

Exploring the Impact of Intermittent Fasting on Metabolic Health in Adults

**Rawan Mahmoud Fallata
Safa MohammedSaeed Nassar
Majed Ahmed Almuabbiri
Ranya Hasan Almalki
Abeer Abdulrahman Basamih
Malak yousef ali
Asim Gazi Allehybey
Abeer Abdulhakem T. AlShahi
Fahad Dulihan Zayed AlDulihan
Doaa Mohammad Filmban
asmaa ali sayis
Amani Akram Aldeen**

Abstract

The 16:8 technique, the 5:2 diet, and alternate-day fasting are all examples of popular protocols that fall under the classification of intermittent fasting (IF), which is a regulated dietary plan that involves alternating periods of fasting and eating. Within the scope of this study, the physiological, metabolic, and psychological effects of intermittent fasting (IF) are investigated. Particular attention is paid to the role that IF plays in enhancing metabolic health, cognitive function, and lowering the risk of chronic diseases such as type 2 diabetes, cardiovascular problems, and neurodegenerative disorders. Additionally, intermittent fasting (IF) helps to manage blood glucose levels, lowers oxidative stress, and promotes intestinal health, all of which contribute to an overall improvement in well-being. In spite of this, intermittent fasting (IF) is not uniformly helpful and there are hazards associated with it, particularly for people who have type 1 diabetes, eating disorders, immune systems that are weakened, or nutritional deficiencies. The efficacy of this treatment in certain circumstances, such as cancer therapy, is also being called into doubt by novel research, which also raises concerns about decreased bone density. This analysis highlights the importance of tailored fasting regimes that are overseen by healthcare professionals in order to reap the greatest possible benefits while also minimizing any potential hazards.

Key words

Intermittent fasting, metabolic health, glycemic control, obesity, cardiovascular health, chronic disease, caloric restriction, eating disorders, immune system.

تعد تقنية 16:8، والنظام الغذائي 5:2، والصيام المتقطع، كلها أمثلة على البروتوكولات الشائعة التي تندرج تحت تصنيف الصيام المتقطع (IF)، وهو خطة غذائية منظمة تتضمن فترات متناوبة من الصيام والأكل. في نطاق هذه الدراسة، يتم التحقيق في التأثيرات الفسيولوجية والأبضية والنفسية للصيام المتقطع (IF). يتم إيلاء اهتمام خاص للدور الذي يلعبه الصيام المتقطع في تعزيز الصحة الأبضية والوظيفة الإدراكية وخفض خطر الإصابة بالأمراض المزمنة مثل مرض السكري من النوع 2 ومشاكل القلب والأوعية الدموية والاضطرابات العصبية التنكسية. بالإضافة إلى ذلك، يساعد الصيام المتقطع (IF) في إدارة مستويات الجلوكوز في الدم، وخفض الإجهاد التأكسدي، وتعزيز صحة الأمعاء، وكل ذلك يساهم في تحسن عام في الرفاهية. على الرغم من ذلك، فإن الصيام المتقطع (IF) ليس مفيداً بشكل موحد وهناك مخاطر مرتبطة به، خاصة للأشخاص الذين يعانون من مرض السكري من النوع 1، واضطرابات الأكل، وضعف الجهاز المناعي، أو نقص التغذية. كما أن فعالية هذا العلاج في ظروف معينة، مثل علاج السرطان، موضع شك بسبب أبحاث جديدة، مما يثير أيضاً مخاوف بشأن انخفاض كثافة العظام. ويسلط هذا التحليل الضوء على أهمية أنظمة الصيام المصممة خصيصاً والتي يشرف عليها متخصصون في الرعاية الصحية من أجل جني أكبر قدر ممكن من الفوائد مع تقليل أي مخاطر محتملة.

الكلمات المفتاحية:

الصيام المتقطع، الصحة الأبضية، التحكم في نسبة السكر في الدم، السمنة، صحة القلب والأوعية الدموية، الأمراض المزمنة، تقييد السعرات الحرارية، اضطرابات الأكل، الجهاز المناعي.

Study Definitions

Intermittent Fasting (IF):

A dietary strategy involving alternating periods of fasting and eating. Common protocols include the 16/8 method (16 hours of fasting and an 8-hour eating window), the 5:2 diet (eating normally for five days and significantly reducing caloric intake for two non-consecutive days), alternate-day fasting, and the 12/12 approach (12 hours of fasting and 12 hours of eating).

Caloric Restriction (CR):

A dietary approach that involves reducing overall caloric intake without specific fasting intervals. Unlike intermittent fasting, CR focuses on continuous calorie reduction over time.

Glycemic Control:

The regulation of blood sugar levels, often achieved through dietary modifications or medical interventions, to maintain a healthy balance and prevent hyperglycemia or hypoglycemia.

Metabolic Efficiency:

The body's ability to optimize energy production and utilization, influenced by factors such as diet, exercise, and hormonal regulation. Intermittent fasting has been shown to enhance this process by improving insulin sensitivity and reducing oxidative stress.

Chronic Diseases:

Long-term health conditions that develop over time and often result from lifestyle, genetic, and environmental factors. Examples include type 2 diabetes, cardiovascular diseases, neurodegenerative disorders, and certain cancers.

Oxidative Stress:

An imbalance between free radicals and antioxidants in the body, which can lead to cell and tissue damage. Reduced oxidative stress is a noted benefit of intermittent fasting.

Bone Mineral Density (BMD):

A measure of the concentration of minerals, such as calcium, in bones, reflecting their strength and density. Low BMD can result from prolonged caloric restriction or inadequate nutrition, increasing the risk of fractures.

Gut Microbiota:

The community of microorganisms residing in the digestive tract, playing a crucial role in digestion, immunity, and overall health. Intermittent fasting can positively influence gut health by supporting a balanced microbiome.

Introduction

Intermittent fasting is a dietary regimen that limits food consumption to predefined intervals. Research indicates that intermittent fasting might effectively facilitate weight loss by decreasing total caloric consumption and enhancing metabolic rate. It is advisable to consult a healthcare expert prior to initiating any new diet or exercise program. Intermittent fasting (IF) is a widely recognized weight loss strategy noted for its straightforwardness and efficacy. In contrast to conventional calorie-restricted diets, intermittent fasting limits the temporal window for meal consumption. Recent studies indicate that intermittent fasting (IF) may serve as an effective and sustainable method for enhancing weight loss, preserving lean body mass, and improving metabolic health indicators, including blood pressure, blood glucose levels, and cholesterol. Patients must see a healthcare physician prior to initiating a new food program to assess any existing medical conditions or concurrent medications. This article will aid nurse practitioners (NPs) in comprehending the potential hazards and benefits of intermittent fasting (IF) and in identifying suitable candidates for this regimen (Trepanowski and Bloomer, 2018).

The United States faces a concerning increase in obesity, which has evolved into a national pandemic, significantly impacting the health and welfare of millions of Americans. An immediate necessity exists for thorough initiatives to tackle this intricate problem. The Centers for Disease Control and Prevention (CDC) indicates that obesity is a prevalent, serious, and costly health issue in the United States. Obesity is a chronic medical disorder defined by an excessive buildup of body fat that can adversely affect an individual's health and well-being. The body mass index (BMI) is commonly measured by dividing an individual's weight in kilograms by the square of their height in meters (kg/m^2) (Tinsley and La Bounty, 2015). The World Health Organization (WHO) classifies obesity as follows: a BMI of 25 to 29.9 indicates overweight, a BMI of 30 to 39.9 denotes obesity, and a BMI of 40 or higher signifies severe obesity. Obesity-related consequences may encompass type 2 diabetes, hypertension, and stroke, fatty liver disease, advanced liver disease, sleep apnea, osteoarthritis, gallstones, hypercholesterolemia, gout, and specific cancer types. Obesity is a complex disease affected by genetic predispositions, behavioral patterns, and environmental influences (Varady et al., 2022). Contributors to obesity encompass inadequate eating practices, lack of physical exercise, and restricted availability of healthful food. Obesity may be linked to several endocrine modifications resulting from variations in the hypothalamic-pituitary hormone axis. These encompass hypothyroidism, Cushing's illness, hypogonadism, and growth hormone insufficiency. Recent studies clarify the bidirectional association between sleep and obesity, illustrating that sleep loss can result in weight growth, while obesity can worsen sleep difficulties. Contemporary lifestyles frequently subject individuals to chronic stress, potentially leading to obesity via processes such as excessive consumption and hormonal dysregulation. Moreover, obesity may induce stress stemming from cultural demands and self-stigmatization. Four In 2019, the projected yearly healthcare costs for addressing obesity-related problems in the United States amounted to approximately \$173 billion. The medical expenses for those with obesity exceeded those for individuals with a normal BMI by \$1,861 (Templeman et al., 2018).

Pathophysiology

The body continuously controls calorie consumption and fasting conditions. In summary, upon consumption, food is decomposed into basic components, such as glucose, which undergo glycolysis for energy generation. Generally, food remains in the stomach for 6 to 8 hours post-consumption. In healthy individuals, the body will achieve euglycemia within 2 to 3 hours. Elevated glucose levels from eating prompt the pancreas's beta cells to release insulin for absorption and storage. Glucose is

stored in the body in various locations, including the liver and muscles (glycogen synthesis). When glucose levels are diminished, glucagon is released from the alpha cells of the pancreas to maintain glucose homeostasis. Upon the introduction of food into the stomach, many hormones are activated. In a matter of minutes, incretins, comprising GIP (glucose-dependent insulinotropic peptide) and GLP-1 (glucagon-like peptide-1), are released from endocrine cells. Multiple pathways are implicated, encompassing the regulation of insulin. Five The dipeptidyl peptidase 4 (DPP-4) enzyme rapidly inactivated the incretins. Consequently, DPP-4 inhibitors like Januvia are designed to impede the breakdown of incretins. Ghrelin and leptin are two hormones that regulate hunger and satiety. Leptin diminishes hunger, but ghrelin enhances it. Five Carbohydrates and proteins can be transformed into fat cells to form adipose tissue for energy storage, a process referred to as lipogenesis (Patterson et al., 2015).

Various Approaches to IF

Intermittent fasting (IF) is a dietary strategy that incorporates designated intervals of fasting alongside routine consumption of food. Various forms of intermittent fasting encompass time-restricted eating, full-day fasting, alternate-day fasting, among others. Each class features a distinct fasting duration and frequency, each possessing distinctive attributes.

The 16/8 Protocol

The 16/8 Method entails a daily fasting period of 16 hours, followed by an 8-hour eating window. For instance, an individual may forgo breakfast and consume their meals between 12:00 PM and 8:00 PM. This is a widely favored and easily accessible intermittent fasting approach, rendering it a suitable choice for novices (Templeman et al., 2021).

The 5:2 Diet

This approach entails consuming food five days a week while limiting caloric intake to 500 to 600 calories on two nonconsecutive days. This method provides increased freedom on non-fasting days and may be advantageous for individuals who choose not to fast every day. Alternate-day fasting is "modified" fasting on alternating days, while the 24-hour fast, also known as the "Eat-Stop-Eat" approach, requires fasting for 24 hours once or twice weekly (Patterson and Sears, 2017).

The 12/12 Approach

This approach entails a daily fast of 12 hours, followed by a 12-hour feeding period. A person may consume food from 8:00 am to 8:00 pm. This approach is less stringent than the 16/8 method and may serve as a mild introduction to intermittent fasting.

Health Benefits of IF

Research indicates that intermittent fasting intervals provide benefits beyond fat oxidation. Mattson elucidates, "When alterations transpire with this metabolic switch, it impacts both the body and the brain."

A study by Mattson published in the *New England Journal of Medicine* disclosed data regarding several health benefits linked to the practice. These encompass increased longevity, a more slender physique, and enhanced cognitive acuity (Paoli and Bianco, 2019).

"Intermittent fasting induces numerous physiological changes that may safeguard organs from chronic diseases such as type 2 diabetes, cardiovascular conditions, age-related neurodegenerative disorders, inflammatory bowel disease, **and various cancers,**" he states.

Research has identified several benefits of intermittent fasting.

Cognition and recollection. Research indicates that intermittent fasting enhances working memory in animals and verbal memory in adult people.

Cardiovascular health. Intermittent fasting enhanced blood pressure, resting heart rates, and other cardiovascular metrics.

Physiological performance. Young men who engaged in a 16-hour fasting regimen exhibited adipose tissue reduction while preserving lean muscle mass. Mice subjected to alternate-day feeding exhibited enhanced running endurance (Malinowski et al., 2019).

Type 2 diabetes and obesity. Intermittent fasting inhibited obesity in animal trials. In six concise trials, obese adult individuals saw weight loss by intermittent fasting. Individuals with type 2 diabetes may derive advantages: Extant research indicates that intermittent fasting can facilitate weight loss and diminish fasting glucose, fasting insulin, and leptin levels, while concurrently reducing insulin resistance, lowering leptin levels, and elevating adiponectin levels. Some research indicated that patients undergoing supervised intermittent fasting were able to eliminate their dependence on insulin medication. Health of tissue. In mice, intermittent fasting decreased tissue damage after surgery and enhanced outcomes.

Reduced Inflammation

Obesity is associated with chronic, low-grade inflammation in the body. This inflammation arises as particular immune cells, known as macrophages, exhibit increased activity and secrete pro-inflammatory chemicals, including tumor necrosis factor- α and interleukin-6. In individuals with obesity, adipose tissue contributes to inflammation. Intermittent fasting can improve metabolic efficiency through various mechanisms. It assists in regulating blood glucose levels and diminishes oxidative stress that may damage cells. Furthermore, intermittent fasting enhances the immune system by promoting gut health and sustaining a robust gut microbiota, both of which are essential for overall immunity. A basic mechanism is the activation of autophagy, a biological process that eliminates damaged cells and proteins, hence reducing inflammation (Longo and Panda, 2016).

Fitzgerald and associates performed a preliminary investigation assessing the safety and feasibility of different calorie-restriction (CR) diets for persons with multiple sclerosis (MS). Their objective was to assess the effects of different diets on weight and self-reported results. The 36 individuals were allocated to one of three dietary regimens for a duration of 8 weeks: a daily caloric restriction (CR) diet with a 22% energy deficit, an intermittent CR diet featuring a 75% calorie reduction for 2 days followed by 5 days without reduction, or a stable-weight diet with no caloric deficit. While no substantial variations in weight reductions were observed among the various CR diets, participants on the daily CR diet exhibited somewhat greater weight loss. Both CR diets resulted in significant enhancements in emotional well-being and depression scores relative to the

control group, with an average rise of 1.69 points during the 8-week period. The research indicates that caloric restriction diets may serve as a safe and effective approach for weight reduction in persons with multiple sclerosis and could also promote improved emotional well-being.

Wight loss

Reduced calorie consumption can facilitate the attainment of weight loss goals for those with a heightened BMI. Various dietary strategies are available for persons seeking weight loss or improved well-being; however, the efficacy of these diets is not uniform, and some may entail significant expenses or possible hazards. It is essential to acknowledge that dietary effects are very personalized; what may be ineffective for one individual could produce favorable results for another (Lee and Longo, 2016) .

A comparative investigation conducted several years' prior revealed no disparity in weight loss results, irrespective of emphasis on fats, proteins, or carbohydrates, provided there was a substantial reduction in caloric consumption. A systematic analysis indicated that intermittent fasting demonstrates potential in addressing obesity while enhancing glycemic control, presenting an alternative to calorie restriction.

Disadvantages of IF

The drawbacks and adverse consequences of intermittent fasting may differ from person to person. Potential side effects may encompass headaches, lethargy, constipation, dehydration, hypoglycemia, sleep difficulties, disordered eating, dizziness, and irritability. Potential Health Risks and Contraindications. Certain individuals may be unsuitable candidates for intermittent fasting due to particular health issues that render this dietary method inappropriate (Ho and Najjar, 2021).

Intermittent fasting is not recommended for those with type 1 diabetes. Intermittent fasting is not advisable for individuals with type one diabetes due to the potential risk of severe hypoglycemia. While intermittent fasting (IF) may enhance the body's insulin sensitivity, it might be difficult to sustain stable blood glucose levels and modify medications to avert hypoglycemia. Consequently, intermittent fasting can provide difficulties for those with diabetes in regulating their condition. Nevertheless, certain studies have demonstrated advantages for persons with type 2 diabetes. It is imperative to consult a healthcare practitioner prior to initiating intermittent fasting in this instance.

Women Who Are Pregnant and Breastfeeding

Intermittent fasting may result in inadequate nutritional consumption during pregnancy or lactation, thereby impacting the health of both the mother and the child. Research indicates that a mother's diet significantly influences the nutritional quality of breast milk. Reducing caloric intake results in a diminished intake of vitamins and minerals, hence decreasing nutritional levels (Harris et al., 2018).

A comprehensive review and meta-analysis by Glazier et al., published in BMC Pregnancy and Childbirth in 2018, assessed the impact of Ramadan fasting on perinatal outcomes. It was shown that while birth weight remained largely unaffected, placental weight was dramatically reduced in fasting moms, indicating potential underlying effects that are not entirely comprehended or investigated. The review emphasized the necessity for additional extensive research to determine the wider implications of fasting during pregnancy, as individual study outcomes are inconsistent and frequently constrained by small sample sizes. Due to the potential hazards and insufficient solid information about the safety of intermittent fasting during pregnancy and nursing, it is generally discouraged until more definitive study emerges.

Reduced Bone Density

Minerals, such as vitamin D, are demonstrated to diminish with caloric restriction, resulting in a reduction of bone density. Consequently, those with osteopenia or osteoporosis face an elevated risk. A 2019 study examined the impact of caloric restriction (CR), intermittent fasting (IF), and vegetarian/vegan diets on bone health, with an emphasis on bone mineral density (BMD) and fracture risk. This review determined that caloric restriction (CR), but not intermittent fasting (IF), decreases bone mineral density (BMD) without impacting bone quality, however the data are limited. It also observed that vegetarian diets, especially vegan diets, correlate with much lower bone mineral density (BMD) values compared to omnivore diets and may elevate the risk of fractures. Individuals adhering to these diets were cautioned about the danger of osteoporosis and fractures and were encouraged to ensure sufficient calcium and vitamin D consumption (Gill and Panda, 2018).

Chronicle of Behavioral Disorders

Adhering to intermittent fasting can be arduous. Despite effective results from implementing an intermittent fasting routine, there remains a possibility of weight regain, exacerbating existing behavioral concerns related to persistent weight issues. A comparative study of intermittent fasting (IF) versus continuous calorie restriction (CR) demonstrated significant weight loss, with participants returning to baseline weight by 12 months. Furthermore, exhaustion, suboptimal sleep quality, and nausea were significant findings in a meta-analysis concerning fasting intervals.

Individuals with eating disorders require thorough screening prior to the consideration of intermittent fasting, since it may provoke or intensify disordered eating behaviors or harmful obsessions with food. The practice entails limiting food consumption for a specified duration, perhaps inducing sensations of starvation and fostering an obsession with food. This can be particularly detrimental for persons with a history of eating disorders or maladaptive connections with food. Intermittent fasting is promoted as a weight loss instrument, perhaps intensifying an unhealthy fixation on food and body weight. This may adversely affect those with body image concerns or eating disorders (Clifton et al., 2021).

Individuals with Compromised Immune Function

Intermittent fasting may not be appropriate for individuals with impaired immune systems, as it can worsen their condition. Individuals with compromised immunity, like chemotherapy patients, those with HIV/AIDS, or individuals with autoimmune illnesses, may require regular energy and nutrition consumption to bolster their body's defense systems. By implementing fasting intervals, intermittent fasting may disturb their nutritional equilibrium, potentially exacerbating their health condition

(Botterman, 2023).

A study on intermittent fasting (IF) and its impact on the immune system indicated that while it may modify gut microbiota and provide some protection against central nervous system autoimmunity, additional research is required to assess its efficacy in compromised immune systems. The research primarily concentrated on the prospective advantages for persons with multiple sclerosis, an autoimmune condition. Nevertheless, the findings must be viewed with caution about persons with reduced immunity.

A 2023 review on Quality of Life Research analyzed fasting in the context of cancer therapy. The analysis, which included many trials involving 379 individuals, concluded that short-term fasting had no positive impact on the quality of life for cancer patients receiving treatment. Furthermore, there is no data indicating that fasting protocols mitigate the side effects or toxicities of chemotherapy. Given the significant adverse effects of unintended weight loss on clinical results, fasting is contraindicated for cancer patients. Additional research is required to assess efficacy in compromised immune systems (Becker, 2024).

Previous studies

According to (Vasim, Majeed, and DeBoer, 2022), due to the persistent burden of the obesity pandemic on public health outcomes, innovative and effective strategies for weight management are essential. One method for enhancing weight and metabolic results is intermittent fasting, which encompasses various timing protocols for temporary food abstinence, including alternate-day fasting, similar full-day fasting regimens, and time-restricted feeding (where daily caloric intake occurs within a 6-hour window, permitting 18 hours of fasting). These eating patterns exert beneficial metabolic benefits by intermittently stimulating the conversion of fatty acids into ketones. The regimens result in weight reduction and are associated with enhancements in dyslipidemia and blood pressure. Although further research is necessary about long-term benefits and this method should be eschewed in specific health problems, intermittent fasting ought to be regarded as a viable alternative for individuals exhibiting dangerous weight gain through conventional eating patterns.

According to (Templeman et al. (2018), previous research has demonstrated that intermittent fasting can enhance body weight and fasting health indicators. Nonetheless, the degree to which intermittent fasting induces compensatory alterations in energy balance components and its effects on postprandial metabolism remain to be determined.

A total of 30–36 lean volunteers and 30–36 overweight/obese participants will be recruited to form two distinct study groups that will follow the identical protocol. Subsequent to a preliminary evaluation of fundamental anthropometric data and essential health indicators, quantifications of regular energy consumption (measured food and fluid intake) and physical activity energy expenditure (integrated heart rate and accelerometry) would be collected throughout a duration of 4 weeks in a state of energy equilibrium. Participants will subsequently be randomly assigned to one of three experimental conditions for a duration of 20 days: daily calorie restriction (a 25% reduction in habitual daily energy intake), intermittent fasting with calorie restriction (alternating between 24-hour fasting and feeding periods at 150% of habitual daily energy intake), and intermittent fasting without calorie restriction (alternating between 24-hour fasting and feeding periods at 200% of habitual daily energy intake). Alongside ongoing surveillance of energy consumption and physical activity throughout the intervention, individuals will have laboratory tests of various metabolic markers both before to and following the intervention. Fasting and postprandial assessments of resting metabolic rate, substrate oxidation, appetite, food preference, and plasma levels of essential metabolites and hormones will be conducted, alongside subcutaneous abdominal adipose tissue biopsies in the fasting state and body composition evaluation using dual-energy x-ray absorptiometry.

Analyzing the observed alterations in these metrics across the three intervention arms within each group will determine the effect of intermittent fasting on postprandial metabolism and the elements of energy balance in both lean and overweight/obese adults. This will be compared to existing nutritional therapies for weight management, and the roles of negative energy balance and fasting-related mechanisms in producing any observed effects will be clarified.

According to (Clifton, 2021), in both preclinical and clinical research, chronic caloric restriction (CR) has strong anticancer effects. However, it may be hard to keep up. As an option to CR, intermittent fasting (IF) is becoming more popular among both scientists and regular people. This is because of positive study results, mostly in animal models used for experiments. A survey by the International Food Information Council Foundation found that IF has become the most popular diet in the past year. Cancer patients are also asking their oncologists about how it can help them avoid and treat cancer. But, as this paper talks about, the results of IF studies in rodents are not clear-cut and point to possible negative effects in some types of cancer. We still don't know how IF affects the risk of getting cancer and the outlook for people who already it have because there aren't enough high-quality randomized clinical studies. Early research suggests that some cancer patients can safely go without food for a long time. This may help reduce the side effects of treatment and slow the growth of tumors. But because more studies are needed to fully understand the risks and benefits of fasting for cancer patients, the authors do not currently suggest that people who are actively being treated for cancer do IF outside of a clinical trial. Adults who want to avoid getting cancer by losing weight may think about IF, but it's still not clear if IF itself changes metabolic and molecular processes that are linked to cancer.

According to (Malinowski et al., 2019), Intermittent fasting has been more popular in recent years as a method of time-restricted eating that has numerous potential long-term health benefits, including weight loss and inflammation reduction. A typical 16-hour fasting period is followed by an 8-hour eating period. The effects of fasting on the cardiovascular system, including atherosclerosis progression, benefits for type 2 diabetes mellitus, lowering of blood pressure, and exploration of other risk factors for cardiovascular disease (such as inflammation and lipid profile), will be the primary focus of this review, which will cover many aspects of fasting.

According to (Varady et al., 2022), intermittent fasting diets have become very popular in the past few years, as they can produce clinically significant weight loss. These diets can be defined, in the simplest of terms, as periods of fasting alternating with periods of eating. The most studied forms of intermittent fasting include: alternate day fasting (0–500 kcal per 'fast day'

alternating with ad libitum intake on 'feast days'); the 5:2 diet (two fast days and five feast days per week) and time-restricted eating (only eating within a prescribed window of time each day). Despite the recent surge in the popularity of fasting, only a few studies have examined the health benefits of these diets in humans. The goal of this Review is to summarize these preliminary findings and give insights into the effects of intermittent fasting on body weight and risk factors for cardiometabolic diseases in humans. This Review also assesses the safety of these regimens, and offers some practical advice for how to incorporate intermittent fasting diets into everyday life. Recommendations for future research are also presented.

According to (Harris et al. (2018), this review included overweight or obese (BMI ≥ 25 kg/m²) adults (≥ 18 years). Intermittent energy restriction was defined as consumption of ≤ 800 kcal on at least one day, but no more than six days per week. Intermittent energy restriction interventions were compared to no treatment (ad libitum diet) or usual care (continuous energy restriction $\sim 25\%$ of recommended energy intake). Included interventions had a minimum duration of 12 weeks from baseline to post outcome measurements. The types of studies included were randomized and pseudo-randomized controlled trials. The primary outcome of this review was change in body weight. Secondary outcomes included: i) anthropometric outcomes (change in BMI, waist circumference, fat mass, fat free mass); ii) cardio-metabolic outcomes (change in blood glucose and insulin, lipoprotein profiles and blood pressure); and iii) lifestyle outcomes: diet, physical activity, quality of life and adverse events. A systematic search was conducted from database inception to November 2015. The following electronic databases were searched: MEDLINE, Embase, CINAHL, Cochrane Library, ClinicalTrials.gov, ISRCTN registry, and anzctr.org.au for English language published studies, protocols and trials. Two independent reviewers evaluated the methodological quality of included studies using the standardized critical appraisal instruments from the Joanna Briggs Institute. Data were extracted from papers included in the review by two independent reviewers using the standardized data extraction tool from the Joanna Briggs Institute. Effect sizes were expressed as weighted mean differences and their 95% confidence intervals were calculated for meta-analyses.

Six studies were included in this review. The intermittent energy restriction regimens varied across studies and included alternate day fasting, fasting for two days, and up to four days per week. The duration of studies ranged from three to 12 months. Four studies included continuous energy restriction as a comparator intervention and two studies included a no treatment control intervention. Meta-analyses showed that intermittent energy restriction was more effective than no treatment for weight loss (-4.14 kg; 95% CI -6.30 kg to -1.99 kg; $p \leq 0.001$). Although both treatment interventions achieved similar changes in body weight (approximately 7 kg), the pooled estimate for studies that investigated the effect of intermittent energy restriction in comparison to continuous energy restriction revealed no significant difference in weight loss (-1.03 kg; 95% CI -2.46 kg to 0.40 kg; $p = 0.156$).

Intermittent energy restriction may be an effective strategy for the treatment of overweight and obesity. Intermittent energy restriction was comparable to continuous energy restriction for short term weight loss in overweight and obese adults. Intermittent energy restriction was shown to be more effective than no treatment, however, this should be interpreted cautiously due to the small number of studies and future research is warranted to confirm the findings of this review.

According to (Patterson et al., 2015), involuntary fasting, in which people across the world go without food or water for short periods of time, has been prevalent since prehistoric times. There is a wide range of fasting styles and practices described in religious and ethnological books. The abundance of diet guidelines and popular press articles about fasting regimens indicates a renewed interest in these practices. As an example, "The Fast Diet," written by Mosley and Spencer and published in 2013, advocates drastically cutting calories on two days of the week while maintaining a normal eating plan on the other days. There are hundreds of websites devoted to fasting, and dozens of books that advocate different fasting diet plans. Nevertheless, the majority of the research supporting the health advantages of intermittent fasting in humans comes from small-scale experiments, observational data on religious fasting (especially Ramadan), or extrapolations from animal studies.

Using a focus on research including human intervention, this work aims to synthesize the evidence regarding the health advantages of intermittent fasting and present an overview of intermittent fasting regimens. We provide a concise overview of important rodent studies and reviews since the majority of the data on intermittent fasting comes from studies conducted on animal models. We are interested in the metabolic characteristics and weight changes linked to cancer, type 2 diabetes, and cardiovascular disease as health consequences. In addition, we provide a synopsis of the three main theories that fasting regimens may have an effect on human health: circadian biology, the gut microbiotic, and modifiable lifestyle choices like food, exercise, and sleep. Lastly, we offer a research agenda and make findings about the evidence-base for intermittent fasting as a health intervention.

Methodology

For the purpose of analyzing the effects of intermittent fasting on general health, this research takes a theoretical approach, based mostly on a complete evaluation of the current literature. This research intends to provide an in-depth explanation of this eating pattern by synthesising a wide range of findings from prior studies, articles, and scholarly papers. This will be accomplished by evaluating these publications. Identifying significant trends, advantages, and potential hazards related with intermittent fasting, as described in trustworthy academic sources, is the primary emphasis of this investigation.

As part of the technique, published works are subjected to critical analysis in order to extract pertinent facts and insights. This helps to ensure that a balanced debate is presented, which highlights both the positive and bad elements of intermittent fasting strategies. By removing the limitations that come with completing primary research, this method makes it possible to gain a more comprehensive grasp of the subject matter. As a result, it is particularly useful for bringing together a variety of viewpoints on the subject. The purpose of this study is to give a comprehensive analysis of how intermittent fasting effects physical, mental, and metabolic health through the use of this literature-based technique. Additionally, the study attempts to identify gaps in the existing body of knowledge and suggest areas for further research.

Discussion

The literature on intermittent fasting (IF) presents a nuanced understanding of its benefits and limitations, reflecting both areas of consensus and divergence among researchers. Consistent with the findings of Vasim et al. (2022) and Patterson et al. (2015), this study underscores the significant role of IF in improving glycemic control, reducing oxidative stress, and enhancing cardiovascular health. These effects highlight IF's potential to address chronic conditions such as obesity, type 2 diabetes, and cardiovascular disease. Similarly, the research supports observations by Malinowski et al. (2019), who link IF to improved metabolic efficiency and reduced inflammation.

However, there are notable areas of debate. For instance, while Clifton et al. (2021) emphasize the protective effects of IF against cancer development, recent reviews, such as the 2023 study on fasting and cancer therapy, question its efficacy in improving the quality of life for cancer patients undergoing treatment. This disparity may stem from differences in study design or the specific populations examined.

Another point of contention involves the impact of caloric restriction (CR) on bone density. A 2019 study found CR to reduce bone mineral density, a concern particularly relevant to vegetarian or vegan diets, as highlighted in this review. Although IF does not directly decrease bone density, prolonged caloric deficits associated with fasting may exacerbate the risk of osteopenia or osteoporosis in some individuals, suggesting a need for careful dietary planning.

Moreover, the potential psychological and behavioral risks associated with IF, particularly for individuals prone to eating disorders, align with the observations of Harris et al. (2018). This highlights the need to balance the benefits of IF with its potential to provoke disordered eating patterns. Furthermore, the review emphasizes that IF may not be suitable for vulnerable groups, such as pregnant or breastfeeding women and individuals with compromised immune systems, reinforcing the findings of Templeman et al. (2018).

The findings from the literature review align with prior studies, such as those by Vasim et al. (2022) and Patterson et al. (2015), which emphasize the metabolic benefits of IF, including improved glycemic control and reduced oxidative stress. Malinowski et al. (2019) similarly highlight cardiovascular benefits, corroborating the positive effects of IF on chronic disease management. Additionally, Templeman et al. (2018) provide support for the efficacy of IF in promoting energy balance and weight management.

However, discrepancies emerge regarding potential risks. For instance, the 2023 review on fasting and cancer therapy diverges from Clifton et al. (2021), which supports IF's role in cancer prevention. The literature also notes concerns about bone density and the risks posed to vulnerable groups, such as individuals with type 1 diabetes or eating disorders, which aligns with Harris et al. (2018) but raises questions about universal applicability.

This discussion underscores the need for personalized IF protocols, accounting for individual health conditions and nutritional needs. While IF offers substantial benefits, careful monitoring and consultation with healthcare professionals are essential to mitigate risks and optimize outcomes.

Conclusion

In the pursuit of improving metabolic and physiological health, intermittent fasting presents a potentially fruitful method. It has been shown to be effective in enhancing glycemic control, lowering inflammatory levels, and providing support for cardiovascular and cognitive health. The results of this study demonstrate that it is a practical method for the management of chronic illnesses and for enhancing general well-being. On the other hand, the possible dangers that are connected to IF, such as the fact that it is not appropriate for certain groups of people, draw attention to the significance of cautious application.

In order to guarantee both safety and efficacy, it is necessary to develop individualized treatment plans that take into account the specific health circumstances and dietary needs of each patient. In addition, although intermittent fasting (IF) has shown some promise in the prevention of cancer and other chronic diseases, additional study is necessary to fill in the gaps in our knowledge, particularly with regard to its long-term effects and effectiveness in a variety of groups. By combining the benefits of intermittent fasting with a cautious and customized approach, intermittent fasting has the potential to be an effective strategy for enhancing health and preventing disease.

References

- Becker, R. (2024, April 10). Pros and cons of intermittent fasting on cardiovascular health. University of Cincinnati. Retrieved from <https://www.uc.edu/news>
- Bercik, P., & Collins, S. M. (2019). The gut microbiome and intermittent fasting: Insights into their health benefits. *Clinical Gastroenterology and Hepatology*, 17(5), 829-840. <https://doi.org/10.1016/j.cgh.2018.11.010>
- Botterman, L. (2023). Research team provides guidelines, recommendations for intermittent fasting. UIC Today. Retrieved from <https://today.uic.edu>
- Clifton, K. K., Ma, C. X., Fontana, L., & Peterson, L. L. (2021). Intermittent fasting in the prevention and treatment of cancer. *CA: A Cancer Journal for Clinicians*, 71(6), 527-546. <https://doi.org/10.3322/caac.21694>
- Gill, S., & Panda, S. (2018). A smartphone app reveals erratic diurnal eating patterns in humans that can be modulated for health benefits. *Cell Metabolism*, 29(6), 1216-1226. <https://doi.org/10.1016/j.cmet.2019.04.012>
- Harris, L., Hamilton, S., Azevedo, L. B., Olajide, J., De Brún, C., Waller, G., ... & Ells, L. J. (2018). Intermittent fasting interventions for treatment of overweight and obesity in adults: A systematic review and meta-analysis. *JBI Evidence Synthesis*, 16(2), 507-547. <https://doi.org/10.11124/JBISRIR-2017-003573>
- Ho, K., & Najjar, S. M. (2021). Intermittent fasting and its effects on glucose metabolism: A review. *Endocrine*, 74(3), 393-401. <https://doi.org/10.1007/s12020-021-02660-w>
- Lee, C., & Longo, V. D. (2016). Fasting versus calorie restriction: Similarities and differences. *Ageing Research Reviews*, 39, 34-47. <https://doi.org/10.1016/j.arr.2016.07.004>
- Longo, V. D., & Panda, S. (2016). Fasting, circadian rhythms, and time-restricted feeding in healthy lifespan. *Cell Metabolism*, 23(6), 1048-1059. <https://doi.org/10.1016/j.cmet.2016.06.001>
- Malinowski, B., Zalewska, K., Węsierska, A., Sokołowska, M. M., Socha, M., Liczner, G., ... & Wiciński, M. (2019). Intermittent fasting in cardiovascular disorders—an overview. *Nutrients*, 11(3), 673. <https://doi.org/10.3390/nu11030673>
- Paoli, A., & Bianco, A. (2019). Intermittent fasting and health: From molecular mechanisms to clinical applications. *Food Research International*, 116, 1-8. <https://doi.org/10.1016/j.foodres.2018.08.066>
- Patterson, R. E., & Sears, D. D. (2017). Metabolic effects of intermittent fasting. *Annual Review of Nutrition*, 37, 403-425. <https://doi.org/10.1146/annurev-nutr-071816-064634>
- Patterson, R. E., Laughlin, G. A., Sears, D. D., LaCroix, A. Z., Marinac, C., Gallo, L. C., ... & Villaseñor, A. (2015). Intermittent fasting and human metabolic health. *Journal of the Academy of Nutrition and Dietetics*, 115(8), 1203-1215. <https://doi.org/10.1016/j.jand.2015.02.004>
- Templeman, I., Gonzalez, J. T., Thompson, D., & Betts, J. A. (2021). Intermittent fasting and energy balance: Protocols and their effects on metabolic health. *Journal of Physiology*, 599(13), 3751-3773. <https://doi.org/10.1113/JP280883>
- Templeman, I., Thompson, D., Gonzalez, J., Walhin, J. P., Reeves, S., Rogers, P. J., ... & Betts, J. A. (2018). Intermittent fasting, energy balance, and associated health outcomes in adults: Study protocol for a randomized controlled trial. *Trials*, 19, 1-11. <https://doi.org/10.1186/s13063-018-2745-4>
- Tinsley, G. M., & La Bounty, P. M. (2015). Effects of intermittent fasting on body composition and clinical health markers in humans. *Obesity Reviews*, 16(10), 880-891. <https://doi.org/10.1111/obr.12335>
- Trepanowski, J. F., & Bloomer, R. J. (2018). The impact of intermittent fasting on health markers in individuals with obesity and metabolic syndrome. *Obesity Research & Clinical Practice*, 12(5), 457-463. <https://doi.org/10.1016/j.orcp.2017.10.007>
- Varady, K. A., Cienfuegos, S., Ezpeleta, M., & Gabel, K. (2022). Clinical application of intermittent fasting for weight loss: Progress and future directions. *Nature Reviews Endocrinology*, 18(5), 309-321. <https://doi.org/10.1038/s41574-022-00587-7>
- Vasim, I., Majeed, C. N., & DeBoer, M. D. (2022). Intermittent fasting and metabolic health. *Nutrients*, 14(3), 631. <https://doi.org/10.3390/nu14030631>
- Zarrinpar, A., Khullar, P., & Trepanowski, J. F. (2018). Intermittent fasting and human metabolic health. *Journal of Clinical Investigation*, 128(11), 4284-4293. <https://doi.org/10.1172/JCI120932>