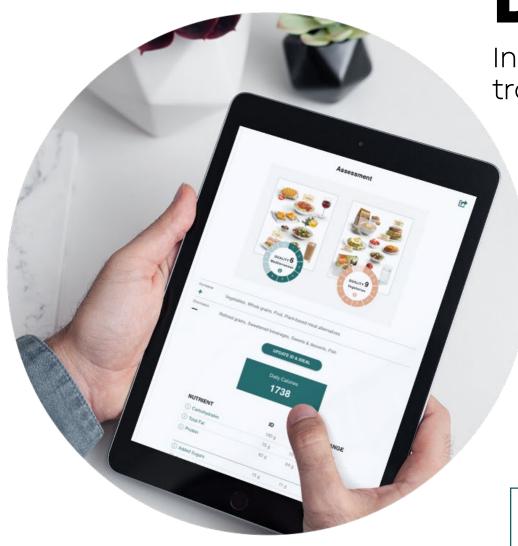
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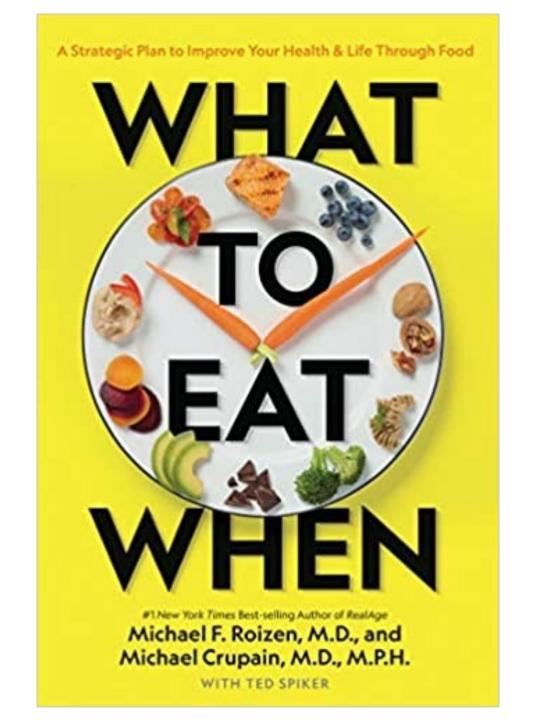
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Intermittent Fasting... And Beyond!

Of Eating, Not Eating, Timing, & Metabolism

David L. Katz, MD, MPH CEO, Diet ID, President, True Health Initiative

Michael Crupain, MD, MPH Chief Medical Officer, Sharecare Medical Unit Chief of Staff, "The Dr. Oz Show"

Michael Roizen, MD Chief Wellness Officer, Cleveland Clinic Developer of the RealAge Concept



March 24, 2020

Everyone has an opinion

HAMILTON CLEM



What does the science say?

Effects of Intermittent Fasting on Health, Aging, and Disease.

de Cabo R. Mattson MP.

N Engl J Med. 2019 Dec 26;381(26):2541-2551. doi: 10.1056/NEJMra1905136.

Metabolic Effects of Intermittent Fasting.

Patterson RE, Sears DD,

Annu Rev Nutr. 2017 Aug 21;37:371-393. doi: 10.1146/annurev-nutr-071816-064634. Epub 2017 Jul 17.

PMID: 28715993

Effects of intermittent fasting on glucose and lipid metabolism.

The objective of this review is to provide an ov Antoni R, Johnston KL, Collins AL, Robertson MD.

intermittent fasting might lead to improved

PubMed and the terms "intermittent fasting

evidence on the health benefits of intermittel Proc Nutr Soc. 2017 Aug;76(3):361-368. doi: 10.1017/S0029665116002986. Epub 2017 Jan 16.

PMID: 28091348

Two intermittent fasting variants, intermittent energy restriction (IER) and time-restricted feeding

(TRF), have received considerable interest as strategies for weight-management and/or improvi Impact of intermittent fasting on the lipid profile: Assessment associated with metabolic health. ...Ultimately, much remains to be learned about intermittent fasting (in its value and weight loss.

forms); however, the findings to date serve to highlight promising avenues for future research.

Impact of **intermittent fasting** on health and disease processes.

Mattson MP, Longo VD, Harvie M.

Ageing Res Rev. 2017 Oct;39:46-58. doi: 10.1016/j.arr.2016.10.005. Epub 2016 Oct 31.

PMID: 27810402 Free PMC article.

(e.g., 16-48h) with little or no energy intake, with intervening period and type-2 diabetes. recurring basis. We use the term periodic fasting (PF) to refer to IF

mimicking diets lasting from 2 to as many as 21 or more days. ...

Intermittent fasting (IF) encompasses eating patterns in which ind The role of low-calorie diets and intermittent fasting in the treatment of obesity ks large sample and detailed information about diet.

Zubrzycki A, Cierpka-Kmiec K, Kmiec Z, Wronska A.

J Physiol Pharmacol. 2018 Oct;69(5). doi: 10.26402/jpp.2018.5.02. Epub 2019 Jan 21.

Intermittent fasting (IF) involves caloric restriction for one or several days a week, or every day as the prolongation of the overnight fast. The results of recent clinical trials have shown that LCDs and intermittent fasting in patients with obesity (including those with coexisting T2D) can lead to a reduction in body fat mass and metabolic parameter improvements. ...

Effects of **intermittent fasting** on body composition and clinical health markers in humans.

Tinsley GM, La Bounty PM.

Nutr Rev. 2015 Oct;73(10):661-74. doi: 10.1093/nutrit/nuv041. Epub 2015 Sep 15.

PMID: 26374764

Intermittent fasting protocols can be grouped into alternate-day fasting, whole-day fasting, and time-

restricted feeding. Alternate-day fasting trials of 3 to 12 weeks in duration appear to be effective at

weight ($\approx 3\%$ -7%), body fat (≈ 3 -5.5 kg), total cholesterol ($\approx 10\%$ -21%), and triglycerides normal-weight, overweight, and obese humans. ... Future studies should examine long-

intermittent fasting and the potential synergistic effects of combining intermittent

cercise....

Santos HO, Macedo RCO.

Clin Nutr ESPEN. 2018 Apr;24:14-21. doi: 10.1016/j.clnesp.2018.01.002.

PMID: 29576352

Intermittent fasting, whose proposed benefits include the improvement of lipid profile and the body weight loss, has gained considerable scientific and popular repercussion. ... However, the majority of studies that analyze the intermittent fasting impacts on the lipid profile and body weight loss are



Research

- Human RCTs
- Observational (e.g. Ramadan)
- Animal
- Meta-analyses
- Reviews



REVIEW ARTICLE

Dan L. Longo, M.D., Editor

Effects of Intermittent Fasting on Health, Aging, and Disease

Rafael de Cabo, Ph.D., and Mark P. Mattson, Ph.D.

CCOR DING TO WEINDRUCH AND SOHAL IN A 1997 ARTICLE IN THE JOURNAL. reducing food availability over a lifetime (caloric restriction) has remarkable effects on aging and the life span in animals. The authors proposed that the health benefits of caloric restriction result from a passive reduction in the production of damaging oxygen free radicals. At the time, it was not generally recognized that because rodents on caloric restriction typically consume their entire daily food allotment within a few hours after its provision, they have a daily fasting period of up to 20 hours, during which ketogenesis occurs. Since then, hundreds of studies in animals and scores of clinical studies of controlled intermittent fasting regimens have been conducted in which metabolic switching from liver-derived glucose to adipose cell-derived ketones occurs daily or several days each week. Although the magnitude of the effect of intermittent fasting on life-span extension is variable (influenced by sex, diet, and genetic factors), studies in mice and nonhuman primates show consistent effects of caloric restriction on the health span (see the studies listed in Section S3 in the Supplementary Appendix, available with the full text of this article at NEJM.org).

Studies in animals and humans have shown that many of the health benefits of intermittent fasting are not simply the result of reduced free-radical production or weight loss. 25 Instead, intermittent fasting elicits evolutionarily conserved, adaptive cellular responses that are integrated between and within organs in a manner that improves glucose regulation, increases stress resistance, and suppresses inflammation. During fasting, cells activate pathways that enhance intrinsic defenses against oxidative and metabolic stress and those that remove or repair damaged molecules (Fig. 1).5 During the feeding period, cells engage in tissue-specific processes of growth and plasticity. However, most people consume three meals a day plus snacks, so intermittent fasting does not occur. 26

Preclinical studies consistently show the robust disease-modifying efficacy of intermittent fasting in animal models on a wide range of chronic disorders, including obesity, diabetes, cardiovascular disease, cancers, and neurodegenerative brain diseases. Periodic flipping of the metabolic switch not only provides the ketones that are necessary to fuel cells during the fasting period but also elicits highly orchestrated systemic and cellular responses that carry over into the fed state to bolster mental and physical performance, as well as disease resistance. 11-12

Here, we review studies in animals and humans that have shown how intermittent fasting affects general health indicators and slows or reverses aging and disease processes. First, we describe the most commonly studied intermittentfasting regimens and the metabolic and cellular responses to intermittent fasting. We then present and discuss findings from preclinical studies and more recent clinical studies that tested intermittent-fasting regimens in healthy persons and in

From the Translational Gerontology Branch (R.C.) and the Laboratory of Neurosciences (M.P.M.), Intramural Research Program, National Institute on Aging, National Institutes of Health, and the Department of Neuroscience, Johns Hopkins University School of Medicine (M.P.M.) — both in Baltimore. Address reprint requests to Dr. Mattson at the Department of Neuroscience, Johns Hopkins University School of Medicine, 725 N. Wolfe St., Baltimore, MD 21205, or at mmattso2@jhmi.edu.

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N Engl J Med 2019;381:2541-51.
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It remains to be determined whether people can maintain intermittent fasting for years and potentially accrue the benefits seen in animal models. Furthermore, clinical studies have focused mainly on overweight young and middleage adults, and we cannot generalize to other age groups the benefits and safety of intermittent fasting that have been observed in these studies.



Types of IF

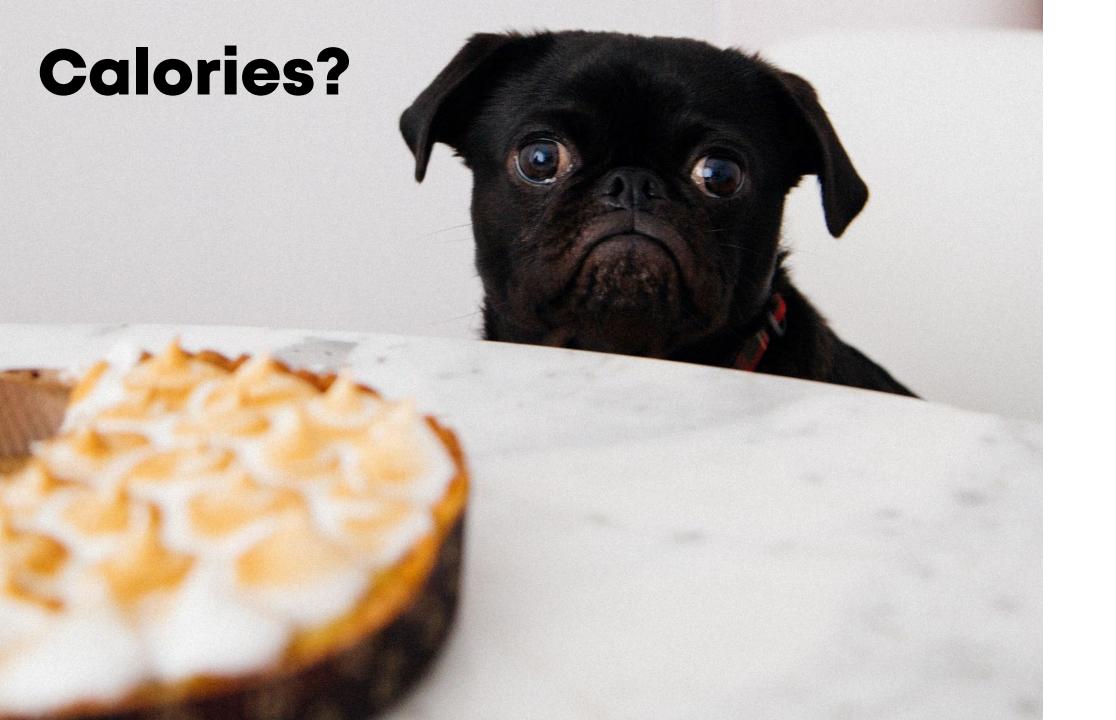
- The 16/8 method
 - fasting for 16 hours daily, keeping the eating window to 8 hours daily
- The 5/2 method
 - eat normally for 5 days, fasting for 2 (or, restrict to 500-600 calories)
- Eat-Stop-Eat
 - do a 24 hour fast 1-2 times per week
- Alternate day fasting: fasting every other day



Strategic Timing

- Metabolic advantage?
- Calorie management?
- Both?











QUESTION & ANSWER



Please submit questions via Q/A zoom feature



Thank you for attending!



Next FOOD TRUTHS Webinar is March 20th. Make sure you're on our email list!