I applaud the #EUCancerPlan BUT beware: putting #meat (a nourishing, evolutionary food) in the same box as to solve a contemporary health challenge, would be basing policy on assumptions rather than robust data. #FollowTheScience, yes, but not just part of it!

THREAD •

1/Granted, some studies have pointed to ASSOCIATIONS of HIGH intake of red & processed meats with (slightly!) increased colorectal cancer incidence. Also, @WHO/IARC is often mentioned in support (usually hyperbolically so). But, let's have a closer look at all this!

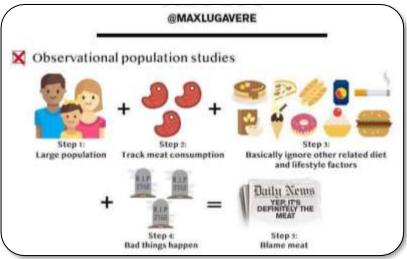
Bacon, burgers and sausages are a cancer risk, say world health chiefs: Processed meats added to list of substances most likely to cause disease alongside cigarettes and asbestos

- · Fresh red meat is also due to join WHO 'encyclopaedia of carcinogens'
- · Rulings will send shock waves through farming and fast food industries
- · Could lead to new dietary guidelines and warning labels on bacon packs
- Mounting concern that meat fuels disease that kills 150,000 a year in UK

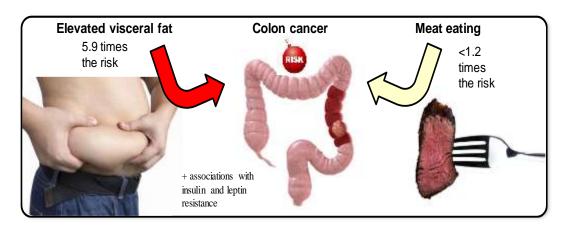
2/First, meat being "associated" with cancer is very different from stating that meat CAUSES cancer. Unwarranted use of causal language is widespread in nutritional sciences, posing a systemic problem & undermining credibility. https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3280017/

3/That's because observational data are CONFOUNDED (even after statistical adjustment). Healthy user bias, eg. is a major problem. Healthy middle classes are TOLD to eat less red meat (due to historical rather than rational reasons, cf link). So, they obey. https://iastatedigitalpress.com/mmb/article/id/9456/

4/ What's captured here is sociology, not physiology. Health-focussed Westerners eat less red meat; those who don't adhere to dietary advice tend to have unhealthier lifestyles. That tells us very little about meat AS SUCH being responsible for disease.



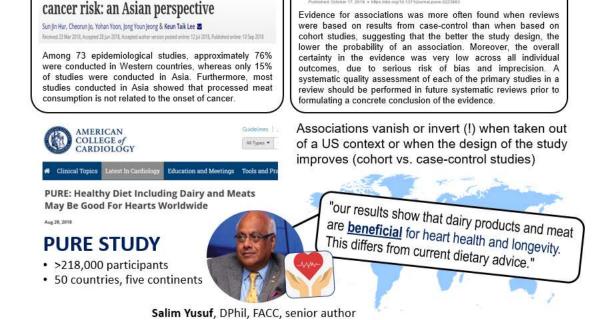
5/At very small relative risks (<<x2), we just CANNOT formulate strong conclusions. Example: someone with elevated visceral fat needs to be worried (6x risk of colon cancer!) For meat, however, risk level is so small (close to x1), that we're out of business.



Controversy on the correlation of red and processed meat consumption with colorectal

6/Worse: the associations are likely mere artifacts. Why? When we look at studies with better design or move out of a US context (e.g. Asia or worldwide), MORE meat is associated with BETTER health!? Indicative of a cultural construct rather than a paradox.

Processed meat intake and chronic disease morbidity and mortality: An overview of systematic reviews and metaanalyses

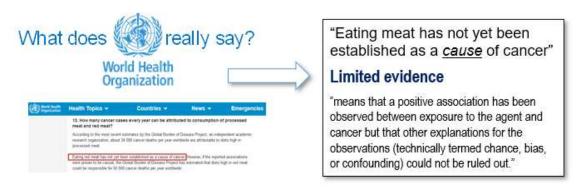


7/To be fair, researchers are usually more nuanced than policy makers. As stated in this highly cited study on meat & mortality, data 'should be interpreted with caution due to the high heterogeneity observed [&] the possibility of residual confounding'

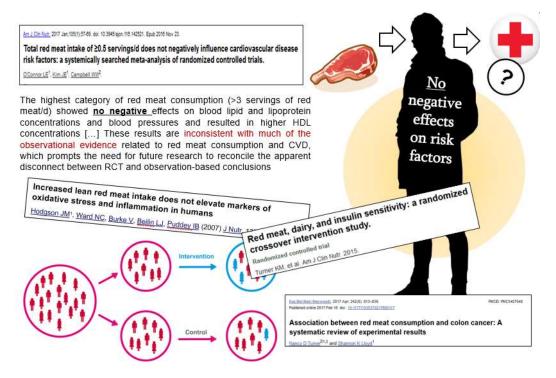
https://www.cambridge.org/core/journals/british-journal-of-nutrition/article/association-between-total-processed-red-and-white-meat-consumption-and-allcause-cvd-and-ihd-mortality-ametaanalysis-of-cohort-studies/35CB32B716F2FBAF6119070029193544

8/Even the WHO/IARC panel looking into the colorectal cancer link declared that 'other explanations for the observations (chance, bias or confounding) could not be ruled out' while 'consumption of red meat has not been established as a cause of cancer' <a href="https://www.who.int/news-room/q-a-detail/cancer-carcinogenicity-of-the-consumption-of-red-carcinogenicity-of-the-consumption-of-red-carcinogenicity-of-the-consumption-of-red-carcinogenicity-of-the-

meat-and-processed-meat



9/Observational data yielding associations between meat intake and disease thus need to be CAREFULLY inspected. At best, this creates a HYPOTHESIS that needs to be validated in intervention studies. But such studies fail to indicate harm!



10/It is true of course that such trials are difficult on long term in humans & rely on biomarkers that are imperfect. Alternatively, one can use animal models or cell cultures. Once more: INSUFFICIENT evidence (not to mention the extrapolation concerns)



Forty studies using animal models or cell cultures met specified inclusion criteria, most of which were designed to examine the role of heme iron or heterocyclic amines in relation to colon carcinogenesis. Most studies used levels of meat or meat components well in excess of those found in human diets. Although many of the experiments used semi-purified diets designed to mimic the nutrient loads in current westernized diets, most did not include potential biologically active protective compounds present in whole foods. Because of these limitations in the existing literature, there is currently insufficient evidence to confirm a mechanistic link between the intake of red meat as part of a healthy dietary pattern and colorectal cancer risk.

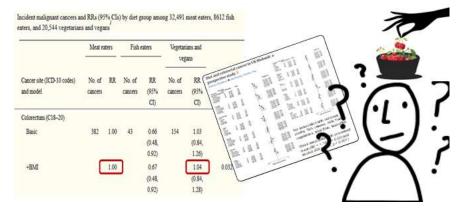


11/Another problem: CHERRY PICKING. Although associated with colorectal cancer, why not mention that meat shows a PROTECTIVE association with melanoma? Or that vegetarians in the UK are WORSE off? Etc.

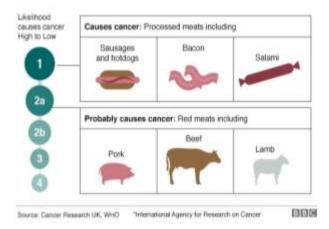


Inconvenient facts

- Red and processed meat intake is associated with colorectal cancer but inversely associated with melanoma (Cross et al. 2007; Yen et al. 2018)
- With respect to colorectal cancer most studies were from 1990s, more up
 to date info from the UK showed no significant association with red meat
 and only a weak one with processed meats (Bradbury et al. 2020)
- (British) vegetarians are not better off than meat eaters: higher incidence of colorectal cancer (Key et al. 2014); mortality from circulatory diseases and all causes is not significantly different (Key et al. 2009)



12/Bringing us to the WHO/IARC assignment of red meat to Group 2A ("probably carcinogenic to humans"). Why did they do this and what does it mean?



13/It's good to have in mind that this is more controversial than it seems. One of the WHO/IARC's own experts, dr. Klurfeld, has severely criticized this. For an overview of his objections, cf:

https://watermark.silverchair.com/vfy009.pdf?token=AQECAHi208BE49Ooan9kkhW_Ercy7Dm3ZL_9Cf3qfKAc485ysgAAApQwg gKQBgkqhkiG9w0BBwagggKBMIICfQIBADCCAnYGCSqGSlb3DQEHATAeBglghkgBZQMEAS4wEQQMsPC9byK57MWgJy8cA gEQgIICR2xgrM7PTPT91bPYAjPsp6SAN9QfbXkSLtn4aa_RusxpW941rZiPQFLCZ0dOSq1D1a1il26gy5SbubQFokQBXpN9Zt8D utBsgW8ttrj2mqqpXoji00ozyAntgzjrdNJ6x298Tpe1sdvzEEdPJi2bBJV3Pn9Ncy6dvEdC1edIOjx_QQnDnSvHlixvmRNlhRABEijaP n4RYn5Hl8Ini1PdBdT4oqVFCWeWUbzthRws2VMTbzI_3yoXSAuTIUSJMY4S6XDI7t_CQqr7vXXtxOYOESteiVZJ0zzMwhhk75U SSgnkkAWrSOSUs20vH-V3t5ZltUU1zN3647xU8VnUMiyoVOosp-

tcFC4g_NK0IVw_V2LRmetPxQlw5B9V0nvrBKDUsi2JbEF0Sa41qQoRbaMfGF2OICvwG5q0vmHy8oaC4N1VLXppc0gmscQd6VRVb7qIJ3mJ1OjF9v9FZ5-UVix-iwDnBRzT8mA87C0rWhrL766AX_SKRhB9J8tP4hdCdJ3vd82KI5HdS0Ck8ELsrtv-SV_UR9xaCahxVnWTAhvO4QRCZflU6X0I_jiQY2JM41IE4ndrl_VkE65suAJn9FwAN06neeQoi3emDgZcJXf0L71HzV0PEbWoEQ7avgLRYTbr9n4DbnT3WcqK-S-pF-zCB4TIXqlCnve7Wi5vHhKATNsxsEH4ZOrzfbWjc7jpaI-

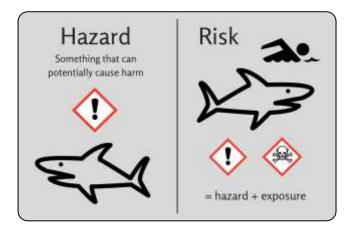
IXw6XujgRwteoF A05QNJGYFc9gq6SWHep8-Z wRamL9d5ZER1IUUnmsRBgumMIF8W1PVYj8HRDM

What is the role of meat in a healthy diet?

David M. Klurfeld

USDA Agricultural Research Service, Beltsville, MD 20705

14/Furthermore, we need to understand that WHO/IARC classifications indicate HAZARDS, not risks. To go from risk to hazard, we need... a risk assessment.



15/Such risk assessment indicates that there is no solid case for concern, especially in the context of a normal diet.

Food Chem Toxicol, 2018 Apr 21. pit: S0278-6915(18)30265-5. doi: 10.1016/j.fct.2018.04.048. [Epub ahead of print]

Red meat and colon cancer: A review of mechanistic evidence for heme in the context of risk assessment methodology.

Kruger C1, Zhou Y2.

In conclusion, the methodologies employed in current studies of heme have not provided sufficient that the mechanisms studied would contribute to an increased risk of promotion of preneoplasia or cusual dietary intakes of red meat in the context of a normal diet.

16/ Because, indeed, CONTEXT is everything. Sunlight is a #hazard (more so than red meat, which is at the level of being a hairdresser) & #risk under certain conditions. But it is fair to say that sunlight is mostly beneficial (vitamin D being just one reason)

Context is everything

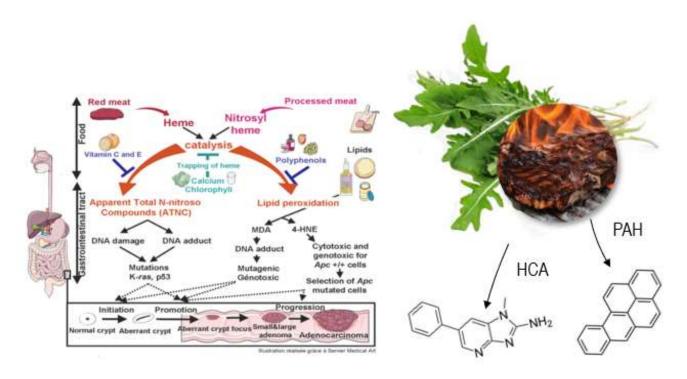




2A: Working as barber or hairdresser

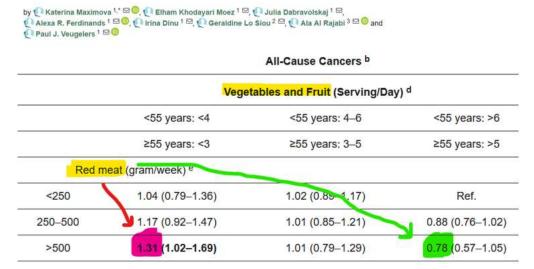


17/Obviously, one shouldn't consume all-too heavily processed meats, or overly charred steaks, all-too often. Or blame the beef patty for the ultraprocessed bun, sauces, fries, & soda consumed with it.



18/So within an overall HEALTHY DIET, whatever potential cancer risk (if any; hard to tell due to confounding & bias) can reasonably be expected to be irrelevant. In the study below, more meat either parallels higher (veg-) or lower (!!) (veg+) risk.

Co-consumption of Vegetables and Fruit, Whole Grains, and Fiber Reduces the Cancer Risk of Red and Processed Meat in a Large Prospective Cohort of Adults from Alberta's Tomorrow Project



https://www.mdpi.com/2072-6643/12/8/2265

19/Some authors are therefore starting to question the usefulness of IARC-type schemes to begin with. As they lead to scaremongering and loss of benefits (meat = valuable nutrition, etc.)

Regul Toxicol Pharmacol. 2016 Dec;82:158-168. doi: 10.1016/j.yrtph.2016.10.014. Epub 2016 Oct 22.

Classification schemes for carcinogenicity based on hazard-identification have become outmoded and serve neither science nor society.

Boobis AR¹, Cohen SM², Dellarco VL³, Doe JE⁴, Fenner-Crisp PA⁵, Moretto A⁶, Pastoor TP⁷, Schoeny RS⁸, Seed JG⁹, Wolf DC¹⁰.

This!

[...] Because a risk-based decision framework fully considers hazard in the context of dose, potency, and exposure the unintended downsides of a hazard only approach are avoided, e.g., health scares, unnecessary economic costs, loss of beneficial products, adoption of strategies with greater health costs, and the diversion of public funds into unnecessary research.

20/No wonder that some top-level scientists, eg. Gordon Guyatt (leading expert in the field of evidence-based medicine), have criticized the WHO/IARC after the release of its report for "doing the public a disservice". They'd likely say the same for the #EUCancerPlan.

Mistaken advice on red meat and cancer

Gordon Guyatt, Distinguished Prof., Dept. of Clinical Epidemiology & Biostatistics, Joint Member, Department of Medicine, McMaster University, Ontario; Benjamin Djulbegovic, Associate Dean, College of Internal Medicine; Professor, Dept. of Oncological Sciences; Distinguished Professor, Clinical and Translational Institute, Director, Evidence-based Medicine Research Group, University of South Florida.

Epidemiology is a science that can establish associations [...] but seldom cause and effect [The] success story of epidemiological science was its ability to link smoking to cancer, with WHO designating tobacco as a "convincing" carcinogen in 1986 [...] Ever since, however, standards for these risks in policy making have been dropping. This week's decisions on meat were based on relative risks of 1.17 to 1.18, a tiny fraction of those for smoking. To keep things in perspective: for colon cancer, which was the focus of the WHO report, the absolute risk of contracting this cancer in one's lifetime is less than 4.5%. An increased relative risk of 1.17 raises the absolute risk to no more than 5.3%. As two of the leaders in evidence based medicine, we were involved in the development an evidence ranking system, called "GRADE," adopted by over 90 groups world-wide, including the WHO. GRADE notes that unless relative risks are greater than 5, epidemiological studies typically provide only low-quality evidence [...] In such cases, the evidence is not convincing, and any recommendations would ordinarily be we would make a "weak", or "optional" recommendation, since the benefits do not clear outweigh the potential harms. The WHO has done the public a disservice in abandoning GRADE in its evaluation of the evidence, and greatly overstating confidence in a causal connection between red meat and cancer. Recent decades are littered with policies based on weak relative risks which, when properly tested in clinical trials, had to be reversed [...] We see the same story with dietary guidelines: recommendations to restrict dietary cholesterol and limit fat to fight cancer were originally based principally on epidemiological 🚧 🗃 hat clinical trials failed to confirm [...] The reason that weak associations are untrustworthy is that they could associated with any number of factors in diet or lifestyle [...] Vegetarians tend to be more alert to good health: very well be ue to they smol ess, exerose more, and have a higher socioeconomic status. By contrast, meat-eaters over the past 30-plus years are nore their doctor's orders and are likely to be engaging in other insalubrious behaviors, all of which alone or in ght explain the small relative risks associated with meat-eating. Bias against red meat is another factor, easily scientific literature and the popular press [...] Small relative risks are therefore just as likely to reflect bias as any omized clinical trials provide far more trustworthy evidence regarding cause and effect. It is therefore perplexing that O document does not even mention the relevant data: two large, multi-year RCTs, both funded by the National Institute of Health [Polyp Prevention Trial and WHI] To say that red or processed meat is equivalent to smoking is profoundly misleading.

21/Last year, Guyatt & others formalized their critique by COMPREHENSIVELY looking at the evidence. Based on sufficiently high standards of evidence, they concluded that advice to reduce red & processed meat is based on weak evidence. https://www.acpjournals.org/doi/10.7326/M19-1621

22/In the editorial of the journal, Carroll & Doherty argued that those who seek to dispute this [assessment] will be hard pressed to find appropriate evidence with which to build an argument"

EDITORIALS | 1 OCTOBER 2019

Meat Consumption and Health: Food for Thought

Aaron E. Carroll, MD, MS; Tiffany S. Doherty, PhI

Annals of Internal Medicine

A fifth article this month is a new guideline [...] based on these reviews. It was voted on by 14 members, including 3 community members, from 7 countries and had strict criteria concerning conflicts of interest. The overall recommendations, contrary to almost all others that exist, suggested that adults continue to eat their current levels of red and processed meat, unless they felt inclined to change them themselves. This is sure to be controversial, but it is based on the most comprehensive review of the evidence to date. Because that review is inclusive, those who seek to dispute it will be hard pressed to find appropriate evidence with which to build an argument [...] Moreover, it may be time to stop producing observational research in this area. These meta-analyses include millions of participants. Further research involving much smaller cohorts has limited value. High-quality randomized controlled trials are welcome, but only if they're designed to tell us things we don't already know.

Johnston et al (2019) recommend "to continue rather than reduce consumption of unprocessed red meat or processed meat" Unprocessed Red Meat and Processed Meat Consumption: Dietary Guideline Recommendations From the Nutritional Recommendations (NutriRECS) Consortium

Bradley C, Privators, PAD, Domo Erranilar, MoC, AM A Han, PAD, Fadris NEAL Verwood), eME, Claudels Vall, MSC, Segine EZ DB, PAD, Carberine MARSHAN, PATERS, Elmow, PAD, Source Intervention—THUE, PAD, Critopose Philips, Ping Ti, Ind. Brainer, Padag Santial of Source, ACC Carbon Services, AMI, PAD, Padris Advances, Carbon Services, Padagonia, Edulatory, Source, Carbon Services, AMI, PAD, Condess, ALD, PAD, Co

Causal inference assessment based on summary of evidence

23/What followed was indeed incoherent rebuttal by anti-meat groups arguing that we should accept lower standards of evidence for nutrition, because it can't do better (?!) Plus a vitriolic smear campaign. MUST READ! https://www.tamus.edu/wp-content/uploads/2020/01/JAMA-Article-1.15.20.pdf

Backlash Over Meat Dietary Recommendations Raises Questions About Corporate Ties to Nutrition Scientists

Rita Rubin, MA

It's almost unheard of for medical journals to get blowback for studies before the data are published. But that's what happened to the Annois of Internol Medicine last fall as editors were about to post several studies showing that the evidence liming red meat consumption with cardiovascular disease and cancer is too weak to recommend that adults eat less of it.

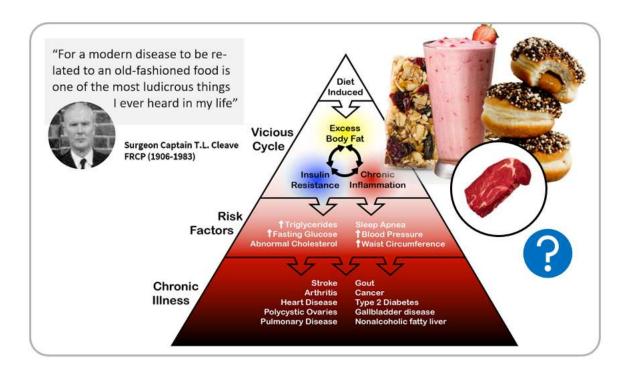
Annols Editor-in-Chief Christine Laine, MD, MPH, saw har inbox flooded with roughly 2000 emails—most bore the same missage, apparently generated by a bot—in a half hour. Laine's inbox had to be shut down, she said. Not only was the volume unprecedented in her decade at the helm of the respected journal, the tone of the emails was particularly caustic.

"We've published a lot on firearm injury prevention," Laine said. "The response from the NRA (National Rifle Association) was less vitrioi; than the response from the Trus Health Initiative."



24/Rather than ideological a "priori", let's return to common sense: "for a modern disease to be related to an old-fashioned food is one of the most ludicrous things I ever heard in my life".

Let's focus on ultraprocessed junk instead?



25/I leave it with the following statement: "we argue that claims about the health dangers of red meat are not only improbable in the light of our evolutionary history, they are far from being supported by robust scientific evidence"

https://www.tandfonline.com/doi/full/10.1080/10408398.2019.1657063

26/...and our @aleph2020 website (brought by a consortium of 35+ scientists). I invite you to visit the Health section where we not only argue that there's no good reason to avoid meat, but also that it may lead to the loss of valuable nutrition: https://aleph-2020.blogspot.com/p/introduction.html

27/Meat, indeed, is an evolutionary food. It made us human. We're *adapted* to it. It would be highly improbable that it harms us to such an extent that we would have to include its restriction in a #EUCancerPlan. https://aleph-2020.blogspot.com/2020/04/the-role-of-asfs-in-historical-diets.html

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