WHEN IT COMES **TO EVIDENCE-BASED** PRACTICE

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TO GET A HANDOUT GO HERE http://therapeuticseducation.org/handouts

peerevidence.ca therapeuticseducation.org





Financial Conflicts of Interest



Entire salary comes through the UBC Faculty of Pharmaceutical Sciences - also some legal/ educational work

I have received no honorarium or research money from the drug industry in the last 30 or so years





Premium podcast subscription Best Science (BS) Medicine podcast - therapeuticseducation.org

I have a self-published book called "The Nutrition Proposition"



My Simple Philosophy on Treatments

- These sorts of terms are uniformly uninformative allopathic, conventional, mainstream, Western medicine, complementary, alternative, integrative, naturopathy, Chinese medicine, natural, homeopathy, herbal
- We all treat people with "things" oral/IV/IM/topical, nutrition, surgery, talk, physical manipulations etc

I don't care HOW treatments work, care IF treatments work

TI Letter #138

The proportion of people over 65 taking prescription medications

14% took 5 or more drugs 42% took 5 or more drugs 50% took 1 to 4 drugs 49% took 1 to 4 drugs 26% took no drugs 9% took no drugs 114% took 5 or more drugs 1100 to 4 drugs 114% took 1 to 4 drugs 9% took 1 to 4 drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114% took 1 to 4 drugs 9% took no drugs 114 to 1 t	14% took 5 or more drugs 42% took 5 or more drugs 50% took 1 to 4 drugs 9% took 1 to 4 drugs 26% took no drugs 1144444 11444 114444 1144 1144444 1144 1144444444 1144 1144444444444444444444444444444444444	1994	2014
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"in the vast majority of circumstances, the only outcome of relevance for EBP is to measure whether a shared decision was made"



doi:10.1136/bmjebm-2018-110922

Satisfaction is linked to shared decisions



Communicating with patients on health care evidence. Discussion Paper, Institute of Medicine, Washington, DC 2012

Where SDM may not work

 OPEN ACCESS
 Shared decision is the only outcome that matters when it comes to evaluating evidence-based practice

 James McCormack,¹ Glyn Elwyn²
 2018

In most societies there are laws that prevent certain harm from occurring, where mental incapacity or strong personal beliefs may threaten the well-being of others

- 1. Jehovah Witness' refusal to transfuse blood to those in dire need
- 2. involuntary detention for psychiatrically unstable patients who risk harming themselves or others
- 3. surrogates are asked to make decisions for those people truly unable to consent to treatment in immediate life-threatening situations
- 4. smoking bans that lead to important reductions in morbidity and mortality
- 5. an intriguing example that some would consider an important exception is mandatory vaccination with the potential of herd immunity. In this case, a shared-decision not to be vaccinated for a transmissible disease could lead to inherent harm of others.



Its not that difficult





My Agenda



Much of what we do, even with the best of intentions, is not that effective Most guidelines are a BIG problem

Some treatments (medications, nutrition, activity) can be effective and even life-saving BUT many aren't and they all have the potential for harm, inconvenience and cost

I believe the size of the effect for many of these treatments is much smaller than people think

Lab test variation makes many tests (especially repeat tests) of questionable use and are simply misleading

The recommended doses for most medications are too high

IT'S GOTTEN SILLY



PRE-EVERYTHING

Pre-diabetes **Pre-hypertension**

Pre-osteoporosis

(osteopenia)

Pre-elderly



Pre-menopausal **Pre-obesity Pre-death Post-birth**

AVOID Misleading Terminology

"Significant" "Use with caution" "Use with extreme caution" "Monitor closely" "High risk" "Very high risk" "Really !@#\$% high risk"

What is "High Risk"



Chance of a heart attack in the next 5 years (%)

A 60 y/o, male, smoker, diabetic, SBP 180, total cholesterol 7.2 mmol/L 5-year risk of heart attack PLUS stroke is at most ~ 25%

The Magnitudinous Problem



Severe Weak Strong Different Faster Shorter Longer Shortened Lengthened Extreme Unlikely Short Many/Most

All these words likely mean something different to everyone

Examples that probably require quantification clarification

Your salary will be **INCREASED**

Turn left after a **MODERATE** number of kilometres

You will be getting a **SHORT** jail sentence

You have an **UNLIKELY** chance of getting an STD

You have a **SIGNIFICANT** chance of a heart attack

A **SMALL** tube will be placed a **CONSIDERABLE** distance into your rectum

Beware of "qualitative quantification"

Qualitative descriptor	EU assigned frequency	Mean frequency by participants (estimated (n=200)
Very common	>10%	65% (24·2)	
Common	1-10%	45% (22·3)	OFF BY
Uncommon	0.1-1%	18% (13·3)	~350% to 18,000%
Rare	0.01-0.1%	8% (7.5)	
Very rare	<0.01%	4% (6·7)	

Values are mean (SD).

Lancet 2002;359:853-54

Let's first address the ELEPHANT in the room









and follow **Clinical Practice Guidelines**

"Standard of Care"

Clinical Practice Guidelines

Medicolegal Sidebar: Clinical Practice Guidelines—Do They Reduce Professional Liability Risk?

Joseph P. McMenamin MD, JD, Wendy Teo BA(Cantab), BM BCh (Oxon), LLM, B. Sonny Bal MD, JD, MBA, PhD

"Clinical practice guidelines, however, are designed to improve care, not to define standard care. They can also limit physician autonomy, impose rules that are adopted mainly to avoid litigation risk, and may be developed by physicians with relevant financial conflicts. In our view, courts should exclude clinical practice guidelines from evidence of the standard of care or of its breach."

Clin Orthop Relat Res (2020) 478:23-25

Patient preferences for shared decisions: A systematic review

Betty Chewning^{a,*}, Carma L. Bylund^b, Bupendra Shah^c, Neeraj K. Arora^d, Jennifer A. Gueguen^e, Gregory Makoul^f

"the number of patients who prefer participation has increased over the past three decades so that the majority of patients prefer to participate in decisions"

Patient Educ Couns (2011), doi:10.1016/j.pec.2011.02.004

Factors involved in deciding to start preventive treatment:qualitative study of clinicians' and lay people's attitudesDavid K Lewis, Jude Robinson, Ewan WilkinsonBMJ 2003;327:841

"Many of the preferences expressed by the clinicians and lay people in this study are at odds with recommendations in guidelines"

> Differing perceptions of intervention thresholds for fracture risk: a survey of patients and doctors Osteoporos Int 2012;23:2135–40

77% of doctors would recommend treatment21% of our patient cohort would consider treatment justified



"There is insufficient evidence to determine whether or not shared decision-making and the use of decision support interventions can reduce medical malpractice litigation. Further investigation is required."



Reducing litigation risk 2 THINGS to DO

Shared decision-making model

1) Use a decision aid

2) Document decision



"I would rather know evidence and try to apply it to each patient, than memorize guidelines and try to apply them to all patients"

Mark McConnell

Making An Example Of Food!



Setting the Stage

Food can absolutely can have a large effect on health



Vitamin C deficiency



Macronutrient deficiency



You really don't need RCTs or cohort studies to evaluate this



The Evidence Issues



Cohort studies and RCTs

Surrogate (blood pressure/glucose/lipids) versus clinical outcomes (heart attack/stroke/mortality)

Relative versus absolute numbers

A 50% reduction (relative) has very different implications if the baseline risk is 10/1,000 versus 10/100

"This food is healthy/unhealthy" – the size of the effect is key Lab tests and what do they mean Examples of BS Food = values and preferences

9 Health Benefits of Broccoli, According to a Nutritionist

Broccoli offers disease-fighting nutrients that can decrease signs of aging, among other benefits.

By **Stochia Sass, MPH, RD** Updated January 27, 2020

Broccoli has a bounty of nutrients Broccoli is high in fiber

It may help prevent cancer

Broccoli offers heart protection

It's linked to brain health

Broccoli can keep bones strong

It fights inflammation

It's a natural detox

Broccoli has antioxidant protection

The article starts with

"You KNOW that broccoli is GOOD for you" It is a "SUPERFOOD"

> At best low quality evidence and in general **NO evidence**

HARVARD HEALTH BLOG

10 superfoods to boost a healthy diet

April 13, 2020



https://www.gousto.co.uk/blog/top-10-superfoods

In 2015 alone - 36% rise in the number of food and drink products launched globally featuring the terms "superfood", "superfruit" or "supergrain".

It's really easy to simply state these things are good or bad for your health

Drinking 2, 4, 6 or 8 glasses of water a day Drinking 0, 1, 2 or 3 alcoholic beverages a day Eating 2, 3, 4, 5, 6 or 7 servings of fruits and vegetables a day Eating 0, 1, 2 or 3 eggs a day Adding salt to food Restricting or increasing the amount of carbs, fat and protein Adding sugar to 1, 2, 3, 4 or 5 cups of coffee or tea a day Being a meat eater, a vegetarian, or a vegan Eating a doughnut, cheesecake, ice cream, or chocolate Drinking a glass of milk or a soft drink a day Eating an apple a day

Everything is "linked"



"Clogged" arteries



"Smart" words



Images with "arrows"



"Medical" References



The Bullshit Asymmetry





The amount of energy needed to refute bullshit is an order of magnitude bigger than to produce it.

Annals of Internal Medicine	IDEAS AND OPINIONS
U.S. Dietary Guidelines: An Evidence-Free Zou Steven E. Nissen, MD	ne 2016

"a detailed review of the new guidelines confirms a disturbing reality: the nearly complete absence of high-quality randomized, controlled clinical trials (RCTs) studying meaningful clinical outcomes for dietary interventions. The report repeatedly makes recommendations based on observational studies and surrogate end points, failing to distinguish between recommendations based on expert consensus rather than highquality RCTs. **Unfortunately, the current and past U.S. dietary guidelines represent a nearly evidence-free zone**"

2015 DGAC: MEETING 7 December 15, 2010 Chole THE FINAL REPORT FROM THE GUIDELINE HOWEVER, THE FINAL REPORT FROM JANUARY 2016 STATED, HOWEVER, THE FELEASED IN JANUARY 2016 STATED, HOWEVER, THE FELEASED IN JANUARY 2016 STATED, HOWEVER, THE FELEASED IN JANUARY 2016 STATED, HOWEVER, THE FINAL REPORT FROM JANUARY 2016 STATED, HOWEVER, HOW Science Base Chapter:

SETTING THE TABLE

The "Studies" "Numbers" "Outcomes"

Simple But Elegant

My Nutrition Proposition

- 1. Eating food is one of life's greatest joys
- 2. Excessively "worrying" about food is just wrong
- 3. If you wish to make nutrition decisions based on health reasons you should know the best available evidence, or lack thereof, and appreciate the context of that evidence
NO RCTS LOOKING AT IMPORTANT CLINICAL OUTCOMES

AN EXAMPLE



The health impact of being a vegetarian vs a carnivore **SURROGATE markers** (lipids, blood pressure etc) The impact that has on **ESTIMATED heart attack/stroke risk**



REASONABLE ESTIMATE OF THE IMPACT ON RISK

~1% absolute decrease over 10 years

~2% absolute decrease over 20 years

Studies of the Mediterranean diet show it produces minimal if any changes on surrogate markers

Very important things I'm not discussing

Animal rights
 Environmental issues



STRICTLY SHOWING HEALTH EVIDENCE

The BIG Question We Have

What foods are delicious healthy?





SHOWS CAUSE AND EFFECT

SHOWS ASSOCIATION

We are bombarded with health claims



Hierarchy of WTF Wellness blogger Celebrity or something overheard on public transit Personal anecdote from your cousin, brother, girlfriend

Dr Google or an info-commercial

"Health professional" newly graduated from a 12-week online course

Highly educated person with university degrees in something totally unrelated to health/nutrition - law, journalism, aerodynamics

Stolen from Kevin Whelan

RCTs can show cause and effect



From 2008 to 2015 20 LARGE TRIALS IN A ROW SHOWED NO BENEFIT FROM CHANGING A SURROGATE MARKER LIPIDS

DIABETES

ACCORD. ADVANCE. VADT (aggressive A1c lowering) ROADMAP (olmesartan) ORIGIN (insulin) SAVOR-TIMI 53 (saxagliptin) EXAMINE (alogliptin) ALECARDIO (aleglitazar)

GENERAL ACTIVE (irbesartan/afib)

CRESCENDO (rimonabant) VISTA-16 (varespladib)

5 cholesterol trials 8 diabetes/glucose trials **4 blood pressure trials 3** general risk reduction trials

AIM-HIGH, HPS2-THRIVE (niacin) ACCORD (fibrates) dalOUTCOMES (dalcetrapib) STABILITY (darapladib)

BI OOD PRESSURE

ALTITUDE (aliskiren) VALISH, AASK, ACCORD (aggressive BP lowering)

FINALLY!!!!2015

1) EMPA-REG OUTCOME (empagliflozin) -1.6% + over 3 years

2) LEADER (liraglutide) - 1.8% + over 4 years

3) SPRINT (120mmHg vs 140mmHg) - $1.6\% \downarrow$ (CVD) over 3 years but also $1.8\% \uparrow$ (Kidney) 4) HOPE 3 - statins YES, BUT blood pressure no benefit

5) FOURIER - 1.6% - over 2 years BUT \$15,000/year

BUT!!!!!

1) ACCELERATE (evacetrapib - increased HDL (130%), reduced LDL (40%) - **NO CVD benefit**



The Numbers - cohort studies Almost always

Look at **10,000-100,000s** of people - looking for a **small** absolute difference Benefits/harms always described as relative numbers - while "**correct**", they are **misleading**

Differences seen typically **ONLY** when extremes of intake are compared

LOWEST quartile/quintile vs the HIGHEST quartile/quintile





NEJM - 229,000 men - 13 years - 1% absolute **↓**

As compared with men who did not drink coffee, men who drank 6 or more cups of coffee per day had a 10% lower risk of death, BMJ - 81,000 people - 8 years - 1% absolute 1

An increase in total red meat consumption of at least half a serving per day was associated with a 10% higher mortality risk

WEIGHT - the Battle of the Bulge













WORDS

words have direct and indirect impacts

on the people who hear or read them

MATTER

WHEN IT COMES TO WEIGHT

WHEN IT COMES TO WEIGHT



Focus on the HEALTH aspect NOT the size







Percentage of the population in select countries worldwide who were trying to lose weight in 2020

45% of people globally are trying to lose weight

What is the "Best Weight"

"The 'best weight,' is whatever weight a person reaches when they're living the healthiest life they can actually enjoy and they don't have any BMI, weight or waist circumference goals."

Yoni Freedhoff Associate Professor, Family Medicine, University of Ottawa

A Weighty Conclusion

Categorize people using BMI/unhealthy weight rather than size words

Waist and hip-waist ratio measurements and what sort of "fruit" you are doesn't really inform much about health risks over and above BMI

The 1/3 of the population that is categorized as "overweight" really have minimal if any associations with worse health outcomes - especially if they are physically active

Alcohol ingestion can absolutely be harmful

The psychosocial impacts of alcohol ABUSE are devastating to individuals, families and the general public - cirrhosis, violence, accidents

Drinking and driving is 1000% wrong - SELFISH!!

Binge drinking can lead to very poor judgments



CONTEXT

MATTERS

Anything more than 3 drinks a day is likely a health issue BUT what about 1, 2, or 3

A History Lesson

2011

REPORT

Alcohol and Health in Canada: A Summary of Evidence and Guidelines for Low-Risk Drinking

Alcohol Health Effects

Publication date: 2011 Author: Canadian Centre on Substance Use and Addiction

> "no more than 10 drinks a week for females and 15 drinks for males"

Do not drink and drive Do not drink when pregnant







vidence. Engagement. Impac

The Red Wine Theory

BEFORE THE 1980s ALCOHOL WAS SIMPLY CONSIDERED HARMFUL

1980s researchers found in France that despite a diet relatively high in saturated fat - lower risk of heart disease - the "French Paradox"

The theory - antioxidants - polyphenols in wine - resveratrol

SURROGATE MARKERS

2 RCT meta-analyses - **resveratrol** - **NO EFFECT** on BP/LDL/HDL/TG, but 4 total cholesterol by ~5%

Beer and wine - NO EFFECT on total cholesterol/LDL/TG/BP, but 1 HDL by 8%

IMPACT THESE SURROGATE MARKER CHANGES WOULD MAKE ON CVD ESTIMATES

10-year cardiovascular risk estimates go from ~5% down to 4.7%

HEALTH

Proposed update to Canada's alcohol guidelines suggests as few as 3 drinks per week

By Cassandra Szklarski · The Canadian Press Posted August 30, 2022 1:23 pm · Updated August 30, 2022 6:42 pm



HEALTH News

Proposed alcohol guidelines recommend no more than 2 drinks per week

A new measure of unhealthy drinking

PUBLISHED SEPTEMBER 1, 2022

If you have three or more alcoholic <u>drinks in a week</u>, you're putting your health at risk. That's according to a <u>new report</u> from the Canadian Centre on Substance Abuse and Addiction (CCSA). The government of Canada's <u>current recommendations</u> are more than a

CALGARY News

Calgarians react to new guidelines for alcohol intake

Having three to six drinks per week increased the risk to moderate, while having more than six was found to contribute to increased risks of cancer, stroke, heart disease and situations of violence.

HOME > LOCAL NEWS

1 drink a day means higher risk of heart disease, stroke, cancer: Report

A recent report highlights the many health risks associated with consuming just one alcoholic drink a day

Michael Ranger Sep 5, 2022 3:00 PM



Living

Are Canadians drinking too much alcohol?

By NetNewsLedger - September 7, 2022

• 182



Are Canadians Drinking too much?



Recommended maximum intake of alcoholic beverages



How Much Do We drink?

	Zero	If you do drink - typical drinking day				
		1-2/day	1-2/day 3-4/day			
Women	23% 74%		17%	9%		
Men	18%	54%	23%	23%		

2005 (over the past year) - https://www.ccsa.ca/sites/default/files/2019-05/ccsa-004028-2005.pdf

Alcohol

Lancet 2018

Alcohol use and burden for 195 countries and territories, 1990–2016: a systematic analysis for the Global Burden of Disease Study 2016

GBD 2016 Alcohol Collaborators*

"We found that the risk of all-cause mortality, and of cancers specifically, rises with increasing levels of consumption and the level of consumption that minimises health loss is zero"





ABSOLUTE NUMBERS - the number who would experience an alcohol related problem

OVER ONE YEAR	Additional people out of 100,000	Extrapolated Increase over 30 years			
1 drink a day	4	0.1% or 1/1000			
2 drinks a day	63	1.5%			
5 drinks a day	338	10%			

TOP 3 HARMS tuberculosis, road injuries, self harm



Lancet 2018

Alcohol use and burden for 195 countries and territories, 1990–2016: a systematic analysis for the Global Burden of Disease Study 2016 CRI 2016 Automatication and Au

Alcoho					
RESULTS FROM THE data taken from the	Absolute change in the number of deaths/100,000				
22 outcomes	Males	Age standardized death rate per 100,000 for each of these outcomes	(calculated by multiplying the age standardized death rate by the relative difference found)		
Ischemic heart disease	Benefit any amount	Benefit any amount	~15% ↓ or 0.85	142	21↓
Hemorrhagic stroke	Harm if >2 drinks/day	None	~25% 1 or 1.25	52	13 🕇
Ischemic stroke	None	Benefit if 1-2 drinks/	~20% ↓ or 0.80	49	10 🖡
Lower respiratory tract infections	None	None	~0% or 1.00	42	0
Unintentional injuries	Harm any amount	Harm any amount	~15% † or 1.15	27	41
Diabetes	None	Benefit	~25% ↓ or 0.75	24	6↓
Transport injuries	Harm any amount	Harm any amount	~20% 🕇 or 1.20	20	41
Cirrhosis	Harm if >1 drink/day	Harm if >1 drink/day	~100 ↑ or 2.00	19	19
Self harm	None	None	~0% or 1.00	17	0 or
Interpersonal violence	Harm if >1 drink/day	Harm if >1 drink/day	~25% † or 1.25	17	4.3 🕇
Tuberculosis	Harm if >1 drink/day	Harm if >1 drink/day	~70% 🕇 or 1.70	16	111
Hypertensive heart disease	Harm if >1 drink/day	Harm if >1 drink/day	~30% † or 1.30	15	51
Colon/rectum cancer	Harm any amount	Harm any amount	~15% † or 1.15	13	21
Liver cancer	None	None	~0% or 1.00	12	01
Breast cancer	Harm any amount	Harm any amount	~25% † or 1.25	8	21
Esophageal cancer	Harm if >1 drink/day	Harm if >1 drink/day	~50% † or 1.50	7	41
Atrial fibrillation/flutter	Harm any amount	Harm any amount	~10% 1 or 1.10	3	<1↑
Pancreatitis	None	None	~0% or 1.00	2	0
Larynx cancer	Harm if >2 drinks/day	Harm if >2 drinks/day	~40% † or 1.40	2	11
Epilepsy	Harm any amount	Harm any amount	~30% † or 1.30	2	11
Lip and oral cavity cancer	Harm any amount	Harm any amount	~50% † or 1.50	2	11
Pharnyx/nasopharynx cancer	Harm any amount	Harm any amount	~80% † or 1.80	1	11
				2	

MAIN ASSOCIATIONS

Relative Numbers

DECREASE 15-25% **↓**

in heart attack/strokes INCREASE

unintentional injuries 15% transport injuries 20% tuberculosis 70% atrial fibrillation 10% colon/esophageal/breast/

nasopharynx/lip cancers 15-80% **↑**

Absolute change in number of deaths/100,000



Even in small quantities, drinking alcohol has consequences for everyone, whether you are male, female, younger or older. In fact, it's biological, it's physical.

That's why drinking less is better!



THIS IS THE PUBLIC SUMMARY (August 2022)

created by the Canadian Center on Substance Use and Addiction and they asked for public consultation

> The terms small, low, moderate, increasingly high risk are too subjective and in no way inform people as to the actual size of the risks

Not sure the weekly amount is all that useful likely better to think about drinks per day given that when people "drink", they drink "daily"

> Not sure of the point of having a weekly target of drinks - kind of sounds like a challenge to achieve either high or low

There are no numbers here and it implies each category has only the risks listed - there is no mention of liver cirrhosis which may numerically be the largest risk

Public Consultation: Summary of Key Actions Taken

The responses received from the open consultation were analyzed and categorized. The table below presents the main categories of comments as well as the actions taken by the LRDG-Scientific Expert Panel (LRDG-SEP) to address comments which fell within the scope of this project's mandate.

There were several suggestions made for knowledge mobilization activities, including knowledge synthesis, dissemination, transfer and exchange. These suggestions have been recorded but are not listed here as they could not be considered for action (i.e., could not lead to edits and revisions of the final report).

Consultation comment or suggestion		Action taken			
Public Summary					
Provide more information about specific cancers.		There are already many consequences of different types presented in the public summary, the suggested consequences were not added. However, the public summary was edited to link the average amount of weekly			
In gene	The objective of the docume information without statistic information and more expla- understood. No statistics we	ent is to communicate s that would need contextual nations to be easily ere added.			

The Top 5 Harms



4 were the same for men and women



Lifetime cancer risk Breast cancer



lifetime risk of dying would increase from 3% to roughly 3.5%

Colorectal cancer

lifetime risk of dying would increase from 3.0% to roughly 3.3%

Cirrhosis

CCSA reports that 1-2 drinks a day 1 the risk of liver cirrhosis in both men and women

But the single paper they use to support these claims states quite clearly that, "although consumption of 1–2 drinks was associated with a substantially elevated risk for liver cirrhosis in women, **this was not the case in men**"

based on the CCSA numbers



My Opinion The 2023 CCSA Alcohol Guidelines:

- 1. Are misleading
- 2. Don't provide appropriate "context"
- 3. Create unnecessary fear and confusion
- 4. In no way inform the public as to the absolute risks/benefits
- 5. Very likely have nothing to do with your values and preferences
- 6. Ignore the research (although it's not great) around the functional social benefits they state it was "out of the scope for this summary" yet their research question clearly states "What are the risks and **benefits** (physical and mental health, and social impact)"

A number of their harm comments are not supported by their own data. Their data show a CVD benefit at 1 drink a day that is greater than the cancer risks and this is not mentioned

Alcohol Risk Visualizer

Based on the latest CCSA report on the lifetime risk of alcohol-attributable death and disability.

This chart shows how many days of life, on average, an individual could lose based on the amount of drinks they have per week.

The CCSA considers one drink as:

- 341 ml (12 oz) of beer 5% alcohol or cooler
- 142 ml (5 oz) of wine 12% alcohol
- 43 ml (1.5 oz) of spirits (whiskey, vodka, gin, etc) 40% alcohol



Combined Risk from 21 Different Health Outcomes

Disease selectors allow you to select the diseases you're interested in. For example, consider removing physical injuries if you don't drink and drive and you are not reckless when you drink.

•	Cancer	Cardiovascular Diseases	Liver Damage	Physical Injuries	2	Other
	Oral cavity and pharynx	Diabetes	 Liver cirrhosis 	Road injuries		Tuberculosis
	cancer	Atrial fibriliation and flutter		Other unintentioal injuries		Lower respiratory infections
	Oesophagus cancer	Hypertension		Intentional injuries		Pancreatitis
	Colorectal cancer	Ischemic heart disease				Z Epilepsy
	Liver Cancer	Ischemic stroke				
	Breast cancer	Intracerebral hemorrhage				
	Larynx Cancer	Subarachnoid baemorrhage				

ASSUMING THEIR NUMBERS ARE CORRECT WHICH IS A BIG ASSUMPTION Remove injuries/cirrhosis/tuberculosis 1 drink a day for life = just over 2 week of life lost 2 drinks a day for life = just over 3 months of life lost

The Bottom Line

If you have a history of an alcohol problem or are pregnant - DON'T DRINK

If you drink and drive, become aggressive when you drink, or have a history of doing stupid things when you drink - DON'T DRINK TO EXCESS

3-5 drinks/day

Very likely a health issue

2 drinks/day

at most a ~1% increase in mortality over a lifetime

1 drink/day

doesn't seem to produce an INDIVIDUAL health risk OR benefit

BUT THE EVIDENCE IS TRICKY AT BEST







Coffee and mortality/heart attack/stroke/cancer associations

CLINICAL OUTCOMES	RESI Risk ratio (confi	RESULTS Risk ratio (confidence intervals)			
	High versus low intake	For each extra cup			
All cause mortality	10% 🖡	4%↓			
Heart attack mortality	No change	Not reported			
Heart attack/stroke mortality	No change	No change			
Stroke mortality	No change	Not reported			
After a heart attack mortality	45%↓	Not reported			
Cancer mortality	No change	No change			
COHORT STUDIES	Meta-analysis - 2017 Poole				

Tea and mortality associations

CUNICAL OUTCOMES	RESULTS Risk ratio (confidence intervals)				
	Highest versus lowes 3-28	Per 1 cup/day 3-28 years			
	Green tea	Black tea	Теа		
All cause mortality (highest vs lowest)	20% I	10% I	Not reported		
All cause mortality (1 cup/day)	4%↓	3%↓	2%↓		
Heart attack/stroke mortality (highest	33%↓	No change	Not reported		
Heart attack/stroke mortality (1 cup/	5%↓	8%↓	4%↓		
Cancer mortality	No change	21%↓	Not reported		
Heart attack/stroke events	Not reported	Not reported	No change		
COHORT STUDIES	Meta-analysis - 18 studies 2015 Tang		Meta-analysis - 39 studies 2020 Chung		

Coffee consumption and health: umbrella review of meta-analyses of multiple health outcomes

9 outcomes

Intakes of three to four cups a day versus none

for all cause mortality 17%↓, cardiovascular mortality 19%↓, and cardiovascular disease 15%↓

High versus low consumption - incident cancer 18%↓

Consumption was also associated with a **lower risk** of several specific cancers and neurological, metabolic, and liver conditions

In pregnancy - high versus low/no consumption

low birth weight 31% ↑, preterm birth in the first 22% ↑ and second 12% ↑ trimester, and pregnancy loss 46% ↑

BMJ 2018; 360 doi: https://doi.org/10.1136/bmj.k194

Effect of coffee on surrogate markers

SURROGATE	RESULTS Change in surrogate marker							
OUTCOMES	45 days (2.4-8 cups) for lipids, 62 days (2- ≥5 cups) for blood pressure							
	Overall	Overall Filtered Unfiltered Caffeinated De						
Total cholesterol	1 4%	1 2%	1 6%	1 5%	No effect			
LDL cholesterol	1 4%	No effect	1 9%	1 4%	No effect			
HDL cholesterol	No effect	Not reported	Not reported	Not reported	Not reported			
Triglycerides	1 8%	No effect	1 3%	1 9%	No effect			
Blood presssure	No effect	Not reported	Not reported	Not reported	Not reported			
CLINICAL TRIALS	Meta-analysis - 2017 Poole							





 \mathbf{x}

Meat - it's about your "values"



					Mortality		Overall cardiovascular	
Message	The two different meta-analyses of cohort studies	# of cohorts	What was examined	Time	Unprocesssed meat	Processed meat	Unprocessed meat	Processed meat
Continue to eat meat group	Zeraatkar October 2019	55	A 3 serving/ week REDUCTION*	11yr	1 8%∗ Absolute 1 ~1%	19%∗ Absolute 1%	1 5%∗ Absolute 1 <0.5%	1 3%∗ Absolute 1 <0.5%
Eat less meat group supported	Zhong February 2020	6	Each additional 2 serving/week INCREASE	19 yr	1 3% Absolute 1 ~1%	↑3% Absolute ↑~1%	↑3% Absolute ↑~0.5%	↑ 7%** Absolute ↑~2%

*Because the Zeraatkar meta-analysis examined a REDUCTION in meat intake and the Zhong meta-analysis examined an INCREASE in meat intake numbers the Zeraatkar numbers have been inverted so they can be directly compared to the Zhong numbers ** for this number 2 versus zero servings a week, not 2 servings/week increase

So Why the Different Response?

NutriRECS

Focused exclusively on health outcomes associated with meat and did not consider animal welfare and environmental issues. Also felt a 1% risk in 11 years was small

THI

Appear to think of this as more of a public health issue and that 1% risk means millions (1% of 300 million) could be affected and also considered the environmental perspective
ARTICLES https://doi.org/10.1038/s41591-022-01968-2 OPEN Systematic review and meta-analysis Health effects associated with consumption of unprocessed red meat: a Burden of Proof study

Oct 2022

Table 2 Strength of the evidence for the relationship between unprocessed red meat consumption and the six health outcomes analyzed									
Health outcome	ROS	Average BPRF	Star rating	RR at 50 g d ^{−1} (conservative 95% UI)	RR at 100 g d ⁻¹ (conservative 95% UI)				
Colorectal cancer	0.06	1.06	2 stars	1.3 (1.01, 1.64)	1.37 (1.01, 1.78)				
Breast cancer	0.03	1.03	2 stars	1.26 (0.98, 1.56)	1.26 (0.98, 1.56)				
IHD	0.01	1.01	2 stars	1.09 (0.99, 1.18)	1.12 (0.99, 1.25)				
Type 2 diabetes	0.01	1.01	2 stars	1.14 (0.97, 1.32)	1.23 (0.96, 1.52)				
Ischemic stroke	-0.02	0.98	1 star	1.05 (0.97, 1.12)	1.15 (0.93, 1.4)				
Hemorrhagic stroke	-0.13	1.14	1 star	0.9 (0.64, 1.26)	0.87 (0.56, 1.35)				

The ROS represents the signed value of the log BPRF averaged across the 15th to 85th percentiles of exposure: the lower (if harmful) or higher (if protective) uncertainty interval—inclusive of between-study heterogeneity—for the RR curve for each risk-outcome pair. ROSs are directly comparable across outcomes and each risk-outcome pair receives an ROS based on the final formulation of the risk curve. For hemorrhagic stroke, the ROS reflects a protective effect of red meat consumption, whereas for the other outcomes it reflects a harmful effect. Negative ROSs indicate that a conservative interpretation of the available evidence suggests there may be no association between risk and outcome. For ease of interpretation, we have transformed the ROS and BPRF into a star rating (1–5), with a higher rating representing a larger effect and stronger evidence.

"In other words, given all the data available on red meat intake and risk of a subsequent outcome, we estimate that consuming unprocessed red meat across an average range of exposure levels increases the risk of subsequent colorectal cancer, breast cancer, IHD and type 2 diabetes at least slightly compared to eating no red meat (by at least 6%, 3%, 1% and 1%, respectively)."



Nature Medicine 2022;28:2038-2044

Data taken from supplementary information document

1/4 pounder 100g

Lifetime risk = 4%

A recent example of Nutrition Evidence Propaganda



Nutrition Propaganda



The Stanford Twin Study It is a VERY well done trial **BUT**...





Food documentary "You Are What You Eat: A Twin Experiment." 4-part series chronicles the study from start to finish



Original Investigation | Nutrition, Obesity, and Exercise Cardiometabolic Effects of Omnivorous vs Vegan Diets in Identical Twins A Randomized Clinical Trial Nov 2023

8 week trial

22 pairs of twins - pretty much removes genetic issues ~40 y/o, 75% female randomized the twins - healthy vegan or healthy omnivorous diet - delivered meals/self-provided Baseline kcal/day started at ~1,950 kcal - then Vegan ~1650 kcal and Omnivore ~1850 kcal Baseline cholesterol started at ~250mg/day - then Vegan ~0mg/day and Omnivore ~ 325-500mg/day

8-week results for vegan versus omnivore

Vegan - LDL ↓ 14mg/dL ~12% - statistically different

Vegan - HDL ↓ 4mg/dL ~ 7% - not statistically different

Vegan - Weight 4 2 kg - statistically different

Vegan - fasting insulin ↓~17% - statistically different

Triglycerides, vitamin B12, glucose, and TMAO- no difference

Diet satisfaction (5 point scale - higher better)

Vegan 3.5 (baseline) \clubsuit to 3.0 (8 weeks)

Carnivore 3.5 1 to 3.6

From Netflix - no change in cognitive scores, telomere lengthening?

Screenshot from Netflix







Impact of the risk marker changes on CVD risk

LDL ↓ 14mg/dL ~12% and HDL ↓ 4mg/dL ~ 7%

BASELINE NUMBERS

Total cholesterol ~200 mg/dL, LDL ~120, HDL ~60, SBP 120?

Impact of vegan diet on surrogate markers

Total cholesterol ~182 mg/dL, LDL ~106, HDL~56, SBP 120?

Best estimate of what these surrogate marker changes would do to 10-year risk for developing CVD

	Baseline	Change in risk factors = Vegan		
40 y/o female	2.1%	2.0%		
40 y/o male	3.1%	3.1%		
60 y/o male	10.3%	10.4%		



Meta-analyses

Health outcomes associated with vegetarian diets: An umbrella review of systematic reviews and meta-analyses

Clin Nutr 2020;11:3283-3307

Effect of vegetarians vs ominivores on surrogate markers

SURROGATE	RESULTS Change in surrogate marker Vegetarians vs Omnivores 8 weeks				
OUTCOMES					
Total cholesterol	↓ 11%				
LDL cholesterol	↓ 14%				
HDL cholesterol	↓6% decrease				
Triglycerides	No effect				
Systolic blood pressure	↓ 4.8mmHg				
Diastolic blood pressure	↓ 2.2mmHg				
SYSTEMATIC REVIEWS	2020 Oussalah				

Studies of the Mediterranean diet show it produces minimal if any changes on surrogate markers

We already knew the answer about surrogate markers and vegetarian/vegan

Vegetarian or vegan diets and blood lipids: a meta-analysis of randomized trials

30 trials - quantified the effect of vegetarian or vegan diets vs. an omnivorous diet on lipids

Plant-based diets

↓ total cholesterol by 0.34 mmol/L (7% from baseline) - 95% CI (5% to 9%)

- LDL by 0.30 mmol/L (10%)
- ↓ apolipoprotein B levels by 12.9 mg/dL (14%)

no effect seen with triglycerides

"this study did not investigate the effect of plant-based diets on HDL-C since we focused on established atherogenic lipids and lipoproteins"

Ultra-processed food

Common examples are carbonated soft drinks, fatty or salty snacks, candies, pastries, cakes and cake mixes, margarine, sweetened cereals, fruit yogurt, pasta, pizza, poultry or fish nuggets, sausages, burgers, hot dogs, powdered or instant soup, noodles, and desserts.



A simple way to figure out if a product is ultra-processed is to see if its list of ingredients contains words such as: hydrolysed proteins, soya protein isolate, gluten, casein, whey protein, mechanically separated meat, fructose, high-fructose corn syrup, fruit juice concentrate, invert sugar, maltodextrin, dextrose, lactose, soluble or insoluble fibre, hydrogenated or interesterified oil

Ultra-processed food

NOVA Food classification

Unprocessed or minimally processed foods	Processed culinary ingredients	Processed foods	Ultra-processed foods
Foods which did not undergo processing or underwent minimal processing technics, such as fractioning, grinding, pasteurization and others.	These are obtained from minimally processed foods and used to season, cook and create culinary dishes.	These are unprocessed or minimally processed foods or culinary dishes which have been added processed culinary ingredients. They are necessarily industrialized.	These are food products derived from foods or parts of foods, being added cosmetic food additives not used in culinary.
	Salt		
Legumes, vegetables, fruits, starchy roots and tubers, grains, nuts, beef, eggs, chicken, milk	Salt, sugar, vegetable oils, butter and other fats.	Bottled vegetables or meat in salt solution, fruits in syrup or candied, bread, cheeses, purees or pastes.	Breast milk substitutes, infant formulas, cookies, ice cream, shakes, ready-to-eat meals, soft drinks and other sugary drinks, hamburgers, nuggets.

Ultra-processed food

The NOVA classification outlines 4 food categories

1.Unprocessed and minimally processed food

2. Processed culinary ingredients

3.Processed food

4.Ultra-processed food (UPF)

% of energy intake US/UK ~50 to 60% from UPF "eat the least" quintile still average 20-30% Canada and Brazil ~50% Spain and Portugal ~20% Italy ~10%

Common examples are carbonated soft drinks, fatty or salty snacks, candies, pastries, cakes and cake mixes, margarine, sweetened cereals, fruit yogurt, pasta, pizza, poultry or fish nuggets, sausages, burgers, hot dogs, powdered or instant soup, noodles, and desserts.

A simple way to figure out if a product is ultra-processed is to see if its list of ingredients contains words such as: hydrolysed proteins, soya protein isolate, gluten, casein, whey protein, mechanically separated meat, fructose, high-fructose corn syrup, fruit juice concentrate, invert sugar, maltodextrin, dextrose, lactose, soluble or insoluble fibre, hydrogenated or interesterified oil

Ultra-processed food and bad outcomes

% of energy intake US/UK ~50 to 60% from UPF Canada and Brazil ~50% Spain and Portugal ~20%										
					QUANTILE					
Italy	/ ~10%	J		1 (reference)	2	3	4	5		
		Years	Outcome	~<25%* total daily energy, or ~<2 servings/day	~25-30% ~2-3.5	~30-40% ~3.5-4.5	~40-45% ~>4.5	~>45%		
	Zhong 2021	13.5	CVD Mortality	1	No change	No change	No change	~20% 🕇		
	Blanco-Rojo 2019	7.7	Mortality	1	No change	No change	~45% 🕇			
	Schnabel 2019	7.1	Mortality	1	No change	No change	No change			
	Srour 2019	5.2	CVD	1	No change	No change	~25% 🕇			
	Kim 2019	10	Mortality CVD mortality	1	No change	No change	~30% 🕇			
		19		1	No change	No change	No change			
	Rico-Campa 2019	200,432	Mortality	1	No change	No change	~45% 🕇			
		years	CVD mortality	1	No change	No change	No change			

* numbers rounded



eClinicalMedicine Part of THE LANCET Discovery Science

Ultra-processed food consumption, cancer risk and cancer mortality: a large-scale prospective analysis within the UK Biobank

Quartiles of % UPF in diet = 9%/17%/24%/41% Looked at **209** comparisons for **25** different cancers **9** were statistically increased - all but **1** were in the **highest** quartile In the highest quartile - **All cancers** 1.07 (1.02-1.14) = **7% relative increase**

The 5 large RCTs of nutrition intervention

People with previous history of heart attacks/strokes





these numbers were reported as statistical different, everything else was not statistically different

People with NO previous history of heart attacks/strokes

these numbers were reported as statistical different, everything else was not statistically different



Absolute differences based on 2 different baseline risk estimates - per 1000 over 5 years

Intermediate risk 5-10% 5 year CVD risk

Dietary programme vminimal intervention	All cause mortality	Cardiovascular mortality	Stroke	Non-fatal myocardial infarction	Unplanned cardiovascular intervention
Mediterranean	-17 (-26 to -5)	-13 (-17 to -6)	-7 (-11 to -1)	-17 (-21 to -11)	-1 (-12 to 16)
Low fat	-9 (-15 to -3)	-6 (-11 to 1)	0 (-5 to 6)	-7 (-13 to -1)	-13 (-20 to -2)
Very low fat	-3 (-14 to 10)	0 (-10 to 14)	-1 (-7 to 9)	6 (-4 to 20)	-2 (-14 to 19)
Modified fat	3 (-12 to 22)	3 (-7 to 17)	13 (-9 to 74)	-4 (-13 to 11)	NA
Combined low fat-low sodium	1 (-11 to 15)	2 (-12 to 25)	-8 (-14 to 5)	21 (-2 to 59)	10 (-12 to 59)
Ornish	76 (-46 to 553)	13 (-22 to 179)	NA	NA	-2 (-22 to 60)
Pritikin	-48 (-61 to 207)	NA	30 (-19 to 561)	NA	NA

Superior to minimal intervention with moderate to high certainty

Little or no benefit relative to minimal intervention with moderate to high certainty

Might be superior to minimal intervention with very low to low certainty

Might have little or no benefit relative to minimal intervention with very low to low certainty

High risk 20-30% 5 year CVD risk

Dietary programme vminimal intervention	All cause mortality	Cardiovascular mortality	Stroke	Non-fatal myocardial infarction	Unplanned cardiovascular intervention
Mediterranean	-36 (-58 to -10)	-39 (-54 to -19)	-16 (-25 to -3)	-42 (-53 to -28)	-4 (-51 to 62)
Low fat	-20 (-33 to -6)	-17 (-34 to 4)	0 (-12 to 13)	-18 (-31 to -3)	-57 (-89 to -9)
Very low fat	-6 (-29 to 22)	-1 (-30 to 40)	-1 (-16 to 20)	15 (-11 to 48)	-6 (-61 to 74)
Modified fat	6 (-25 to 44)	8 (-21 to 49)	29 (-21 to 151)	-9 (-32 to 26)	NA
Combined low fat-low sodium	2 (-23 to 32)	5 (-36 to 70)	-17 (-32 to 11)	42 (-7 to 116)	41 (-54 to 199)
Ornish	145 (-104 to 659)	38 (-67 to 380)	NA	NA	-10 (-101 to 202)
Pritikin	-109 (-141 to 338)	NA	63 (-43 to 718)	NA	NA

Superior to minimal intervention with moderate to high certainty

Little or no benefit relative to minimal intervention with moderate to high certainty

Might be superior to minimal intervention with very low to low certainty

Might have little or no benefit relative to minimal intervention with very low to low certainty

Absolute differences - over 5 years

Dietary programme <i>v</i> minimal intervention (usual or no/ minimal advice)	All cause mortality		Cardiovascular mortality		Stroke		Non-fatal myocardial infarction		Unplanned cardiovascular intervention	
BASELINE RISK	INTER- MEDIATE	HIGH	INTER- MEDIATE	INTER- HIGH INTER- HIGH MEDIATE HIGH		INTER- MEDIATE	HIGH	INTER- MEDIATE	HIGH	
Mediterranean	2%↓	4%↓	1%↓	4%↓	%↓ 1%↓ 2%↓		2%↓	4%↓	No diffe	rence
Low fat (20-30%)	1%↓	2%↓	No difference No difference		1%↓	2%↓	No difference			
Very low fat (10-20%)		No difference								
Modified fat (1 in PUFA/SF)						No difference				
Combined low fat-low sodium						No difference				
Ornish (<10% fat)			No difference							
Pritikin (<10% fat)					No diffe	rence				
Low carb					NO TR	IALS				

Intermediate risk 5-10% 5 year CVD risk High risk 20-30% 5 year CVD risk

What is the answer?

Teasing out the benefits and harms of things we eat is **EXTREMELY** complicated **SINGLE NUTRIENTS**

Not enough robust data to ascribe causality

Some interesting associations - eggs, salt, coffee, alcohol

MULTIPLE NUTRIENTS AND BEHAVIOURS

Issues of RCTs and Cohorts - bias and confounding - answer may be unknowable

How to best lose weight is very individual - low carb/higher fat/protein maybe somewhat better? - is the difference important?

Overall nutrition is hugely personal and emotional



Nutrition advice to which pretty much everyone agrees

But the magnitude of the effect is "smaller than you may think"

based on the Best Available Evidence

- 1. Eat a greater percentage of whole foods (food that has not been overly processed or refined as little as possible)
- 2. Eat more vegetables
- 3. Eat less added sugar
- 4. Eat more whole grains
- 5. Eat in a style that fits your food preferences, tolerances, and lifestyle
- 6. Eat in a style you can sustain
- 7. When it comes to weight, how much you eat is really the KEY issue
- 8. The "best" weight is the weight you are when living the healthiest life you can enjoy
- 9. Avoid any food that has, for you, been properly shown to cause unacceptable intolerances

BUT THERE ARE BIG CAVEATS

Almost all the nutrition "benefits and harms" evidence comes from cohort studies

- real possibility of important publication bias because 100s to 1000s of researchers are looking at 100s of different databases
- many potential confounders let alone data collection issues
- many associations seen in cohort studies are quite small (<10% relative) and only seen when you compare "lots quantiles" to "not much at all quantiles"
- in general single cohorts unless that is all you have should not be used as solid evidence

Much of nutrition research is on surrogate markers (blood pressure, lipids, glucose)

- the changes seen **IF** they translated into effects on clinical outcomes would only amount to a 1% (at most 2%) absolute change in CVD risk over 10 years
- in general single RCTs of surrogates should not be considered high quality evidence

There are only 5 large RCTs (2+years) that have looked at important clinical outcomes

the "best evidence" is for the "Mediterranean Diet" and even that only showed a 1-2% absolute ↓ in stroke over 5 years - but a bigger decrease (↓3-8%) if secondary prevention

THESE ARE ACTUALLY PRETTY REASONABLE CONSIDERING THE EVIDENCE





CANADA

Anything else is likely

assuming you wish your eating to be informed by the best available evidence





















1. ENJOY EATING

- Differences in outcomes are typically found from "extremes" and are "small"
- 3. The **Mediterranean diet** (whatever it is) seems reasonable also CFG/ USDA/DASH
- 4. Eat in moderation/moderation/moderation
- 5. Avoid eating "lots" of ultra processed food
- 6. You can easily justify some red meat, butter etc
- 7. Eggs, coffee, salt, and alcohol in moderation seem fine
- 8. Saturated fats OK trans-fat?
- 9. Added sugars (beverages mainly) at the high end seem to increase risk of obesity
- 10.It is **VERY unlikely** a single "nutrient" would have an important effect
- 11.Animal rights/environmental issues are a whole other topic









IF THERE IS CAUSATION, WHAT DO ALL THESE NUMBERS MEAN

BALLPARK

2 drinks a day ~15% relative increase for oral/GI/breast cancers **"Three slices of bacon" a day** ~15% relative increase for oral/GI cancers

Lifetime risks of cancers, for the two foods associated with an increased risk of cancer

ORAL/GI CANCERS Male/Female - 6% BASELINE RISK

~15% relative increase = ~1% absolute increase over a lifetime **BREAST**

Female - 13% BASELINE RISK

~15% relative increase = ~2% absolute increase over a lifetime