

The MIND Diet

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If four and a half billion years of evolution, twelve thousand years of agriculture and a hundred years of nutritional science (Mozaffarian *et al*, 2018) has taught us anything, it is that the importance of food simply cannot be underestimated. Indeed, the dramatic increase in human life span to double that of ancestral humans, was largely due to improved access to nutrition and medicine (Finch and Caleb, 2010). Furthermore, there is now significant evidence to suggest that many major diseases such as cardiovascular disease (Mente *et al*, 2009), or cancer (Bringham and Sheila *et al*, 2004) are better combated by prevention with effective diet than direct medical treating which can be very expensive, and imply unpleasant side effects. The role of diet, in basic nutrition and disease prevention is especially evident in old age, with the increased incidence of degenerative diseases such as Alzhiemers (Arendt and Bigl, 1987). However, despite the well established importance of diet, what would constitute the ideal diet, is still not entirely clear, although many suggestions have been made. The MIND (Mediterranean-DASH Intervention for Neurodegenerative Delay) diet, is one such suggestion. It combines aspects of the classic Mediterranean diet, consisting predominately of fish, fruit and vegetables, with the newer DASH (Dietary Approaches to Stop Hypertension) diet, which includes increased dairy consumption and decreased sodium intake, to prevent neurodegenerative disease. This dietary plan has not gone without criticism, however, its overall efficacy is well supported by field trials, and it serves as a much needed guideline in the complex and confusing arena of dietary science.

To truly appreciate the advantages offered by the MIND diet requires a more detailed view of its constituents and their metabolic effects. The mind diet is based around splitting common food sources into two general groups, brain healthy and brain unhealthy, encouraging increased consumption of the former, and decreased consumption of the latter. (Di Fiore N., 2015) Brain healthy foods include vegetables (especially green leafy) vegetables, whole grains, berries, nuts, beans, wine (in limited quantities), fish, poultry, and olive oil, whereas brain unhealthy foods include cheese, red meats, fried foods, butter/margarine, or pastries and sweets (Marcason and Wendy, 2015). All of these food groups, both healthy and unhealthy are broken down by specific anabolic pathways into their basic constituents, which can then be catabolised into new macromolecules, with the production of side/waste products along the way. (Garrett and Grisham, 4th edition). Hence, which food groups are consumed will affect the extent and amount of raw materials available for anabolism, and the exposure to toxic metabolic by products.

The metabolism of leafy green vegetables provides a poignant example of the importance of consuming the appropriate substrates for metabolism, and the potential advantages offered by the MIND diet. Leafy greens contain significant component of proteins and carbohydrates (Menasah *et al*, 2018), with very low unsaturated or trans fat components. (Singh *et al*, 2001). The starch component of the carbohydrates present can be hydrolyzed to glucose molecules, which provide an important energy source to brain cells, which can not use any other substrate as an energy source, while many of the proteins may contain metal ion prosthetic groups, which form a vital source of micro-nutrients, for the cell. In particular the Mg^{2+} ions present in the chlorophyll of such leaves is a vital co-factor for many of the enzymes involved in glycolysis, (the break down of glucose to release energy), hence without a steady supply of Mg^{2+} ions the brain cells would again be starved of an energy source, among many other deleterious consequences. Finally the low trans and unsaturated fat concentration in leafy green helps specifically to reduce the changes of neurodegenerative disease with highly unsaturated trans fats (such as in red meat or pastries), associated with the incidence of Alzheimer's. (Morris *et al*, 2003). Similar metabolic themes can be identified for the other components of the MIND diet, to provide a thorough explanation for its contribution to decreased risk of neurodegenerative disease.

The efficacy of the diet plan does not rest solely on theory however, as there is also a wealth of empirical evidence linking its application to decreased incidence of neurodegenerative disease. Stronger adherence to the diet has been strongly correlated to lower rates of decline in working memory, episodic memory, visiospatial ability, and perceptual speed, as general cognition scores, with strict adherence effectively delaying the deterioration process by as much as 7.5 years (Morris *et al*, 2015). Perhaps even more importantly even intermediate levels of compliance with the diet show a dramatic (35%) reduction in the incidence of Alzheimer

disease. (Moris *et al*, 2015). Furthermore the MIND diet is demonstrably more effective at Alzheimer’s prevention than either of its predecessors the Mediterranean or DASH diets. (Morris *et al*, 2014). Indeed, what experimental data has been collected strongly supports the MIND diet as not only an effect, but possibly the most effective dietary choice for Alzheimer’s prevention.

However, despite the theoretical and empirical support for the diet’s efficacy, it is not without limitations. Chief among these limitations is that despite the general correlation between the diet and Alzheimer’s prevention discussed above, the effect of the diet on any given individual is variable, dependent upon both genetics, and behavioral patterns. (Schelke, *et al*, 2018). Inclusion of personalized nutrition, as well as cognitive activity, excessive and even social engagement regimes, in addition to the basic MIND diet, proved considerably more effective in treatment and prevention of cognitive decay. For example, insulin resistance results in glucose hypo-metabolism in the brain, and increased expression of potentially neurotoxic amyloid precursor proteins, associated with cognitive decline. (Wiette, *et al*, 2015). Adherence to the MIND diet, specifically leafy vegetables and berries which contain high concentrations of oleic acid, can show significant reductions in insulin resistance (Ryan, 2000). However, the additional consumption of cocoa flavanols for individuals already following the mind diet limits insulin resistance even further. In this regard the MIND diet is not by its self the full solution for Alzheimer’s prevention, as other lifestyle, and even dietary factors can have serious impacts. Nonetheless, these limitations in no way mitigate the usefulness of the diet as a general indication of cognitive health. Individuals who are often without the resources or inclination to adhere to more fundamental lifestyle changes can still decrease their risk of developing neurodegenerative diseases by reasonable accordance to this relatively simple diet.

In conclusion, the MIND diet combines both the time tested Mediterranean diet, and its more recent derivative the DASH diet, to form a diet specifically tailored to prevention, and mitigation of neurodegenerative disease. The diet groups common food types generally into brain healthy and brain unhealthy foods, based on the biological consequences of the metabolites which they generate, both in terms of the nutritional value of end products of metabolism as well as the potential deleterious effects of the waste/side products. The theory behind this classification relies on considerations of well established and experimentally verified metabolic pathways, and empirical trials show strong links between the diet and decreased neurodegenerative disease. While the effectiveness of the diet may vary to a degree between individuals due to genetically or behaviorally based idiosyncrasies in metabolism. However, in general it provides an easily understandable and useful guideline for neurodegenerative disease prevention, even at relatively low levels of diet compliance. High levels of diet compliance combined with bespoke modifications to the basic diet plan can lead to truly extraordinary improvements in neurodegenerative disease prevention.

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