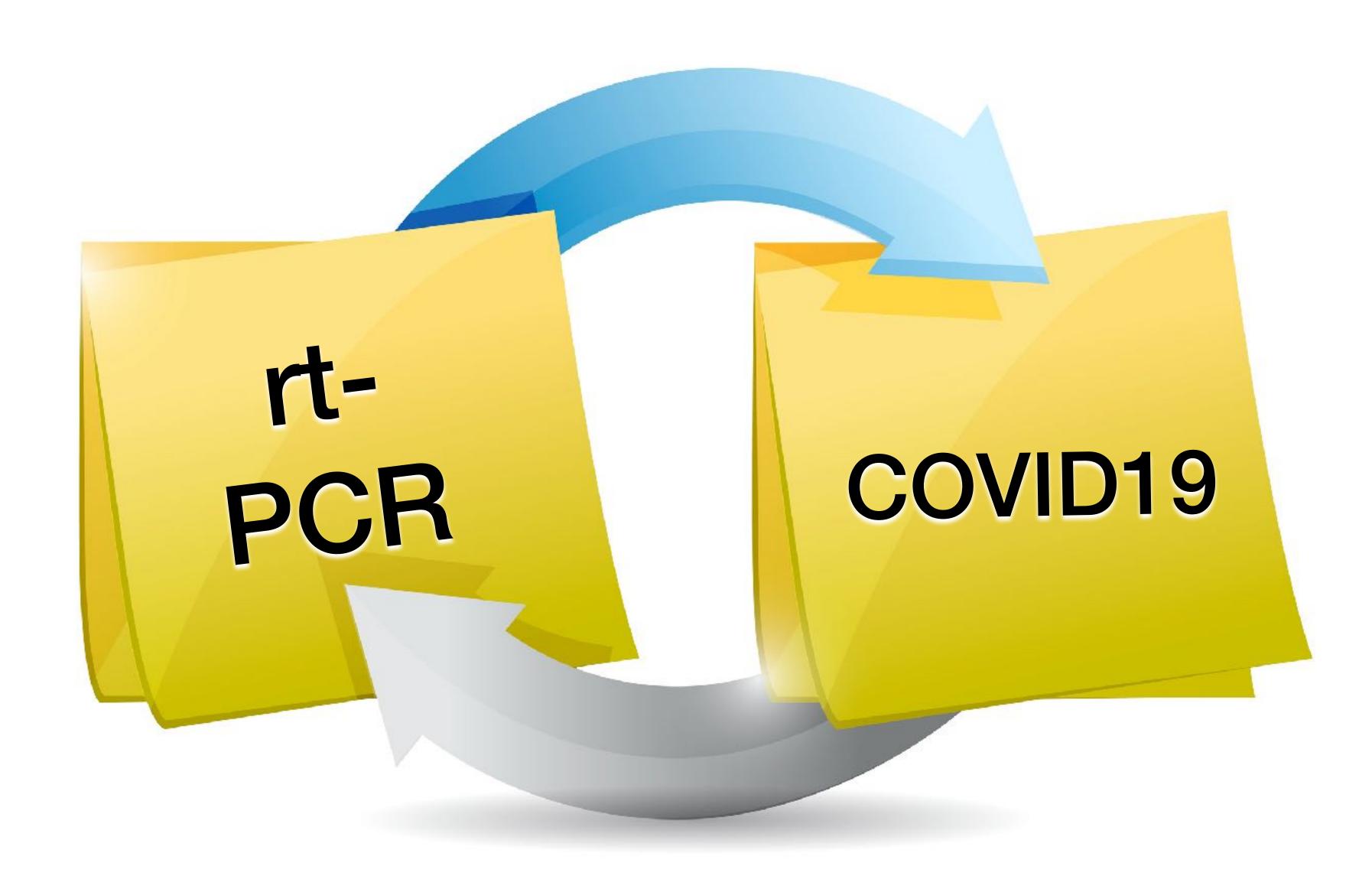




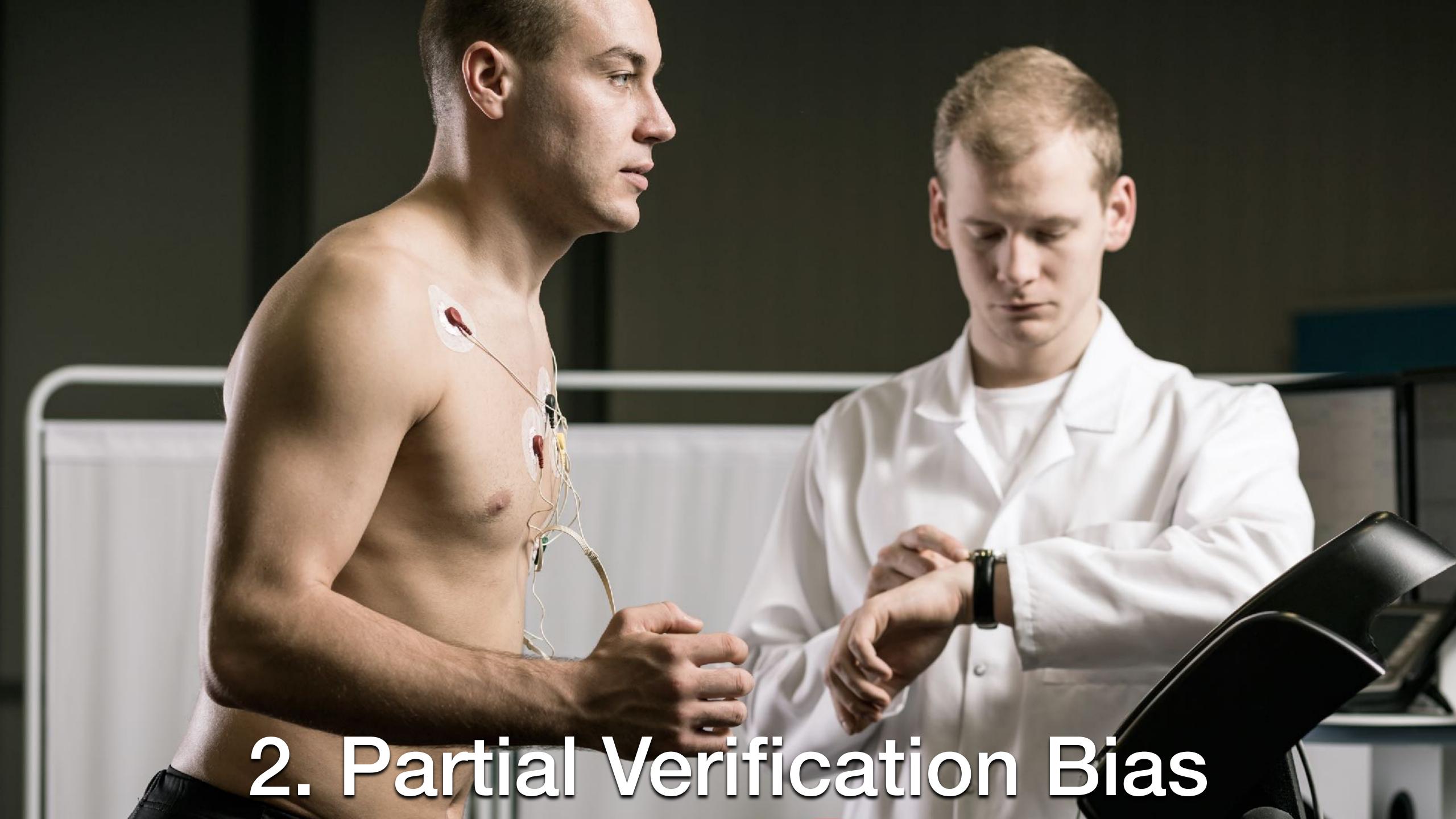
Type of Bias in Diagnostic Studies



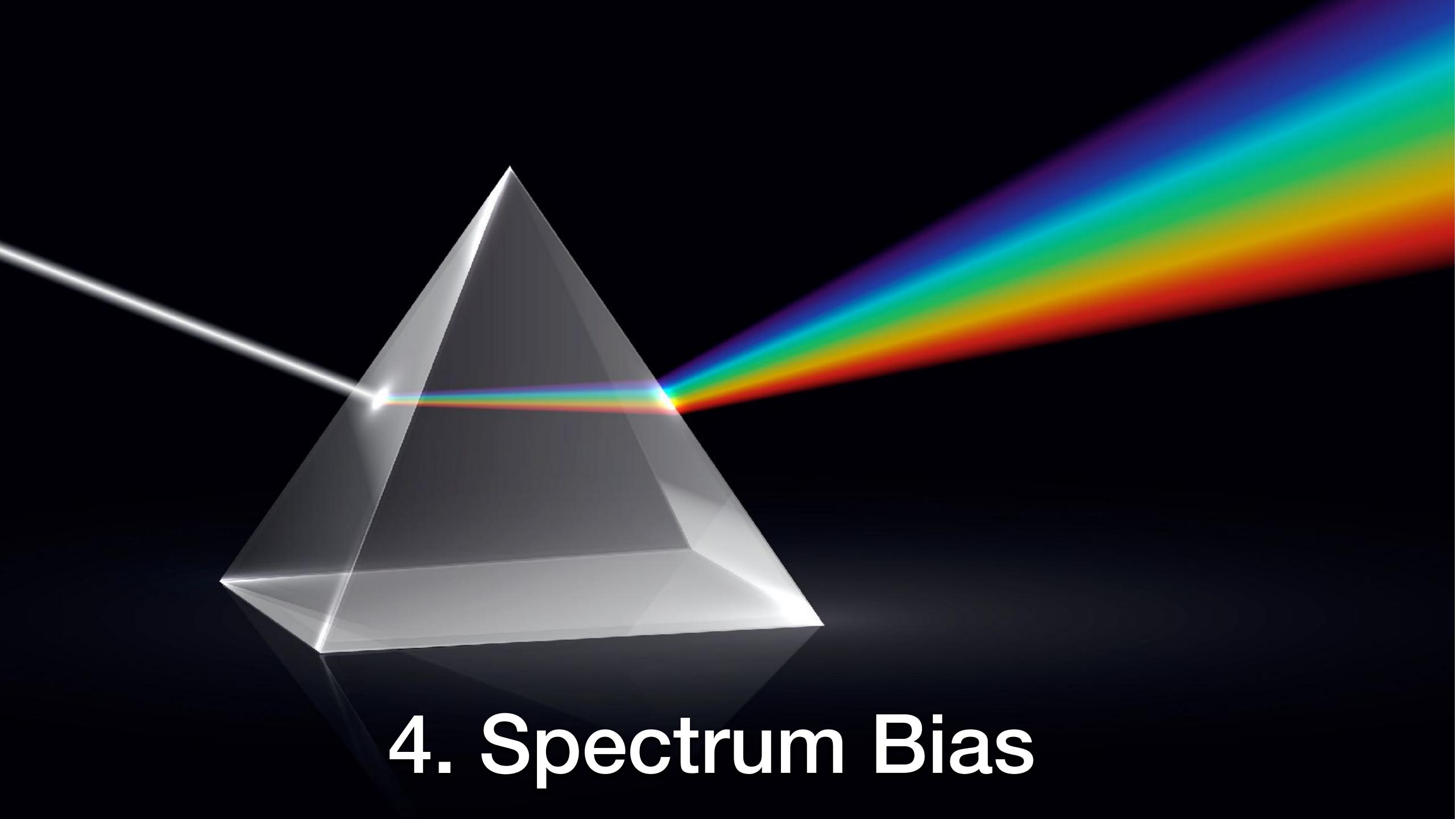
- 1. Incorporation
- 2. Partial Verification
- 3. Differential Verification
- 4. Spectrum
- 5. Imperfect Gold Standard



1. Incorporation Bias







5. Imperfect Gold Standard Bias



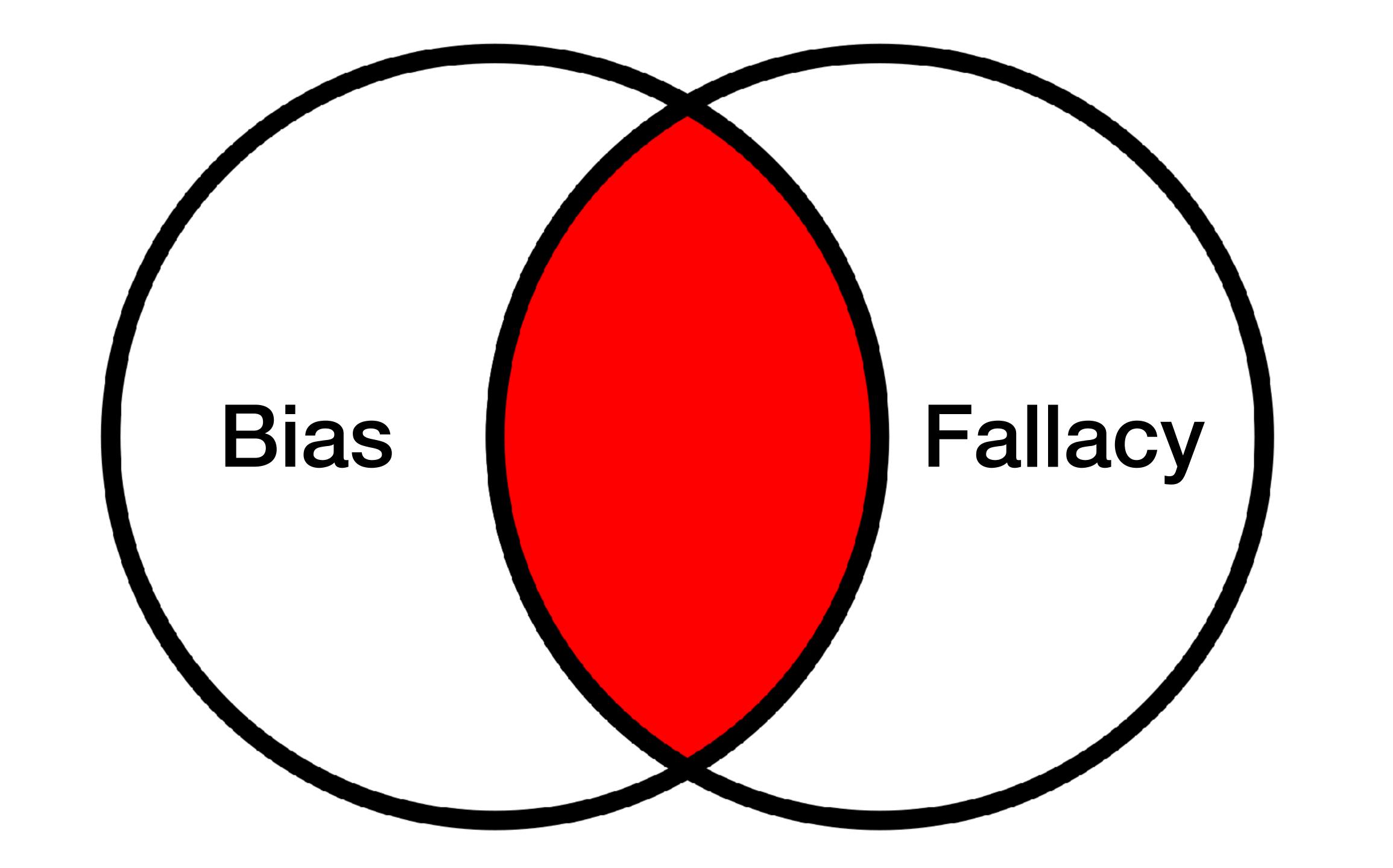
Research Methods & Statistics

Understanding the Direction of Bias in Studies of Diagnostic Test Accuracy

Michael A. Kohn, MD, MPP, Christopher R. Carpenter, MD, MSc, and Thomas B. Newman, MD, MPH









Misrepresenting someone's argument to make it easier to attack.

After Will said that we should put more money into health and education. Warren responded by saying that he was surprised that Will hates our country so much that he wants to leave it defenceless by cutting military spending.



slippery slope

Asserting that if we allow A to happen, then Z will consequently happen too, therefore A should not happen.

Colin Closet asserts that if we allow same-sex couples to marry, then the next thing we know we'll be allowing people to marry their parents, their cars and even monkeys.



pleading

Moving the goalposts to create exceptions when a claim is shown to be false.

Edward Johns claimed to be psychic, but when his 'abilities' were tested under proper scientific conditions, they magically disappeared. Edward explained this saying that one had to have faith in his abilities for them to work.



false cause

Presuming that a real or perceived relationship between things means that one is the cause of the other.

Pointing to a fancy chart, Roger shows how temperatures have been rising over the past few centuries, whilst at the same time the numbers of pirates have been decreasing; thus pirates cool the world and global warming is a heax.



ad hominem

Attacking your opponent's character or personal traits in an attempt to undermine their argument.

After Sally presents an eloquent and compelling case for a more equitable taxation system, Sam asks the audience whether we should believe anything from a woman who isn't married, was once arrested, and smells a bit weird.



loaded question

Asking a guestion that has an assumption built into it so that it can't be answered without appearing guilty.

Grace and Helen were both romantically interested in Brad. One day, with Brad sitting within earshot, Grace asked in an inquisitive tone whether Helen was having any problems with a fungal infection.





Manipulating an emotional response in place of a valid or compelling argument.

Luke didn't want to eat his sheep's brains with chopped liver and brussels sprouts, but his father told him to think about the poor, starving children in a third world. country who weren't fortunate enough to have any food at all.



Presuming that because a claim has been poorly argued, or a fallacy has been made, that it is necessarily wrong.

Recognising that Amanda had committed a fallacy in arguing that we should eat healthy food because a nutritionist said it was popular, Alyse said we should therefore eat bacon double cheeseburgers every day.



tu quoque

Avoiding having to engage with criticism by turning it back on the accuser answering criticism with criticism.

The blue candidate accused the redicandidate of committing the tu quoque fallacy. The red candidate responded by accusing the blue candidate of the same, after which ensued an hour of back and forth criticism with not much progress.



burden of proof

Saying that the burden of proof lies not with the person making the claim, but with someone else to disprove.

Bertrand declares that a teapot is, at this very moment, in orbit around the Sun between the Earth and Mars, and that because no one can prove him wrong his claim is therefore a valid one.



personal incredulity

Saying that because one finds something difficult to understand that it's therefore not true.

Kirk drew a picture of a fish and a human and with effusive disdain asked Richard if he really thought we were stupid enough to believe that a fish somehow turned into a human through just, like, random things happening over time.



ambiguity

Using double meanings or ambiguities of language to mislead or misrepresent the

When the judge asked the defendant why he hadn't paid his parking fines, he said that he shouldn't have to pay them because the sign said 'Fine for parking here' and so he naturally presumed that it would be fine to park there.



the gambler's

Believing that 'runs' occur to statistically independent phenomena such as roulette wheel spins.

Red had come up six times in a row on the roulette wheel, so Greg knew that it was close to certain that black would be next up. Suffering an economic form of natural selection with this thinking, he soon lost all of his savings.



black-or-white

Where two alternative states are presented as the only possibilities, when in fact more possibilities exist.

Whilst rallying support for his plan to fundamentally undermine citizens' rights, the Supreme Leader told



bandwagon

Appealing to popularity or the fact that many people do something as an attempted form of validation.

Sharnus pointed a drunken finger at Sean and asked him to explain how so many people could believe in leprechauns if they're only a stily old superstition. Sean, however, had had a few too many Guinness: himself and fell off his chair.



begging the question

A circular argument in which the conclusion is included in the premise.

The word of Zorbo the Great is flawless and perfect. We know this because it says so in The Great and Infallible Book of Zorbo's Best and Most Thuest Things that are Definitely True and Should Not Ever Be Questioned



appeal to authority

Using the opinion or position of an authority figure, or institution of authority, in place of an actual argument.

Not able to defend his position that evolution 'isn't true' Bob says that he knows a scientist who also questions evolution (and presumably isn't a primate).



composition /division

Assuming that what's true about one part of something has to be applied to all, or other, parts of it.

Daniel was a precocious child and had a liking for logic. He reasoned that atoms are invisible, and that he was made of atoms and therefore invisible too. Unfortunately, despite his thinky skills, he lost the game of hide and go seek.



no true scotsman

Making what could be called an appeal to purity as a way to dismiss relevant criticisms or flaws of an argument.

Angus declares that Scotsmen do not put sugar on their porridge, to which Lachlan points out that he is a Scotsman and puts sugar on his porridge. Furious, like a true Scot, Angus yells that no true Scotsman sugars his porridge.



genetic

Judging something good or bad on the basis of where it comes from, or from whom it comes.

Accused on the 6 o'clock news of corruption and taking bribes, the senator said that we should all be very wary of the things we hear in the media, because we all know how very unreliable the media can be.



appeal to nature

Making the argument that because something is 'natural' it is therefore valid, justified, inevitable, good, or ideal.

The medicine man rolled into town on his bandwagon offering various natural remedies, such as very special. plain water. He said that it was only natural that people should be wary of 'artificial' medicines such.



anecdotal

Using personal experience or an isolated example instead of a valid argument, especially to dismiss statistics.

Jason said that that was all cool and everything, but his grandfather smoked, like, 30 cigarettes a day and lived. until 97 - so don't believe everything you read about meta analyses of sound studies showing proven causal relationships.



the texas sharpshooter

Cherry-picking data clusters to suit an argument, or finding a pattern to fit a presumption.

The makers of Sugarette Candy Drinks point to research showing that of the five countries where Sugarette drinks sell the most units, three of them are in the top ten healthiest countries on Earth, therefore Sugarette drinks are healthy.



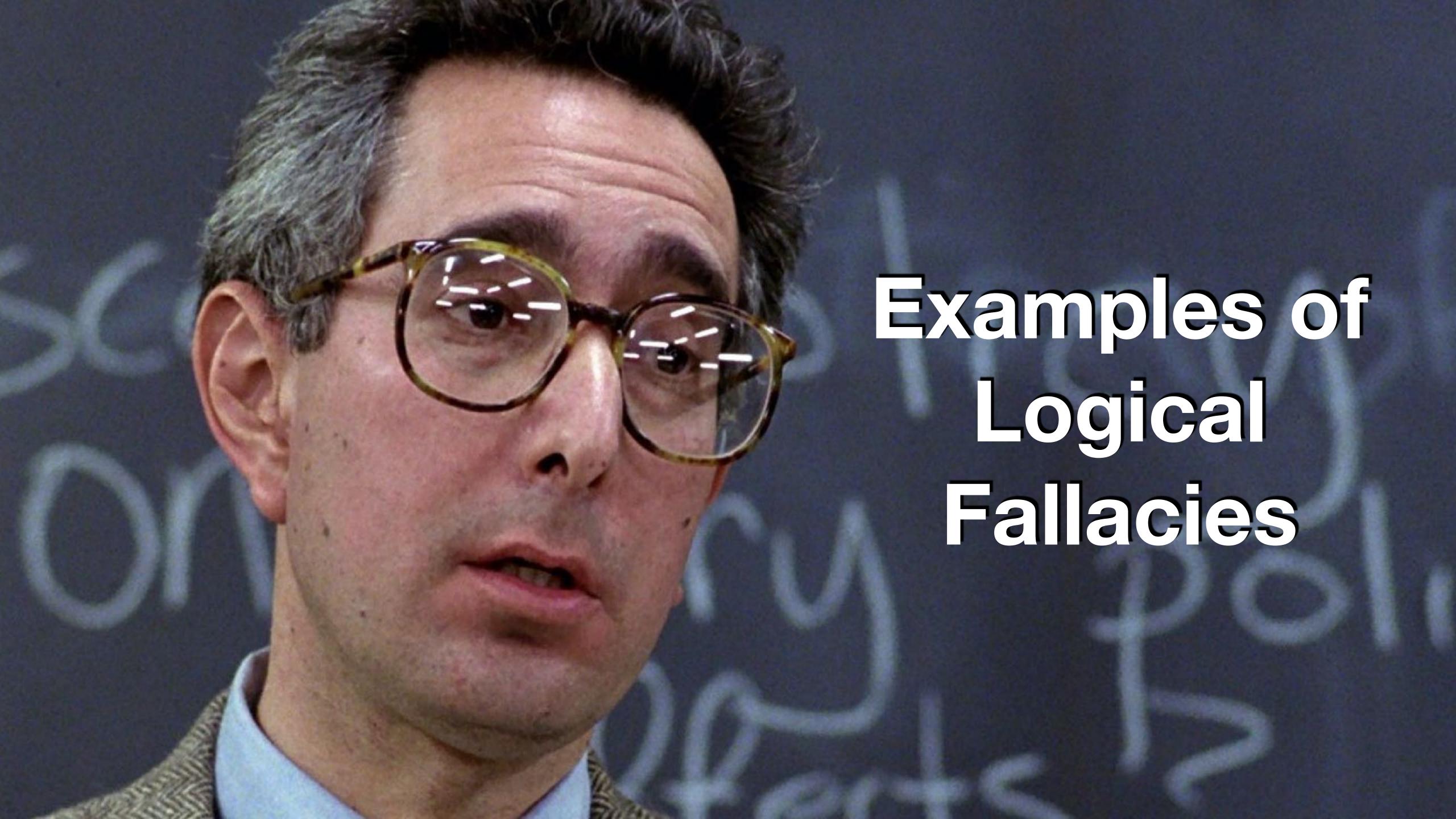
middle ground

Saying that a compromise, or middle point, between two extremes is the truth.

Holly said that vaccinations caused autism in children, but her scientifically well-read friend Caleb said that this claim had been debunked and proven false. Their friend

thou shalt not commit logical fallacies











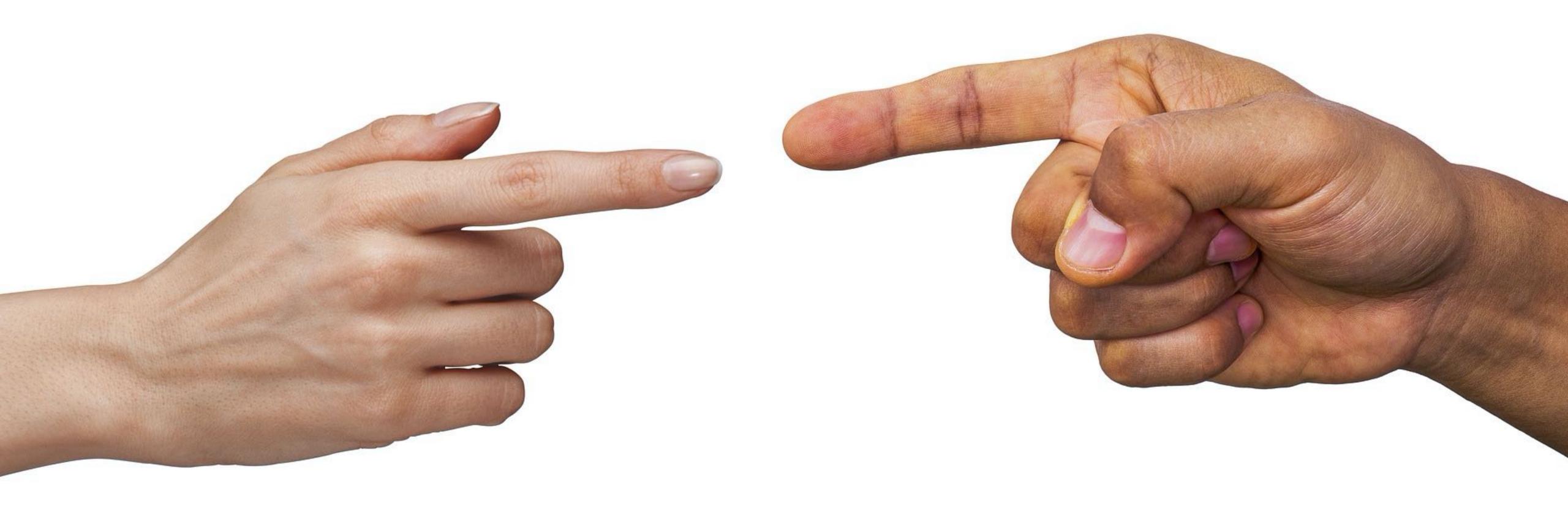












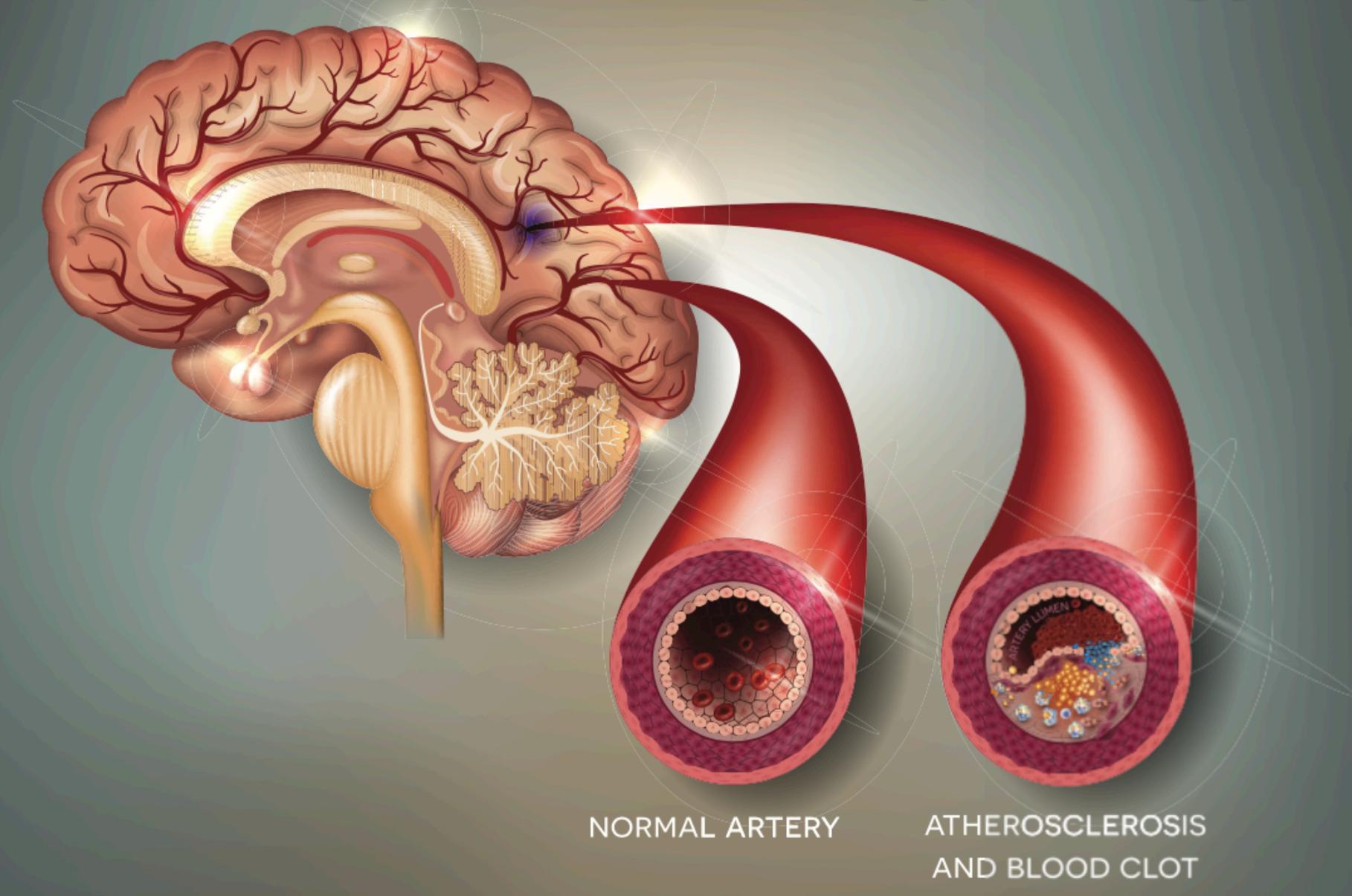
Tu Quoque







Appeal to Pathophysiology





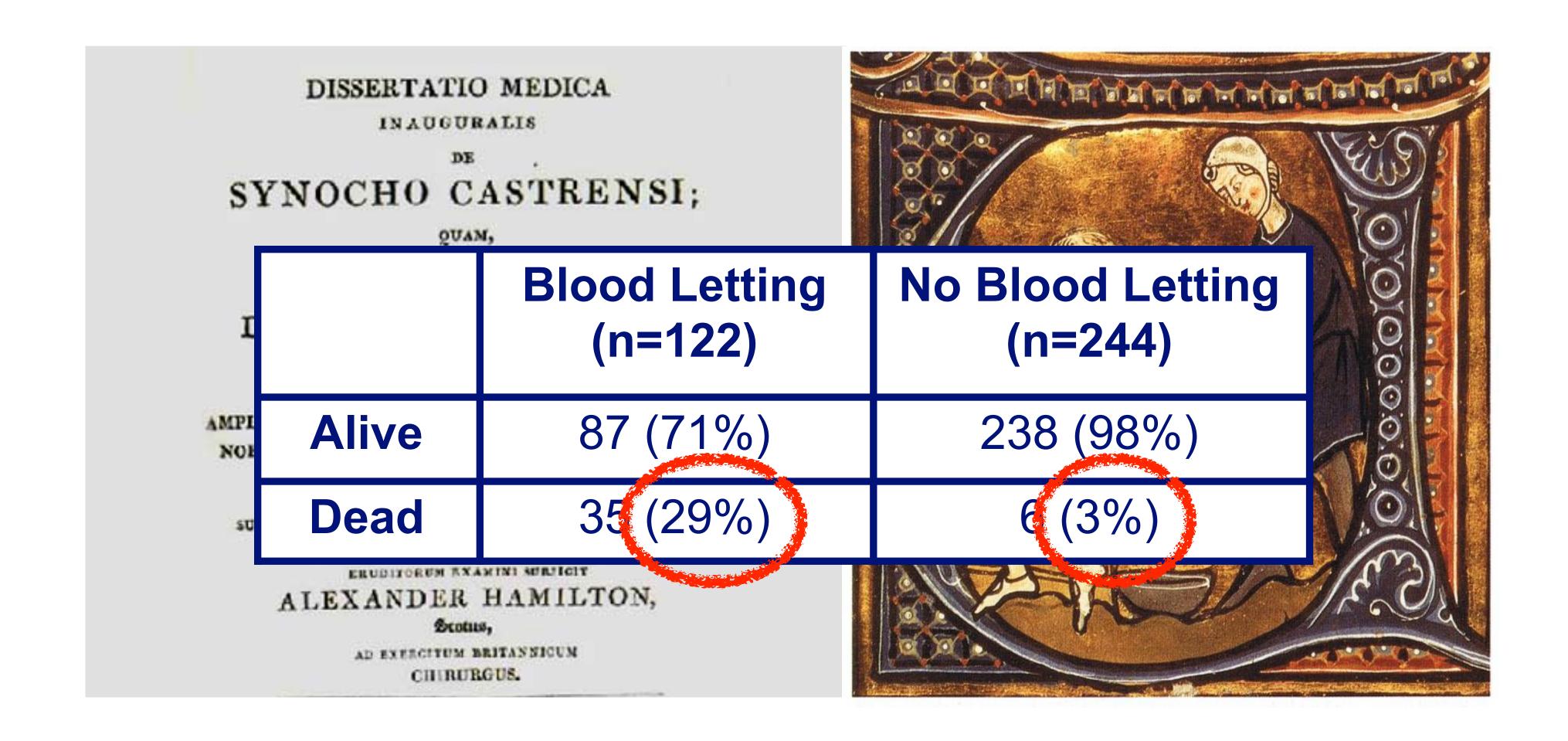




Right Patients



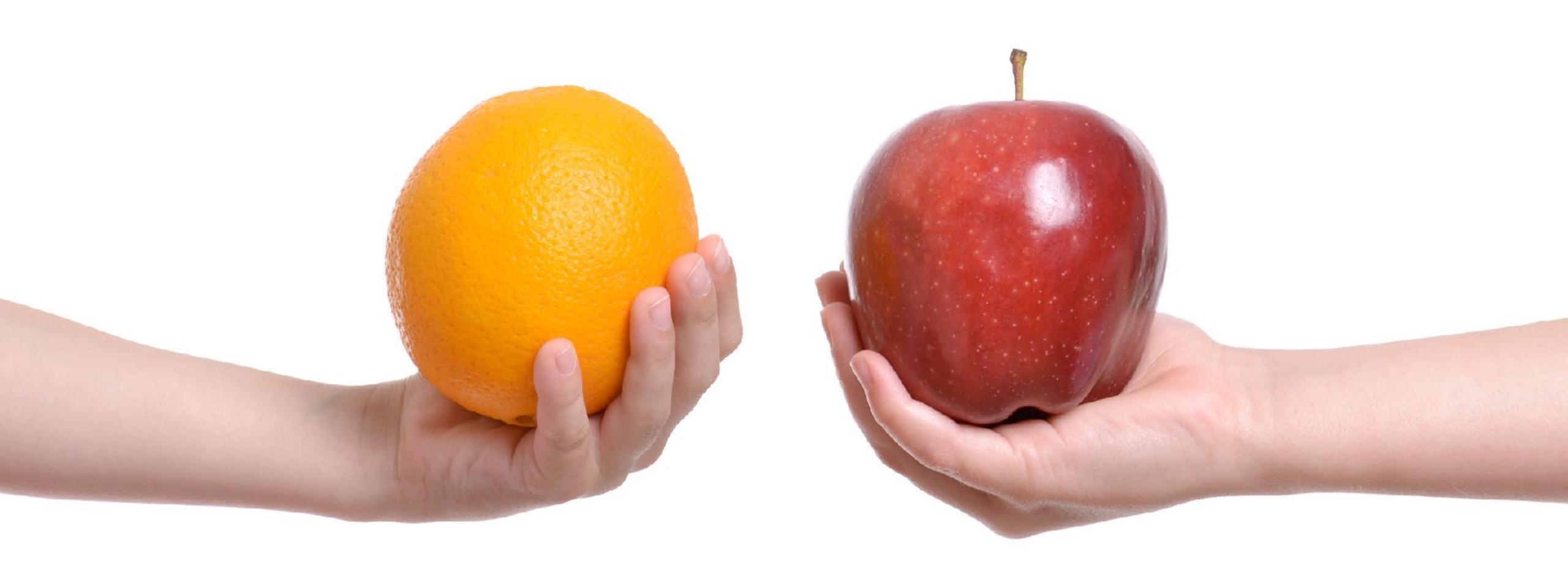
Randomization



NNH 4



Similar



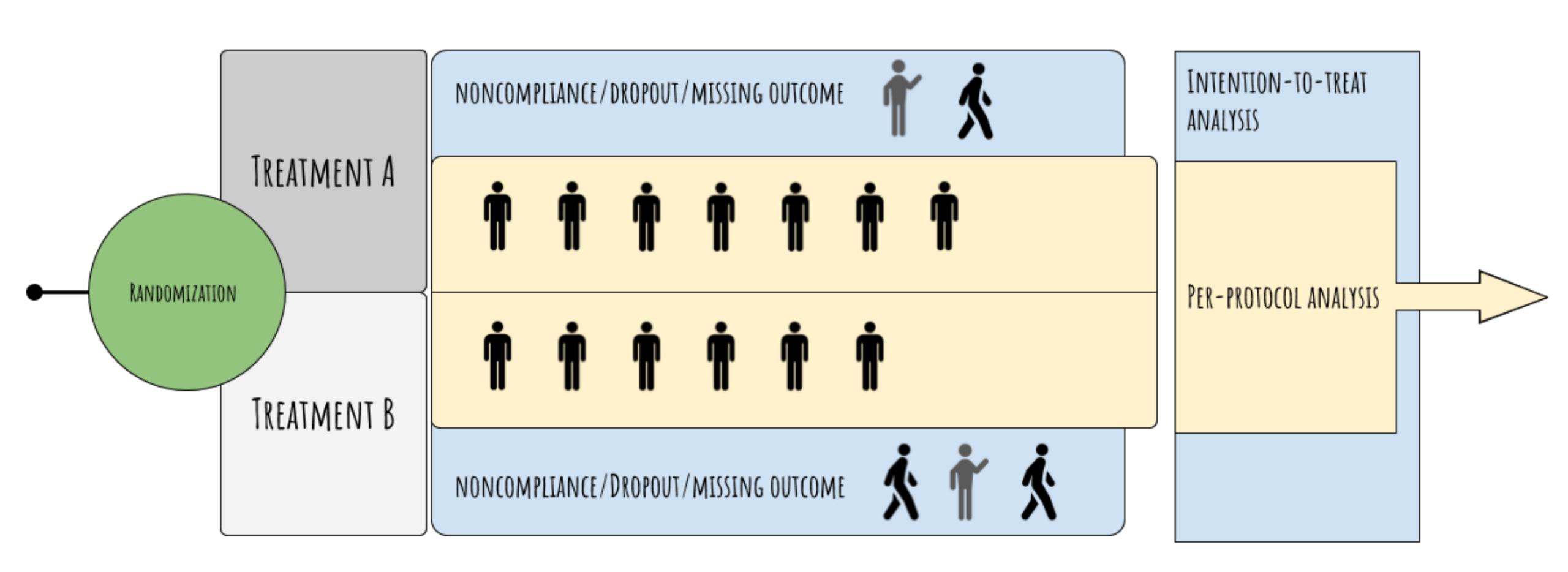
Blinding







ITT vs. PP



www.MedicoWiki.com

Outcome (POO/DOO/LOO/MOO



Large and Precise



Practice Changing?



NNT & NNH





Sackett et al BMJ 1996

There are 13 RCTs of thrombolytics for acute ischemic stroke, 4 were stopped early for harm (bleeding) or futility, and all 13 failed to show a statistical benefit after the reanalysis of NINDS-2 & ECASS-III.

Summary of Thrombolytics for Acute Ischemic Stroke

Trial	Number of patients	Journal	Time to treatment	Results
MAST -	622	Lancet 1995	<6 hours	No difference in primary benefit, increased
Italy				chance of early death
ECASS-I	620	JAMA 1995	<6 hours	No difference on disability scores and 7%
NUMBE I	201	NEINA 1005	e2 hauna	increase in mortality
NINDS-I	291	NEJM 1995	<3 hours	No difference in symptoms or 3-month outcomes
NINDS-II	333	Annals of	<3 hours	No difference in favorable mRS at 90 days,
•		EM 2009		6% absolute increase in brain bleeds and no mortality difference
MAST-	310	NEJM 1996	<6 hours	No difference in death or disability at 3 to 6
Europe				months, 18% increase in brain bleed and
				stopped early due to harm
ASK	340	JAMA 1996	<4 hours	No difference in death or disability at 3
				months, 10% increase in brain bleeds and
				stopped early due to harm
ECASS-II	800	Lancet 1998	<6 hours	No difference in outcomes on the mRS or
				mortality and 7% increase in brain bleeds
ATLANTIS-	613	JAMA 1999	3-5 hours	No difference in neurologic recovery and
В				stopped early because "unlikely to prove beneficial"
ATLANTIS-	142	Stroke 2000	<6 hours	No benefit in NIH stroke scale at 30 days,
Α				18% greater risk of mortality and stopped
				early due to harm
ECASS-III	821	BMJ_EBM	3-4.5 hours	No difference in favorable mRS score after
**		2020		90 days and 9% increased rate of brain bleed
DIAS-2	193	Lancet 2009	3-9 hours	No difference in clinical response and
				increased rate of brain bleed
IST-3	3035	Lancet 2012	<6 hours	No difference in mortality or independence
				after 6 months, 4% increase in death at 1
				week and 6% increase in fatal or non-fatal
				brain bleeding
DIAS-III	492	Lancet	3-9 hours	No difference in favorable mRS at 90 days
		Neuro 2015		and no difference in major adverse events

Reanalysis of NINDS-2 by Hoffman and Schriger Annals of EM 2009

This list does not include two newer RCTs looking at extending the therapeutic window to 4.5-9hrs. These newer trials were done with more advanced brain imaging selecting patients with a perfusion mismatch. Both RCTs were stopped early which can introduce bias towards efficacy. In addition, the majority of patients included in these trials would now qualify for endovascular therapy (EVT) clot retrieval. EVT has more evidence for efficacy than systemic thrombolysis and a recent RCT has shown that EVT alone is non-inferior to EVT plus tPA.

Lytics and AIS?

- 13 RCTs
- Zero Show Efficacy*
- Four Stopped Early Due to Harm or Futility
- Not Include Wake-Up

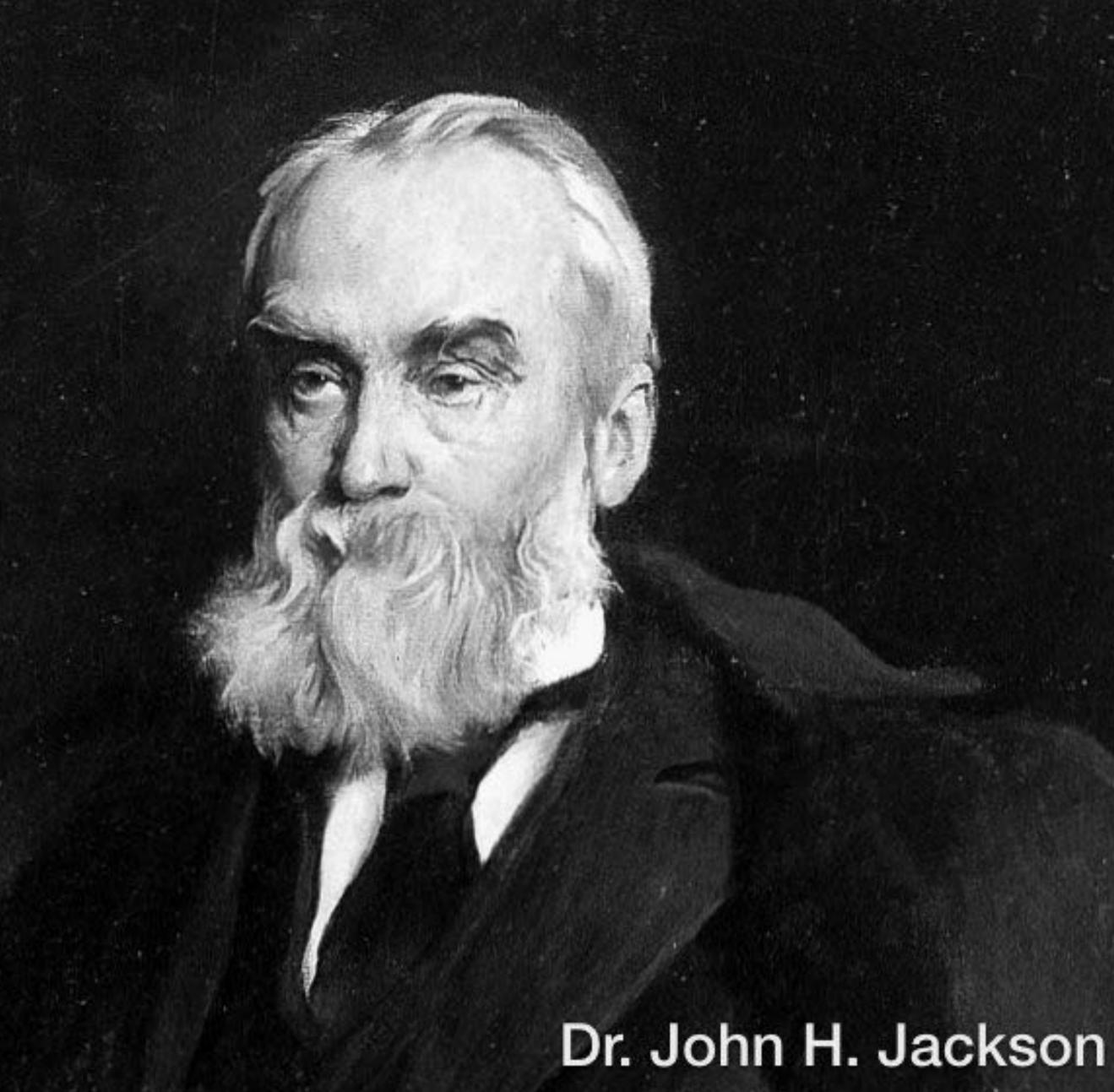
^{**} Reanalysis of ECASS-III by Alper et al BMJ_EBM 2020

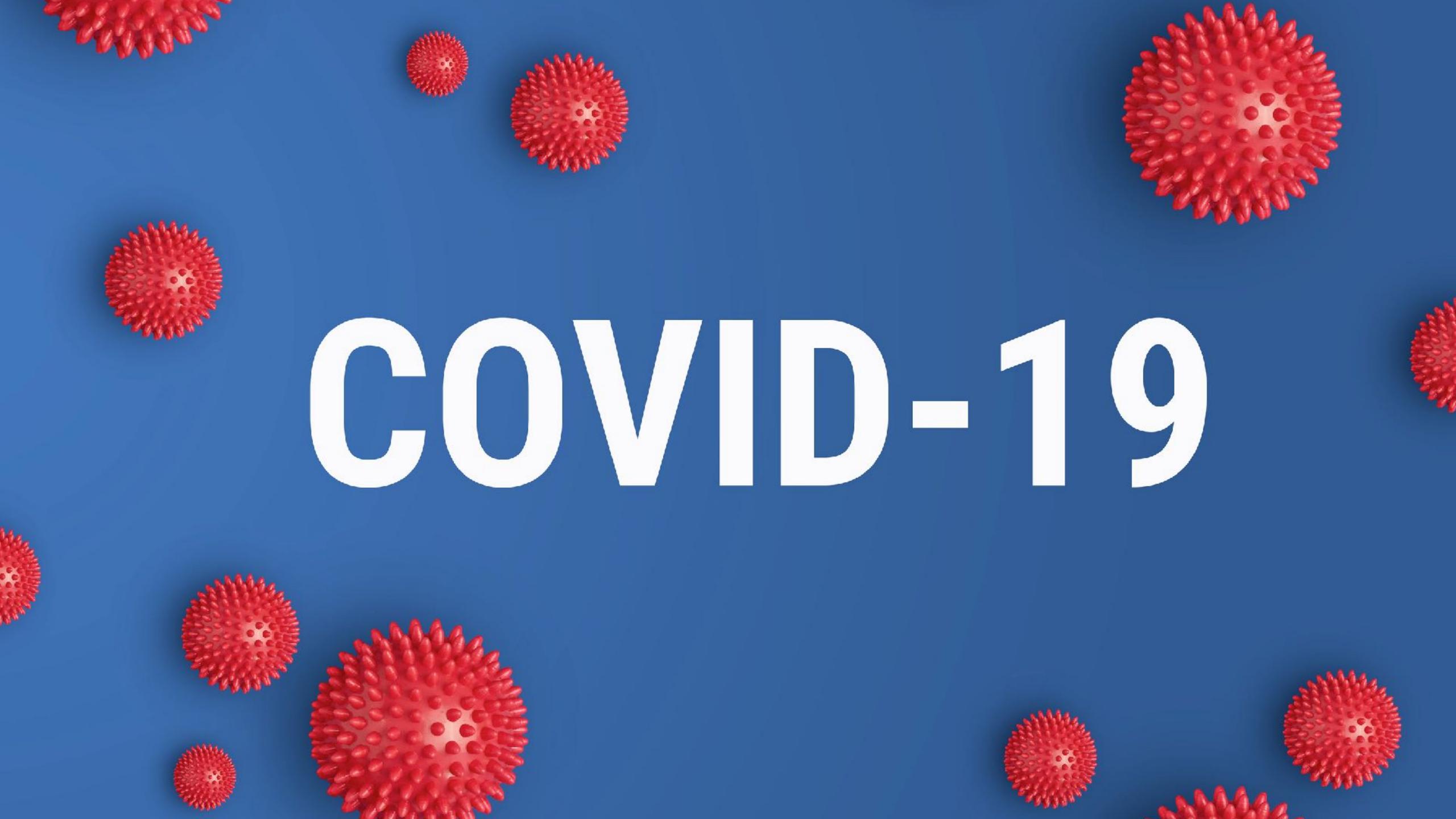
Lytics for AIS?





"It takes 50 years to get a wrong idea out of medicine, and 100 years a right one into medicine."

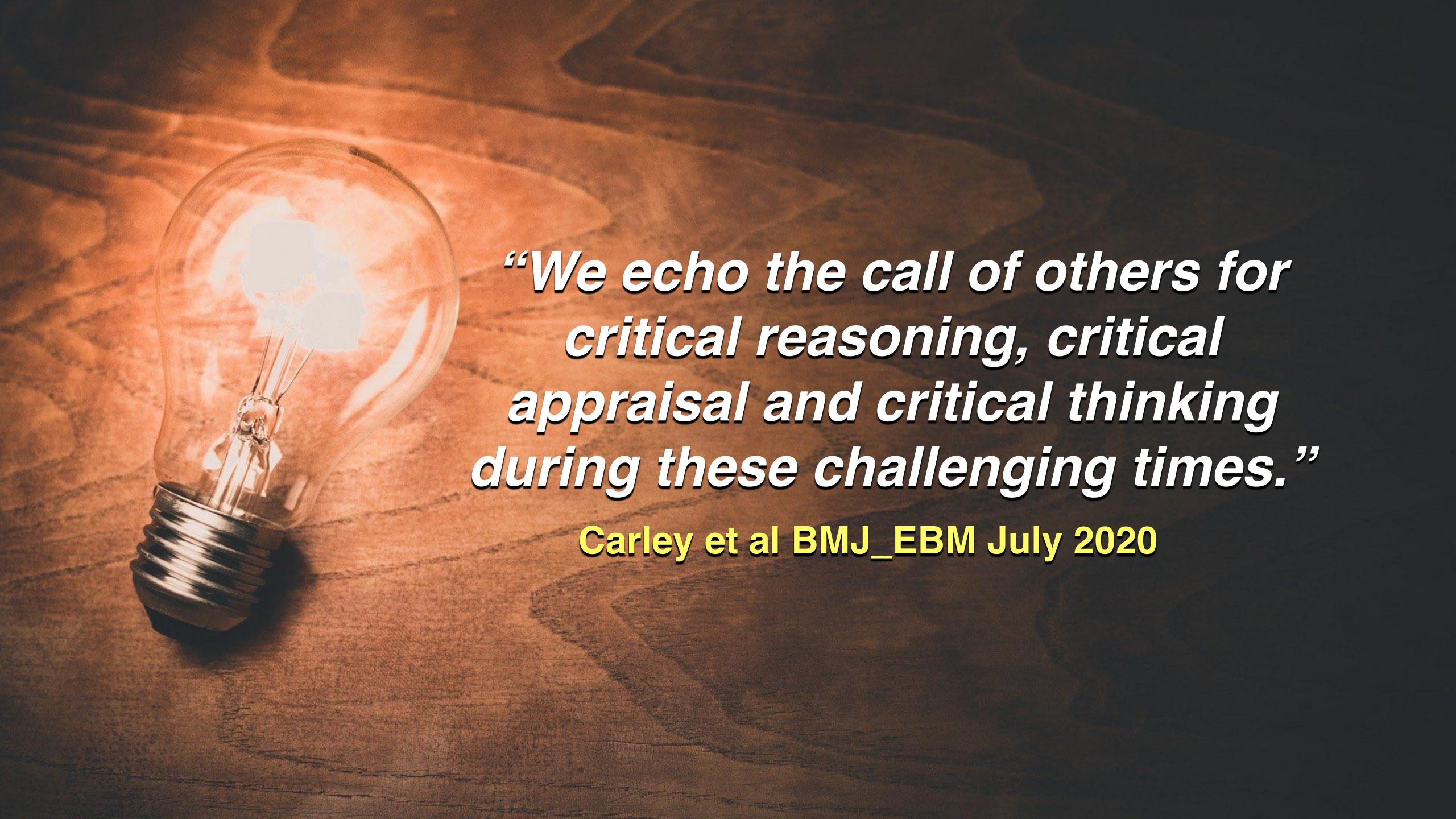








"The principles of EBM are more important now than at any other time in our careers. We must collectively do all that we can to ensure that our response to the pandemic is based on the science and not on the emotional, political or economic issues that challenge it. Carley et al BMJ_EBM July 2020



Take Home Message

