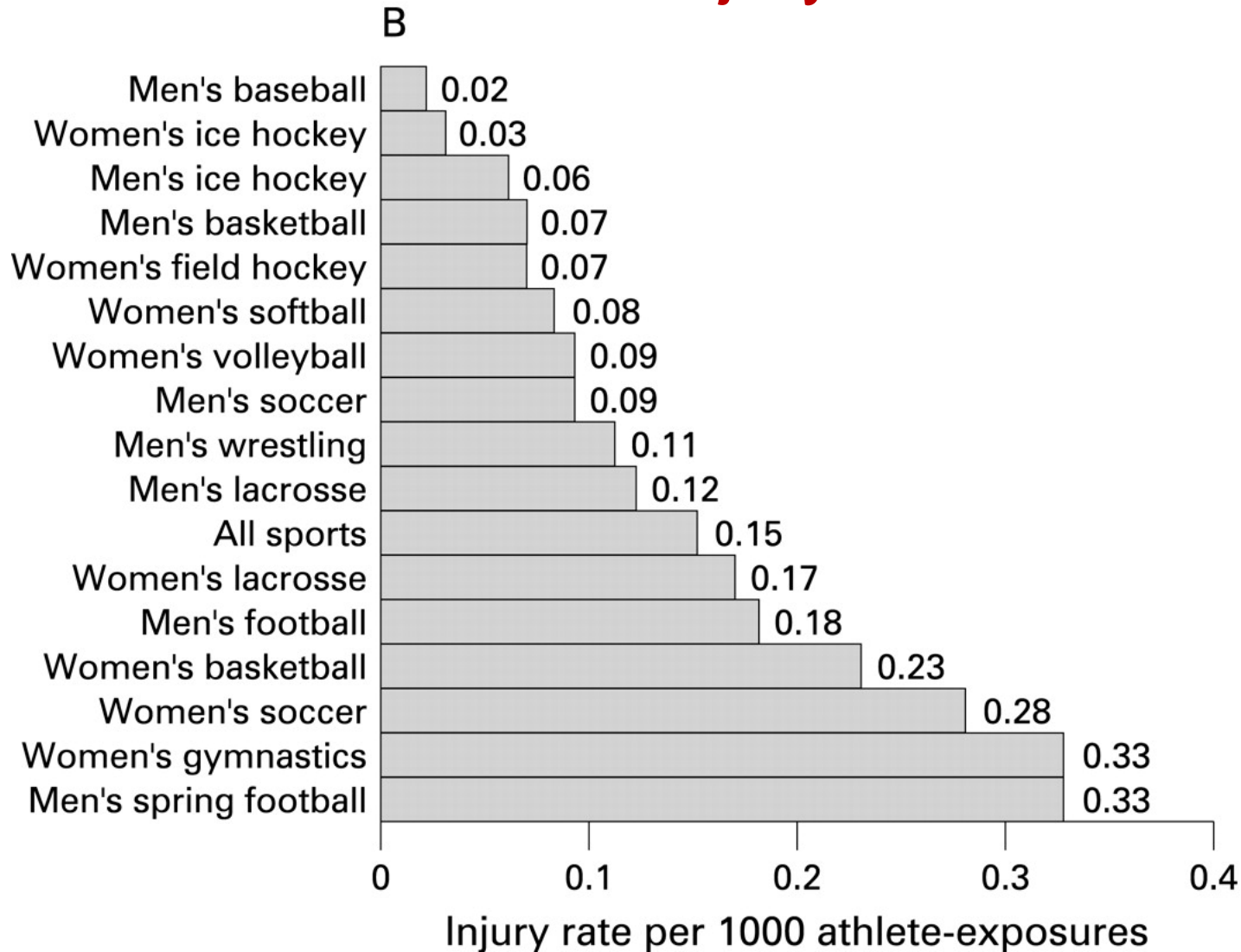
How much is what...?

- **Incidence:**

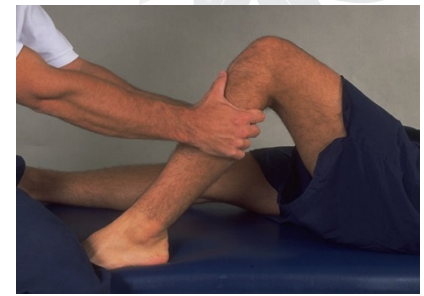
- **Rate** (in month/year/etc.) of occurrence of new cases of a disease or condition
- (# new cases (over time course) / total population)
- # cases per 100,000



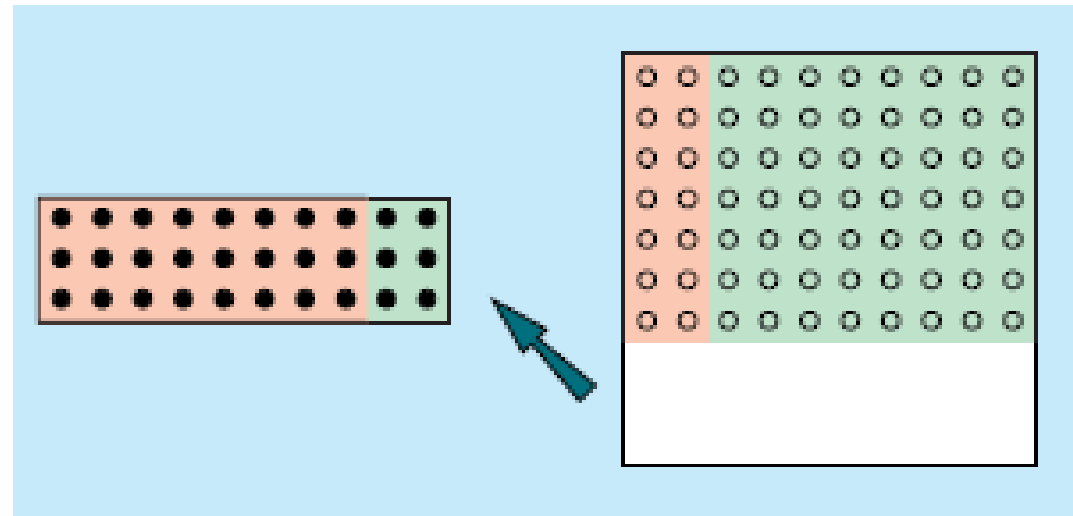
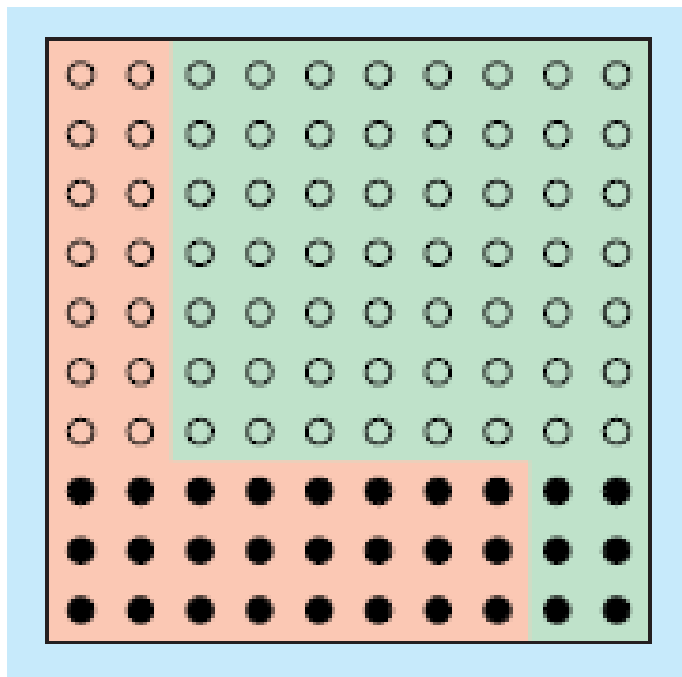
ACL injury



Sensitivity



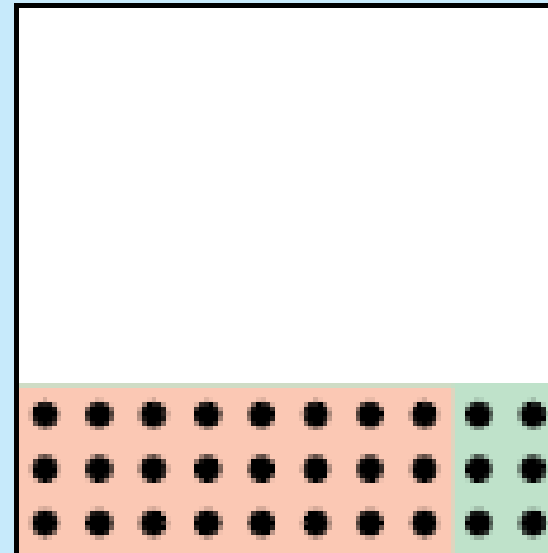
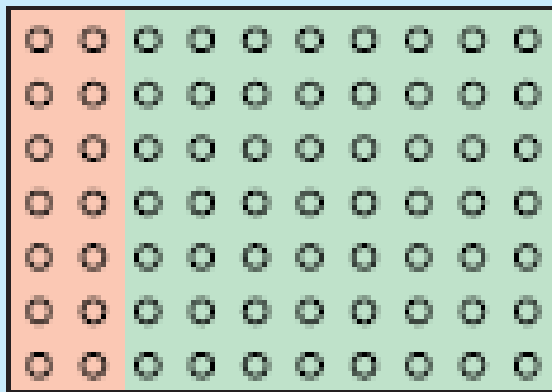
- How good a test is at correctly identifying people who have a “disease/condition”
- “...test's ability to identify positive results.”
 - 24 out of 30 → $[24/(24+6)]=0.80$



Specificity

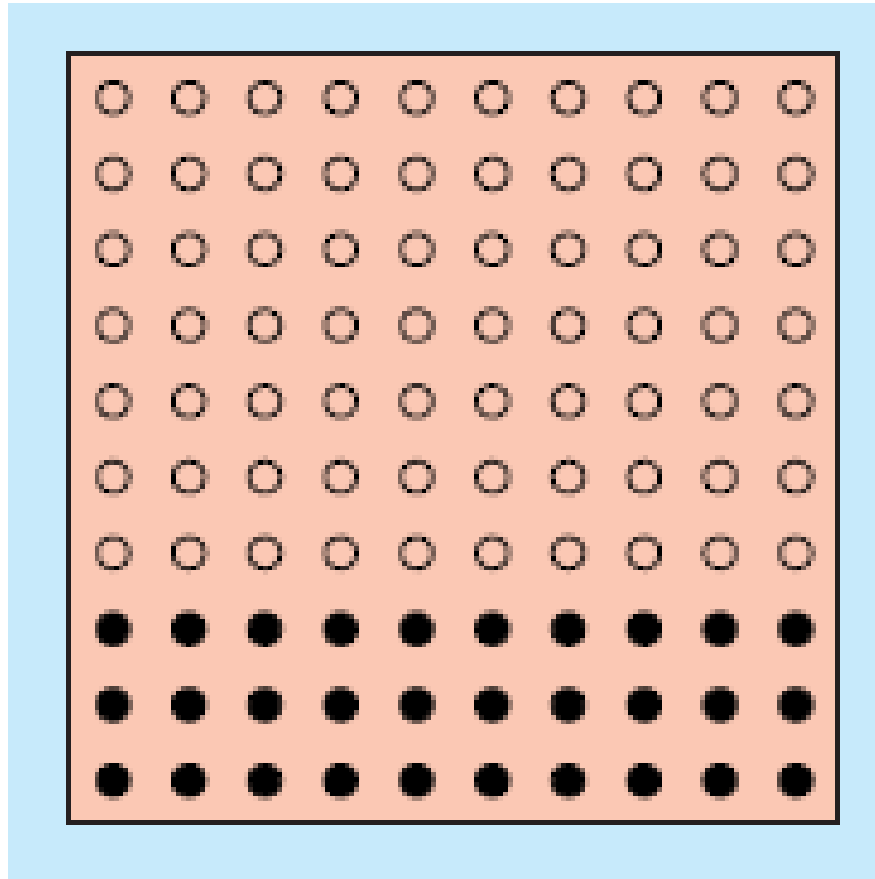


- How good a test is at correctly identifying people who are well
- “...ability of the test to identify negative results.”
 - 56 out of 70 → $[56/(14+56)]=0.80$



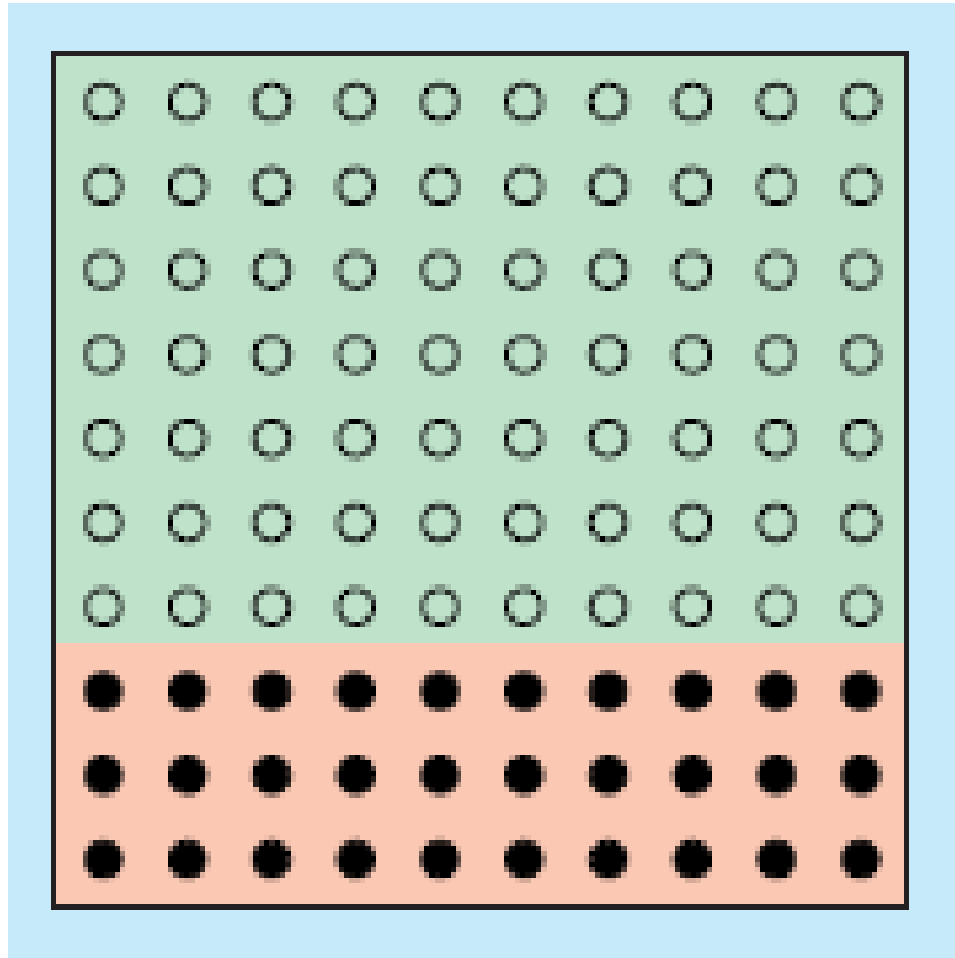
100% Sensitivity

- “...test's ability to identify positive results.”



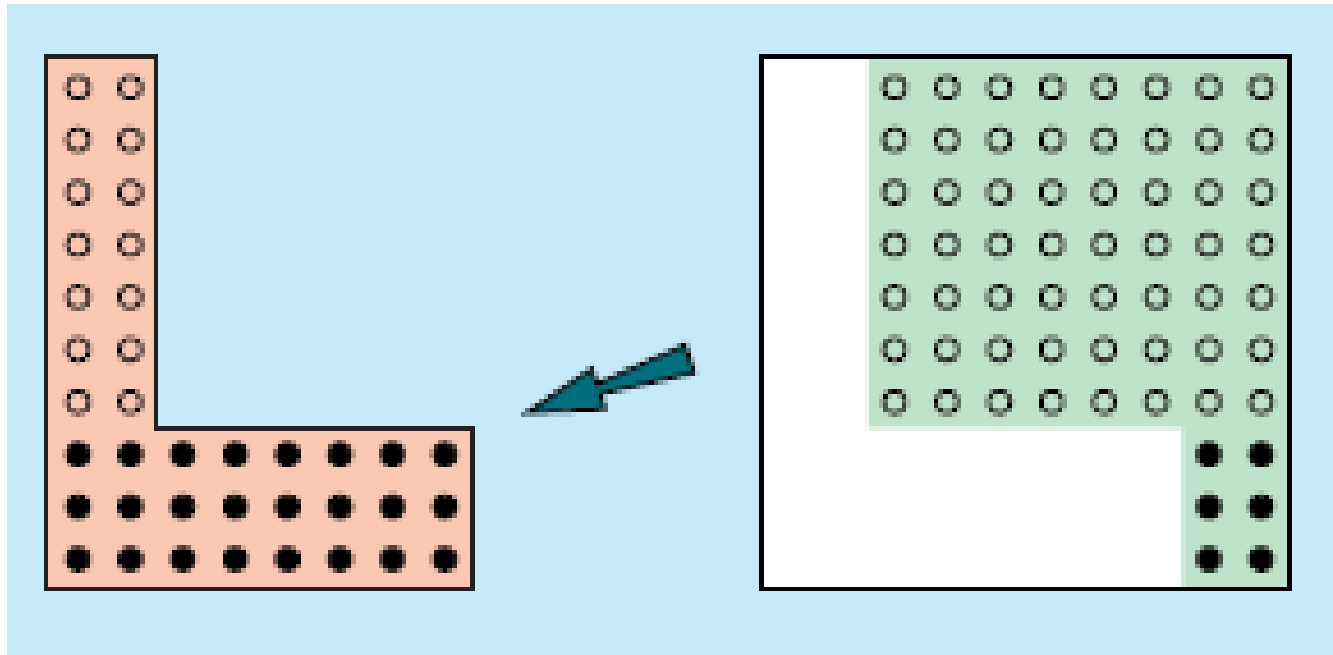
Perfect Test

A+



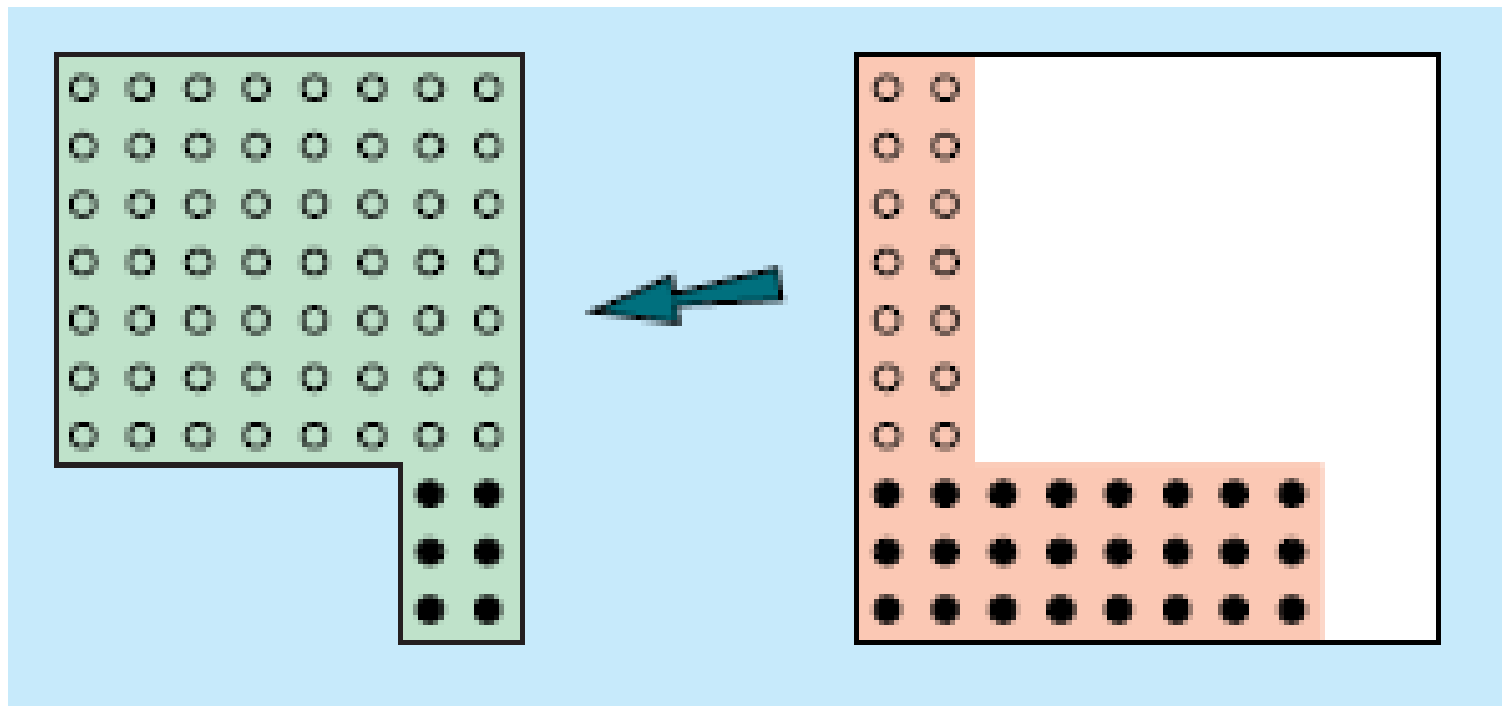
Positive Predictive Value

- The chance that a positive test result will be correct.
- 24 out of 38 positive tests correct: $[24/(24+14)]= 0.63$



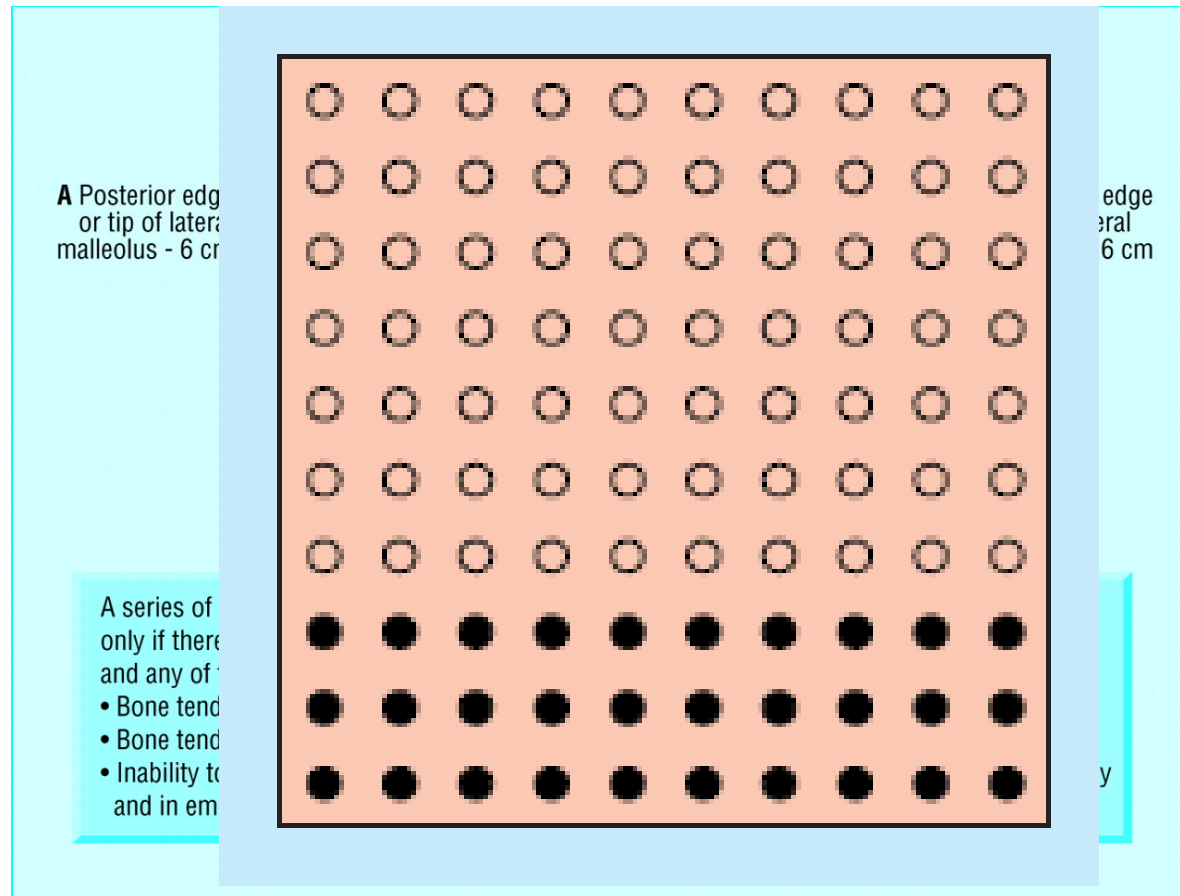
Negative Predictive Value

- The chance that a negative test result will be correct
- 56 out of 62 neg. results correct: $[56/(6+56)] = 0.90$



Ottawa Ankle Rules example...

- Sensitivity ~100%
- Specificity: 48%
- PPV: 15%
- NPV: ~100%



What is the Likelihood Ratio (LR)

The probability of a clinical finding in patients with a condition divided by the probability of the same finding in patients without the condition

Direct estimate of how much a test result will change the odds of having a disease/condition.

Likelihood of a disorder or condition being present



LR “+” / “-”

Increased diagnostic confidence:

“Probability estimate of presence/absence of the condition of interest”

- **LR+** tells you how much the odds of the condition increase when a test is positive.
- **LR-** tells you how much the odds of the condition decrease when a test is negative.

LR -				LR+		
0 - .1	.1 - .2	.2 - .5	.5 - 2	2 - 5	5 - 10	>10
Important		Unimportant			Important	



Likelihood Ratios

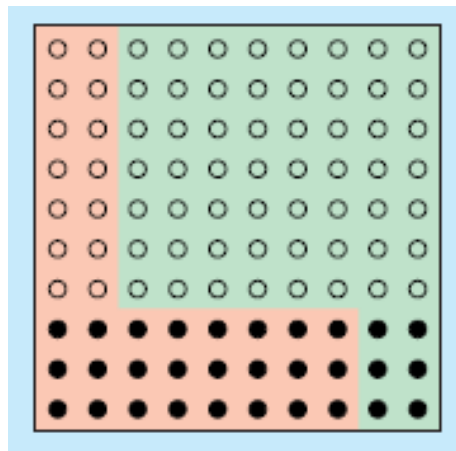
		Condition		
		+	-	
Test	+	True Positive	False Positive	PPV
	-	False Negative	True Negative	NPV
		Sensitivity	Specificity	Total

$$LR+ = \frac{\Pr(T+|D+)}{\Pr(T+|D-)} = \frac{\text{True Positive}}{\text{False Positive}} = \frac{\text{sensitivity}}{1-\text{specificity}}$$

$$LR- = \frac{\Pr(T-|D+)}{\Pr(T-|D-)} = \frac{\text{False Negative}}{\text{True Negative}} = \frac{1-\text{sensitivity}}{\text{specificity}}$$



		ACL Tear		
		Positive	Negative	
Lachman	Positive	24	14	PPV 24/(24+14)
	Negative	6	56	NPV 56/(6+56)
		Sensitivity 24/(24+6)	Specificity 56/(14+56)	



		Actual ACL Tear		
		Positive	Negative	
Anterior Drawer	Positive	24	14	PPV 24/(24+14)
	Negative	6	56	NPV 56/(6+56)
Total		30	70	
		Sensitivity 24/(24+6) = 0.80	Specificity 56/(14+56)	

What is the proportion of patients with an ACL tear who have a “+” Lachman?

$$[24/(24+6)] = 0.80 \text{ (sensitivity)}$$

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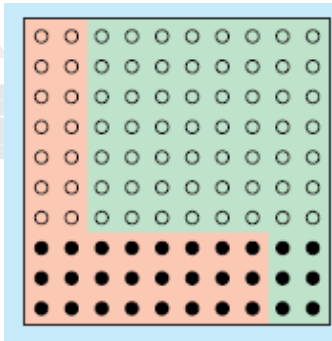
-LR = (1-sensitivity)/specificity = .20/.80 = .25

In other words, a “+” Lachman is **4x’s** more likely in a patient who has an ACL tear than a patient who does not have an ACL tear

AND

A “-” Lachman is only ¼ (**0.25**) more likely in those who have an ACL tear.





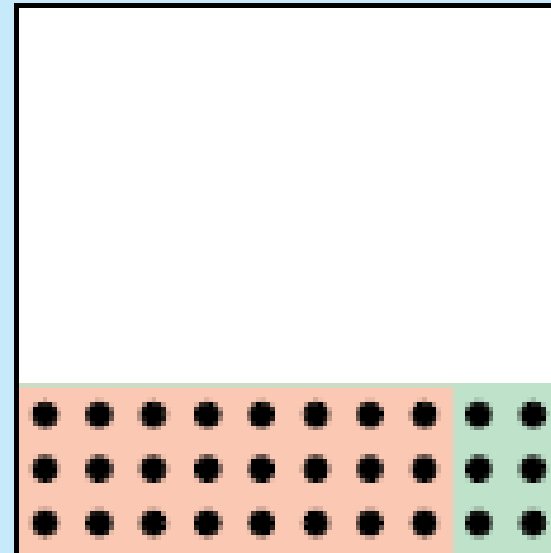
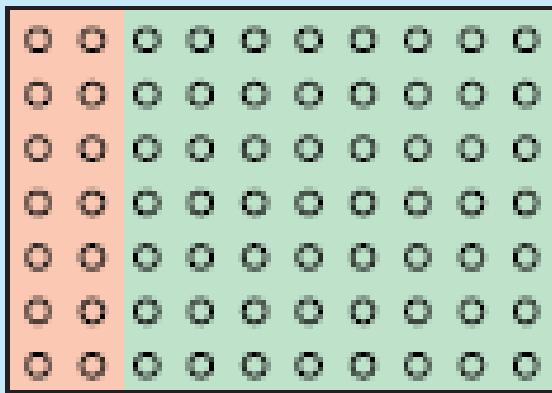
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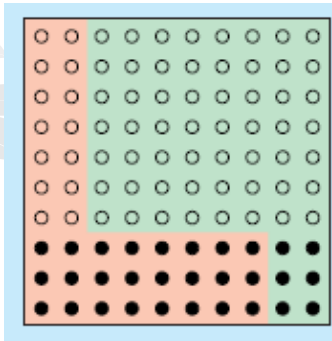
What is the proportion of patients without an ACL tear who have a “-” Lachman?

$$[56/(14+56)] = 0.80 \text{ (specificity)}$$



In other words, ...

... have an ACL tear



		Actual ACL Tear		
		Positive	Negative	
Anterior Drawer	Positive	24	14	PPV 24/(24+14)
	Negative	6	56	NPV 56/(6+56)
Total		30	70	
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What is the proportion of patients without an ACL tear who have a “-” Lachman?

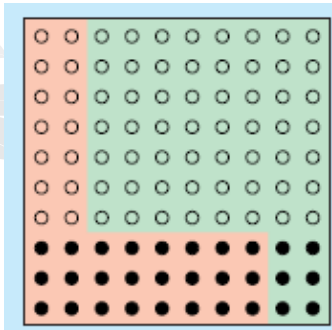
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What is the proportion of patients with a “+” Lachman have an ACL tear?

What is the proportion of patients with a “+” Lachman have an ACL tear?

In other words, a “+” Lachman have an ACL tear

STATE UNIVERSITY
CENTER



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		Positive	Negative	
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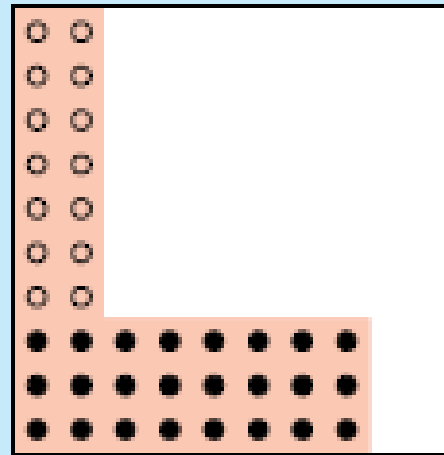
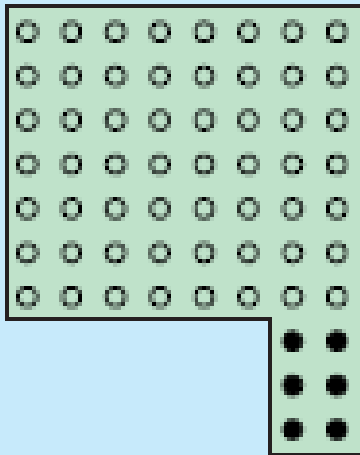
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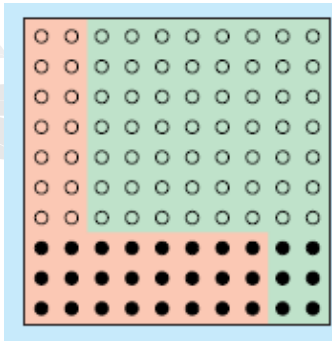
$$[24/(24+14)] = 0.63\% \text{ (PPV)}$$

What is the proportion of patients with a “-” Lachman who don’t have an ACL tear?



In other words, a “+” La

does not have an ACL tear



		Actual ACL Tear		
		Positive	Negative	
Anterior Drawer	Positive	24	14	PPV 24/(24+14)
	Negative	6	56	NPV 56/(6+56)
Total		30	70	
		Sensitivity 24/(24+6) = 0.80	Specificity 56/(14+56)	

What is the proportion of patients with an ACL tear who have a “+” Lachman?

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What is the proportion of patients without an ACL tear who have a “-” Lachman?

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If the Lachman’s is “+”, what are the odds favoring an ACL tear?

$$+LR = \text{sensitivity}/(1-\text{specificity}) = .0.80/0.20 = 4$$

If the Lachman’s “-” what are the odds favoring an ACL tear?

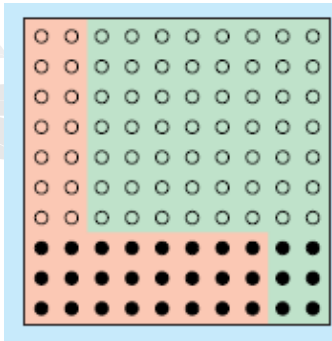
$$-LR = (1-\text{sensitivity})/\text{specificity} = .20/.80 = .25$$

In other words, a “+” Lachman is **4x’s** more likely in a patient who has an ACL tear than a patient who does not have an ACL tear

AND

A “-” Lachman is only ¼ (**0.25**) more likely in those who have an ACL tear.





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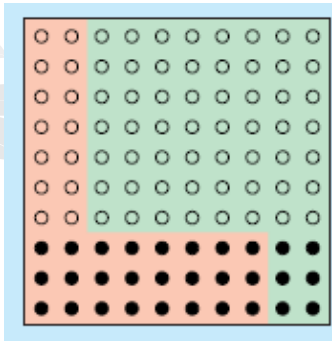
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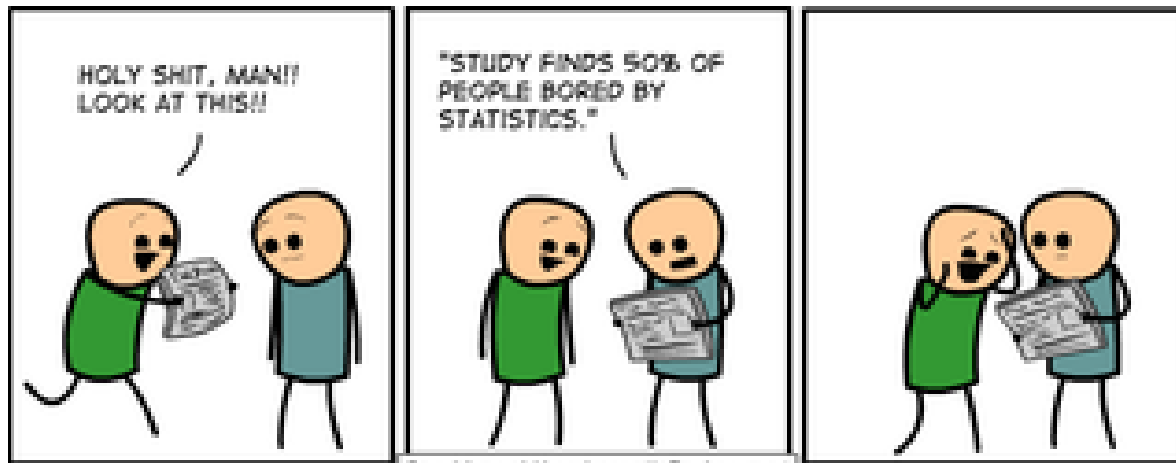
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