



Nutritional Psychiatry: Focuses on Improving Mental Health through Diet

¹Souvik Tewari,²Lina Sarkar³Sandhya Ranee Bhoi and⁴Rojina Swayamsiddha Sahu

¹Assistant Professor, dept.of Food and Nutrition, Swami Vivekananda University, Barrackpore, West Bengal, India.

²Assistant Professor, dept. of Psychology, Swami Vivekananda University, Barrackpore, West Bengal, India.

³Jr. Lecturer, dept. of Home Science, Govt. Women's College, Sambalpur, Odisha, India.

⁴Guest faculty, dept. of Home Science, Govt. Women's College, Sambalpur, Odisha, India.

To Cite this Article

Souvik Tewari, Lina Sarkar, Sandhya Ranee Bhoi and Rojina Swayamsiddha Sahu. Nutritional Psychiatry: Focuses on Improving Mental Health through Diet. *International Journal for Modern Trends in Science and Technology* 2021, 7 pp. 85-91. <https://doi.org/10.46501/IJMTST0712015>

Article Info

Received: 24 October 2021; Accepted: 01 December 2021; Published: 04 December 2021

ABSTRACT

Diet and nutrition have a significant role in both physical health as well as mental health, according to a growing body of studies. A growing body of evidence shows that a poor diet may contribute to the development of mental health issues such as anxiety and depression. Despite the lack of scientific evidence, scientists are now looking at the link between diet and mental health, which has hitherto been mostly ignored. Epidemiology does not show the fundamental reasons or processes of a good diet and mental health. What is going on should be the primary goal of future studies. High-quality, well-powered randomised controlled trials (RCTs) are needed to go from population-based observations to individualised nutrition. Nutritional psychiatry is a relatively new discipline, and this overview examines the empirical evidence that supports the necessity for a well-balanced diet for mental health. Evidence on nutrition and mental health is scant in terms of causation and underlying processes. In the future, researchers should focus on understanding out how everything works together. High-quality, well-powered randomised controlled trials should focus on transitioning from population-based observations to individual nutrition. You may learn more about why a healthy diet is so important for mental health by looking at the scientific data.

Keywords: mental health, nutrients, attention deficit hyperactivity disorder (ADHD), brain function, depression, minerals, vitamins.

Nutritional advice on mental health has a weak scientific evidence base.

“Whether the goal is to improve mood, enhance cognitive function, prevent decline, or even provide beneficial effects in certain brain diseases, such as epilepsy, attention deficit hyperactivity disorder (ADHD), and autism, information about the link between nutrition and mental health is increasingly invading our daily lives in the popular press.”

A common misunderstanding seems to exist that dietary advice for mental health are supported by solid research. Even in the best-case scenario, it might be difficult to prove that certain diets or certain food components have an effect on mental health.

One of the most important societal concerns of our day is the growth in mood disorders, stress-induced cognitive weaknesses, and mental diseases across Europe and the rest of the world during the next several

decades, according to all available statistics. Preventative measures are essential in the realm of public health. Research on nutrition's role in predicting mental health is vital, despite the difficulty of doing and understanding it.

There are a number of essential nutrients needed to keep the brain healthy and functioning properly.^{1,2,3} A person's ability to enhance their mental well-being, mood, and cognitive performance may be influenced by the amount and kind of food they consume and the quality of that food.^{4,5} Eating particular foods causes our bodies to release certain hormones, neuropeptides and neurons that influence our internal environment.^{6,7,8,9,10}

There is no way to prove a link between a person's mental health and their nutrition, even if they are linked by cross-sectional epidemiological research. Nutritional interventions with proper length and specificity seldom exhibit favourable mental health results, although there are a handful of notable exceptions. It is difficult to blind participants to the food intervention's nature, and there are not enough randomised studies to properly stratify people across populations. A nutritional intervention cannot be blinded to its nature because there is no random allocation to treatment conditions or observers who are not blinded in intervention studies. In general, nutritional interventions in healthy individuals have small effect sizes, making detection challenging. However, there is reason to be optimistic, because nutritional interventions could have a significant impact in cases of poor functioning or disease. Nutritional deficiencies (or excesses) in the diet, as well as specific dietary needs in sickness contexts, may contribute to disease development or severity, or even begin disease genesis. Nutrient deficiencies It is hoped that the emerging subject of "Nutritional Psychology"¹¹ would shed light on the question of whether dietary components are in fact essential for mental health, even in the case of mental diseases. It is possible to discover which dietary components are crucial for mental health, especially in the case of psychiatric disorder, and to whom, under what conditions, and at whose exact doses these nutritional treatments may be helpful as both preventive and therapeutic interventions.

Meta-analyses show that food has an impact on mental health.

Science has failed to demonstrate a clear link between food and mental health. Nevertheless, some studies have shown significant links between a good diet and mental health, which may have an influence on future dietary advice.¹² Those who eat a diet abundant in fresh fruits and vegetables, for example, are more likely to be happy and mentally well.^{13,14,15,16,17}

Diet and mental health have been the subject of several research and meta-analyses. According to the findings of four cohorts and nine cross-sectional studies, eating a diet rich in fruits, vegetables, seafood, and whole grains reduces one's risk of depression.¹⁸ According to the results of the second meta-analysis, which comprised eight cohort studies and one case-control research, Mediterranean diet consumers are less likely to suffer from depression.¹⁹ There is substantial evidence that the Mediterranean diet helps protect against depression, according to a recent systematic review of 20 long-term research and 21 cross-sectional studies.²⁰ Depression may be less common thanks to dietary therapy, according to a meta-analysis of 16 randomised, controlled research.²¹ Depression is not linked to the Mediterranean diet in a meta-analysis of cohort studies.²² In cross-sectional data, adherence to the Mediterranean diet was revealed to have an inverse correlation with depression risk. These studies, when considered together, provide a strong platform for further study into the impact of particular dietary modifications on mental health.

Vitamin deficiencies and disease-specific diets have an impact on mental health.

The ketogenic diet for children with epilepsy may have a positive influence on brain health.²³ There is no recognised procedure in this case; nevertheless, since ketone bodies are the brain's primary source of energy, this suggests that a shift in energy supply might be crucial.²⁴ Phenylketonuria may also be omitted from a diet in order to minimise cognitive damage.²⁵ Cognition may be affected by deficiencies in vitamins, particularly vitamin D, according to study (A shortage of vitamin B12 and thiamine can produce weariness, lethargy; depression; decreased memory; mania; and psychosis).^{26,27,28,29} Wernicke's encephalopathy and numbness in the central nervous system (CNS) may be

caused by folic acid insufficiency (vitamin B9; deficiency has detrimental effects on neurodevelopment during pregnancy and infancy; these deficiencies are connected to an increased risk of depression in adulthood).^{30,31,32}

Sub-clinical or mild deficits may play a role in the development of mental dysfunction. Numerous research have looked at the connection between vitamin D and mental well-being, yielding conflicting results on the subject. It has been shown that greater levels of blood vitamin D in the 65+ age group are associated with better attention and working memory.³³ An impact on depression and attention deficit/hyperactivity disorder has been suggested by randomised controlled trials (RCTs)³⁴ (albeit not universally).^{34,35} Bone health assessment cutoffs show that a considerable proportion of the general population is vitamin D deficient, underscoring the need of strong evidence for the vitamin's benefit in neuropsychiatric diseases. To enhance men's health, including their cognitive function, mood, stress sensitivity, and neuroinflammation, a diet high in polyphenols and polyunsaturated fatty acids (PUFAs) has been identified.³⁶⁻⁴⁶

Diets for ADHD and autism

There have been a variety of dietary therapy tested in the case of ADHD. If you remove food additives from your diet according to reports from your children, you may expect to see an impact size of 0.18, but this decreases to just 0.12 when publication bias is taken into account.⁴⁷ Additionally, several clinical investigations have shown that stricter elimination diets are beneficial. Six controlled studies were analysed by two separate meta-analyses and reported effect values ranging from 0.29 to 0.51, suggesting that about one-third of children with ADHD were responsive (symptom reduction of more than 40 percent).^{47,48,49} Supplementing with free fatty acids may help reduce ADHD symptoms, with an impact size of between 0.18 and 0.31, according to meta-analyses of studies on the subject.⁴⁹ Vitamins and minerals (micronutrients) have been related to decreased aggression and improved mood control in children with ADHD. An increase in the risk of ADHD or hyperactivity may be associated with a diet high in refined sugar and saturated fat, according to a new meta-analysis, whereas a diet high in fruits and

vegetables may have a protective effect.⁵⁰ Also highlighted is that the lack of study and design limitations invalidate current data; thus, longitudinal research should be undertaken in the future. Meta-analyses have not yet been conducted on nutritional therapy for autism, which include vitamin D supplementation, micronutrient supplementation, and gluten and casein-free diets. As a result, more rigorous studies are required, particularly in the field of autism, and a variety of mechanisms may account for its effectiveness.⁵¹

Towards diets for mental health

Most RCTs assessing the effectiveness of diet modifications in men's health are lacking. During one of the earliest intervention trials ever undertaken, a 12-week Mediterranean diet was utilised. There has been a marked improvement in mood and anxiety among people with severe depression, according to new research.⁵² People with depression who follow a Mediterranean-style diet, as studied by HELFIMED and PREDI DEP, reap the benefits.^{53,54} MoodFOOD RCT found that multinutrient supplementation had no impact on bouts of major depression in overweight or obese persons with subsyndromal depressive symptoms.^{55,56} For the development of nutritional psychiatry, it is essential to conduct research that seek to prevent and cure common mental health conditions. A greater number of blinded, placebo-controlled clinical studies is also required, as is Combining findings from all of the preceding scientific studies, it is possible that dietary modifications might alter brain function and mental health. Finding out how nutrition affects brain function in health and sickness will be an important step in the future. If specific minerals or whole-food diets might benefit mental health, we need to find out.⁵⁷ Dietary treatments may also be evaluated using experimental medicine methodologies; in order to make the greatest use of existing information, including the identification of acceptable biomarkers, we must optimise our selection of nutrients or diets to be examined in costly and protracted trials.

Diet, mental health, and cognition throughout the course of a lifetime

When it comes to illness susceptibility, research suggests that early life experiences might have an impact. As a result, early life development should be

taken into account when tailoring diet to improve mental health. It is possible that nutritional treatment administered during the early stages of brain development may have a greater impact on long-term health outcomes (the so-called first 1000 days, e.g. from conception to age two). Further progress in this field may be made with a greater understanding of how foods influence numerous signalling pathways such as metabolic and endocrine, immunological and other processes, including those that function via the gut microbiome.^{58,59,60}

There are both energy and nutritional restraints on the brain's growth and development in newborn humans, which limits its ability to expand and develop.⁶¹ Food security is essential during this time of increasing population increase. Nutritional psychiatry has focused on the cognitive effects of childhood malnourishment in the past several years.⁶²⁻⁶⁶

There is a clear link between the nutritional health of vulnerable groups, such as preterm and small for gestational age (SGA) children, and the possibility of neurological disorders.^{67,68} Nutrients like iron, choline and folate are crucial for brain development but other nutrients such as vitamins A, B6, B12 and fatty acids like omega-3 are also important.⁶⁹ The cyto-architecture of the cerebral cortex may be irreversibly altered during foetal development by an iodine deficiency, leading to aberrant neuron migration patterns and cognitive impairment in offspring. This has been shown in experiments. Infancy's iron deficiency anaemia has been linked to changes in the brain, despite the fact that the opposite has also been shown.⁷⁰ Also, even slight dietary changes may have an influence on the development of the brain in the womb.⁷¹ Lipids in breast milk include omega-3 and omega-6 polyunsaturated fatty acids DHA and ARA, however the mother's food diet affects the amount of these fatty acids in the milk.⁷² Neuronal membrane integration of omega-3 fatty acids was positively affected by diets high in omega-3 or low in omega-6 fatty acids, according to research on mice.^{73,74} Reducing the omega-6 content of adults' diets has been shown to completely reverse stress-induced cognitive impairments.⁷⁵ Stressed rats fed a high-fat diet rich in omega-3 polyunsaturated fatty acids and vitamin A showed enhanced cognitive behaviour and indicators of plasticity in the adolescent brain after the stress.⁷⁶ Early life stress was reduced in rats when given

MFGM and a polydextrose/galacto-oligosaccharide prebiotic mixture in a mother separation paradigm.⁷⁷ It is important to consume a healthy diet in order to support brain growth and cognitive function later in life, and this may explain at least in part why treatment outcomes vary so widely.

New concepts

Pharmacological therapies are not the same as nutritional interventions. When a drug has a high affinity for its target, it is able to attach to just one or a few of these targets. It is only with vitamins that such comparative study can be done, given their affinity is within a similar range to that of medicines, that we know of. Most nutrients are ingested in far larger quantities than medications, and they have a lower affinity for proteins than drugs. The greater quantities of nutrients and their metabolites allow them to bind and influence a wide spectrum of targets. Nutrition and its metabolites in general affect different organs and diverse targets. Rather of concentrating on specific nutrients or supplements, nutritional research is increasingly examining the influence of dietary patterns on the brain and behaviour, rather than the effects of specific nutrients or supplements.⁷⁸ Nutrients can interact to modify particular processes and interactions among nutrients can affect bioavailability, making it more difficult to identify individual substrates. This is crucial. Even more complicating the picture is the fact that certain nutrients have synergistic advantages, while others have been demonstrated to be ineffectual (for a recent review of nutraceuticals and a related stance on nutrition-based psychiatry, see).⁷⁹

Conclusion:

A growing body of evidence suggests that a person's diet has an impact on everything from stress tolerance to mental health and cognitive performance. However, the evidence is only correlative, and we still do not know exactly how these effects manifest themselves. In order to improve food-related public health policy, new breakthrough research on the bidirectional links between nutrition and brain function are urgently required. In the meanwhile, no such findings have surfaced. People's ability to maintain brain fitness throughout their lives will be improved by better understanding of how diet effects mental health and

cognition, leading to the development of innovative nutritional therapies and evidence-based recommendations. There would be an increase in healthcare system resiliency, along with lower costs, if dietary practises that benefit mental health were promoted and specific nutritional components were identified and confirmed.

REFERENCES

- [1] Castro AI, Gomez-Arbelaez D, Crujeiras AB, Granero R, Aguera Z, Jimenez-Murcia S, Sajoux I, Lopez-Jaramillo P, Fernandez-Aranda F, Casanueva FF. Effect of a very low-calorie ketogenic diet on food and alcohol cravings, physical and sexual activity, sleep disturbances, and quality of life in obese patients. *Nutrients*. 2018 Oct;10(10):1348.
- [2] Delpech JC, Madore C, Nadjar A, Joffre C, Wohleb ES, Layé S. Microglia in neuronal plasticity: influence of stress. *Neuropharmacology*. 2015 Sep 1;96:19-28.
- [3] Lépinay AL, Larrieu T, Joffre C, Acar N, Gárate I, Castanon N, Ferreira G, Langelier B, Guesnet P, Brétilon L, Parnet P. Perinatal high-fat diet increases hippocampal vulnerability to the adverse effects of subsequent high-fat feeding. *Psychoneuroendocrinology*. 2015 Mar 1;53:82-93.
- [4] de la Torre R, de Sola S, Hernandez G, Farré M, Pujol J, Rodriguez J, Espadaler JM, Langohr K, Cuenca-Royo A, Principe A, Xicota L. Safety and efficacy of cognitive training plus epigallocatechin-3-gallate in young adults with Down's syndrome (TESDAD): a double-blind, randomised, placebo-controlled, phase 2 trial. *The Lancet Neurology*. 2016 Jul 1;15(8):801-10.
- [5] Dinan TG, Stanton C, Long-Smith C, Kennedy P, Cryan JF, Cowan CS, Cénit MC, van der Kamp JW, Sanz Y. Feeding melancholic microbes: MyNewGut recommendations on diet and mood. *Clinical Nutrition*. 2019 Oct 1;38(5):1995-2001.
- [6] El Aidy S, Dinan TG, Cryan JF. Gut microbiota: the conductor in the orchestra of immune–neuroendocrine communication. *Clinical therapeutics*. 2015 May 1;37(5):954-67.
- [7] Sandhu KV, Sherwin E, Schellekens H, Stanton C, Dinan TG, Cryan JF. Feeding the microbiota-gut-brain axis: diet, microbiome, and neuropsychiatry. *Translational Research*. 2017 Jan 1;179:223-44.
- [8] Schellekens H, Finger BC, Dinan TG, Cryan JF. Ghrelin signalling and obesity: at the interface of stress, mood and food reward. *Pharmacology & therapeutics*. 2012 Sep 1;135(3):316-26.
- [9] Torres-Fuentes C, Schellekens H, Dinan TG, Cryan JF. The microbiota–gut–brain axis in obesity. *The lancet Gastroenterology & hepatology*. 2017 Oct 1;2(10):747-56.
- [10] Van de Wouw M, Schellekens H, Dinan TG, Cryan JF. Microbiota-gut-brain axis: modulator of host metabolism and appetite. *The Journal of nutrition*. 2017 May 1;147(5):727-45.
- [11] Adan RA, van der Beek EM, Buitelaar JK, Cryan JF, Hebebrand J, Higgs S, Schellekens H, Dickson SL. Nutritional psychiatry. *European Neuropsychopharmacology*. 2019:000-1.
- [12] Sarkar A, Harty S, Lehto SM, Moeller AH, Dinan TG, Dunbar RL, Cryan JF, Burnet PW. The microbiome in psychology and cognitive neuroscience. *Trends in cognitive sciences*. 2018 Jul 1;22(7):611-36.
- [13] Conner TS, Brookie KL, Carr AC, Mainvil LA, Vissers MC. Let them eat fruit! The effect of fruit and vegetable consumption on psychological well-being in young adults: A randomized controlled trial. *PloS one*. 2017 Feb 3;12(2):e0171206.
- [14] Emerson SD, Carbert NS. An apple a day: Protective associations between nutrition and the mental health of immigrants in Canada. *Social psychiatry and psychiatric epidemiology*. 2019 May;54(5):567-78.
- [15] Fresán U, Bes-Rastrollo M, Segovia-Siapco G, Sanchez-Villegas A, Lahortiga F, de la Rosa PA, Martínez-Gonzalez MA. Does the MIND diet decrease depression risk? A comparison with Mediterranean diet in the SUN cohort. *European journal of nutrition*. 2019 Apr;58(3):1271-82.
- [16] Moreno-Agostino D, Caballero FF, Martín-María N, Tyrovolas S, López-García P, Rodríguez-Artalejo F, Haro JM, Ayuso-Mateos JL, Miret M. Mediterranean diet and wellbeing: Evidence from a nationwide survey. *Psychology & health*. 2019 Mar 4;34(3):321-35.
- [17] Mujcic R, J. Oswald A. Evolution of well-being and happiness after increases in consumption of fruit and vegetables. *American Journal of Public Health*. 2016 Aug;106(8):1504-10.
- [18] Lai JS, Hiles S, Bisquera A, Hure AJ, McEvoy M, Attia J. A systematic review and meta-analysis of dietary patterns and depression in community-dwelling adults. *The American journal of clinical nutrition*. 2014 Jan 1;99(1):181-97.
- [19] Psaltopoulou T, Sergentanis TN, Panagiotakos DB, Sergentanis IN, Kosti R, Scarmeas N. Mediterranean diet, stroke, cognitive impairment, and depression: a meta-analysis. *Annals of neurology*. 2013 Oct;74(4):580-91.
- [20] Lassale C, Batty GD, Baghdadli A, Jacka F, Sánchez-Villegas A, Kivimäki M, Akbaraly T. Healthy dietary indices and risk of depressive outcomes: a systematic review and meta-analysis of observational studies. *Molecular psychiatry*. 2019 Jul;24(7):965-86.
- [21] Firth J, Marx W, Dash S, Carney R, Teasdale SB, Solmi M, Stubbs B, Schuch FB, Carvalho AF, Jacka F, Sarris J. The effects of dietary improvement on symptoms of depression and anxiety: a meta-analysis of randomized controlled trials. *Psychosomatic medicine*. 2019 Apr;81(3):265.
- [22] Shafiei F, Salari-Moghaddam A, Larijani B, Esmailzadeh A. Adherence to the Mediterranean diet and risk of depression: A systematic review and updated meta-analysis of observational studies. *Nutrition reviews*. 2019 Apr 1;77(4):230-9.
- [23] Neal EG, Chaffe H, Schwartz RH, Lawson MS, Edwards N, Fitzsimmons G, Whitney A, Cross JH. The ketogenic diet for the treatment of childhood epilepsy: a randomised controlled trial. *The Lancet Neurology*. 2008 Jun 1;7(6):500-6.
- [24] Morris AA. Cerebral ketone body metabolism. *Journal of inherited metabolic disease*. 2005 Apr;28(2):109-21.
- [25] Borghi L, Salvatici E, Riva E, Giovannini M, Vegni EA. Psychological and psychosocial implications for parenting a child with phenylketonuria: a systematic review. *Minerva pediatrica*. 2017 May 4;71(2):181-95.
- [26] Gaudio S, Wiemerslage L, Brooks SJ, Schiöth HB. A systematic review of resting-state functional-MRI studies in anorexia nervosa: evidence for functional connectivity impairment in

- cognitive control and visuospatial and body-signal integration. *Neuroscience & Biobehavioral Reviews*. 2016 Dec 1;71:578-89..
- [27] Giannunzio V, Degortes D, Tenconi E, Collantoni E, Solmi M, Santonastaso P, Favaro A. Decision-making impairment in anorexia nervosa: New insights into the role of age and decision-making style. *European Eating Disorders Review*. 2018 Jul;26(4):302-14.
- [28] Smith AD, Warren MJ, Refsum H. Vitamin B12. *Advances in food and nutrition research*. 2018 Jan 1;83:215-79.
- [29] Tangney CC, Aggarwal NT, Li H, Wilson RS, Evans DA, Morris MC. Vitamin B12, cognition, and brain magnetic resonance imaging measures.
- [30] Black MM. Effects of vitamin B12 and folate deficiency on brain development in children. *Food and nutrition bulletin*. 2008 Jun;29(2_suppl1):S126-31.
- [31] Enderami A, Zarghami M, Darvishi-Khezri H. The effects and potential mechanisms of folic acid on cognitive function: a comprehensive review. *Neurological Sciences*. 2018 Oct;39(10):1667-75.
- [32] Hegyi J, Schwartz RA, Hegyi V. Pellagra: dermatitis, dementia, and diarrhea. *International journal of dermatology*. 2004 Jan;43(1):1-5.
- [33] Brouwer-Brolsma EM, Dhonukshe-Rutten RA, van Wijngaarden JP, van de Zwaluw NL, in't Veld PH, Wins S, Swart KM, Enneman AW, Ham AC, van Dijk SC, van Schoor NM. Cognitive performance: a cross-sectional study on serum vitamin D and its interplay with glucose homeostasis in dutch older adults. *Journal of the American Medical Directors Association*. 2015 Jul 1;16(7):621-7.
- [34] Föcker M, Antel J, Ring S, Hahn D, Kanal Ö, Öztürk D, Hebebrand J, Libuda L. Vitamin D and mental health in children and adolescents. *European child & adolescent psychiatry*. 2017 Sep 1;26(9):1043-66.
- [35] Mohammadpour N, Jazayeri S, Tehrani-Doost M, Djalali M, Hosseini M, Effatpanah M, Davari-Ashtiani R, Karami E. Effect of vitamin D supplementation as adjunctive therapy to methylphenidate on ADHD symptoms: a randomized, double blind, placebo-controlled trial. *Nutritional neuroscience*. 2018 Mar 16;21(3):202-9.
- [36] Bazinet RP, Layé S. Polyunsaturated fatty acids and their metabolites in brain function and disease. *Nature Reviews Neuroscience*. 2014 Dec;15(12):771-85.
- [37] Firth J, Stubbs B, Teasdale SB, Ward PB, Veronese N, Shivappa N, Hebert JR, Berk M, Yung AR, Sarris J. Diet as a hot topic in psychiatry: a population-scale study of nutritional intake and inflammatory potential in severe mental illness. *World Psychiatry*. 2018 Oct;17(3):365.
- [38] McGrattan AM, McGuinness B, McKinley MC, Kee F, Passmore P, Woodside JV, McEvoy CT. Diet and inflammation in cognitive ageing and Alzheimer's disease. *Current nutrition reports*. 2019 Jun;8(2):53-65.
- [39] Pusceddu MM, Kelly P, Ariffin N, Cryan JF, Clarke G, Dinan TG. n-3 PUFAs have beneficial effects on anxiety and cognition in female rats: Effects of early life stress. *Psychoneuroendocrinology*. 2015 Aug 1;58:79-90.
- [40] Rapaport MH, Nierenberg AA, Schettler PJ, Kinkad B, Cardoos A, Walker R, Mischoulon D. Inflammation as a predictive biomarker for response to omega-3 fatty acids in major depressive disorder: a proof-of-concept study. *Molecular psychiatry*. 2016 Jan;21(1):71-9.
- [41] Su KP, Lai HC, Yang HT, Su WP, Peng CY, Chang JP, Chang HC, Pariante CM. Omega-3 fatty acids in the prevention of interferon-alpha-induced depression: results from a randomized, controlled trial. *Biological psychiatry*. 2014 Oct 1;76(7):559-66.
- [42] Delpuch JC, Madore C, Joffre C, Aubert A, Kang JX, Nadjar A, Layé S. Transgenic increase in n-3/n-6 fatty acid ratio protects against cognitive deficits induced by an immune challenge through decrease of neuroinflammation. *Neuropsychopharmacology*. 2015 Feb;40(3):525-36.
- [43] Labrousse VF, Nadjar A, Joffre C, Costes L, Aubert A, Gregoire S, Bretillon L, Laye S. Short-term long chain omega3 diet protects from neuroinflammatory processes and memory impairment in aged mice. *PLoS one*. 2012 May 25;7(5):e36861.
- [44] Larrieu T, Hilal LM, Fourrier C, de Smedt-Peyrusse V, Sans N, Capuron L, Laye S. Nutritional omega-3 modulates neuronal morphology in the prefrontal cortex along with depression-related behaviour through corticosterone secretion. *Translational psychiatry*. 2014 Sep;4(9):e437.
- [45] Lépinay AL, Larrieu T, Joffre C, Acar N, Gárate I, Castanon N, Ferreira G, Langelier B, Guesnet P, Brétilon L, Parnet P. Perinatal high-fat diet increases hippocampal vulnerability to the adverse effects of subsequent high-fat feeding. *Psychoneuroendocrinology*. 2015 Mar 1;53:82-93.
- [46] Zamroziewicz MK, Talukdar MT, Zwilling CE, Barbey AK. Nutritional status, brain network organization, and general intelligence. *Neuroimage*. 2017 Nov 1;161:241-50.
- [47] Nigg JT, Lewis K, Edinger T, Falk M. Meta-analysis of attention-deficit/hyperactivity disorder or attention-deficit/hyperactivity disorder symptoms, restriction diet, and synthetic food color additives. *Journal of the American Academy of Child & Adolescent Psychiatry*. 2012 Jan 1;51(1):86-97.
- [48] Bloch MH, Qawasmi A. Omega-3 fatty acid supplementation for the treatment of children with attention-deficit/hyperactivity disorder symptomatology: systematic review and meta-analysis. *Journal of the American Academy of Child & Adolescent Psychiatry*. 2011 Oct 1;50(10):991-1000.
- [49] Sonuga-Barke EJ, Brandeis D, Cortese S, Daley D, Ferrin M, Holtmann M, Stevenson J, Danckaerts M, Van der Oord S, Döpfner M, Dittmann RW. Nonpharmacological interventions for ADHD: systematic review and meta-analyses of randomized controlled trials of dietary and psychological treatments. *American Journal of Psychiatry*. 2013 Mar;170(3):275-89.
- [50] Del-Ponte B, Quinte GC, Cruz S, Grellert M, Santos IS. Dietary patterns and attention deficit/hyperactivity disorder (ADHD): a systematic review and meta-analysis. *Journal of affective disorders*. 2019 Jun 1;252:160-73.
- [51] Ly V, Bottelier M, Hoekstra PJ, Vasquez AA, Buitelaar JK, Rommelse NN. Elimination diets' efficacy and mechanisms in attention deficit hyperactivity disorder and autism spectrum disorder. *European child & adolescent psychiatry*. 2017 Sep 1;26(9):1067-79.
- [52] Jacka FN, O'Neil A, Opie R, Itsiopoulos C, Cotton S, Mohebbi M, Castle D, Dash S, Mihalopoulos C, Chatterton ML, Brazionis L. A randomised controlled trial of dietary improvement for adults

- with major depression (the 'SMILES' trial). *BMC medicine*. 2017 Dec;15(1):1-3.
- [53] Parletta N, Zarnowiecki D, Cho J, Wilson A, Bogomolova S, Villani A, Itsiopoulos C, Niyonsenga T, Blunden S, Meyer B, Segal L. A Mediterranean-style dietary intervention supplemented with fish oil improves diet quality and mental health in people with depression: A randomized controlled trial (HELFI-MED). *Nutritional neuroscience*. 2019 Jul 3;22(7):474-87.
- [54] Sanchez-Villegas A, Cabrera-Suárez B, Molero P, Gonzalez-Pinto A, Chiclana-Actis C, Cabrera C, Lahortiga-Ramos F, Florido-Rodríguez M, Vega-Pérez P, Vega-Pérez R, Pla J. Preventing the recurrence of depression with a Mediterranean diet supplemented with extra-virgin olive oil. The PREDI-DEP trial: study protocol. *BMC psychiatry*. 2019 Dec;19(1):1-7.
- [55] Berk M, Jacka FN. Diet and Depression—From Confirmation to Implementation. *Jama*. 2019 Mar 5;321(9):842-3.
- [56] Bot M, Brouwer IA, Roca M, Kohls E, Penninx BW, Watkins E, Van Grootheest G, Cabout M, Hegerl U, Gili M, Owens M. Effect of multivitamin supplementation and food-related behavioral activation therapy on prevention of major depressive disorder among overweight or obese adults with subsyndromal depressive symptoms: the MooDFOOD randomized clinical trial. *Jama*. 2019 Mar 5;321(9):858-68.
- [57] Gibson-Smith D, Bot M, Brouwer IA, Visser M, Giltay EJ, Penninx BW. Association of food groups with depression and anxiety disorders. *European journal of nutrition*. 2020 Mar;59(2):767-78.
- [58] Dinan TG, Stanton C, Long-Smith C, Kennedy P, Cryan JF, Cowan CS, Cénit MC, van der Kamp JW, Sanz Y. Feeding melancholic microbes: MyNewGut recommendations on diet and mood. *Clinical Nutrition*. 2019 Oct 1;38(5):1995-2001.
- [59] Fernandez-Real JM, Serino M, Blasco G, Puig J, Daunis-i-Estadella J, Ricart W, Burcelin R, Fernández-Aranda F, Portero-Otin M. Gut microbiota interacts with brain microstructure and function. *The Journal of Clinical Endocrinology & Metabolism*. 2015 Dec 1;100(12):4505-13.
- [60] Wang S, Harvey L, Martin R, van der Beek EM, Knol J, Cryan JF, Renes IB. Targeting the gut microbiota to influence brain development and function in early life. *Neuroscience & Biobehavioral Reviews*. 2018 Dec 1;95:191-201.
- [61] Cunnane SC, Crawford MA. Energetic and nutritional constraints on infant brain development: implications for brain expansion during human evolution. *Journal of human evolution*. 2014 Dec 1;77:88-98.
- [62] Innis SM. Dietary omega 3 fatty acids and the developing brain. *Brain research*. 2008 Oct 27;1237:35-43.
- [63] Laus MF, Duarte Manhas Ferreira Vales L, Braga Costa TM, Sousa Almeida S. Early postnatal protein-calorie malnutrition and cognition: a review of human and animal studies. *International Journal of Environmental Research and Public Health*. 2011 Feb;8(2):590-612.
- [64] McNamara RK, Carlson SE. Role of omega-3 fatty acids in brain development and function: potential implications for the pathogenesis and prevention of psychopathology. *Prostaglandins, Leukotrienes and Essential Fatty Acids*. 2006 Oct 1;75(4-5):329-49.
- [65] Prado EL, Dewey KG. Nutrition and brain development in early life. *Nutrition reviews*. 2014 Apr 1;72(4):267-84.
- [66] Schwarzenberg SJ, Georgieff MK, Daniels S, Corkins M, Golden NH, Kim JH, Lindsey C, Magge SN. Advocacy for improving nutrition in the first 1000 days to support childhood development and adult health. *Pediatrics*. 2018 Feb 1;141(2).
- [67] Castanys-Muñoz E, Kennedy K, Castañeda-Gutiérrez E, Forsyth S, Godfrey KM, Koletzko B, Ozanne SE, Rueda R, Schoemaker M, van der Beek EM, van Buuren S. Systematic review indicates postnatal growth in term infants born small-for-gestational-age being associated with later neurocognitive and metabolic outcomes. *Acta Paediatrica*. 2017 Aug;106(8):1230-8.
- [68] Ong KK, Kennedy K, Castañeda-Gutiérrez E, Forsyth S, Godfrey KM, Koletzko B, Latulippe ME, Ozanne SE, Rueda R, Schoemaker MH, Van Der Beek EM. Postnatal growth in preterm infants and later health outcomes: a systematic review. *Acta paediatrica*. 2015 Oct;104(10):974-86.
- [69] Georgieff MK, Ramel SE, Cusick SE. Nutritional influences on brain development. *Acta Paediatrica*. 2018 Aug;107(8):1310-21.
- [70] Velasco I, Bath SC, Rayman MP. Iodine as essential nutrient during the first 1000 days of life. *Nutrients*. 2018 Mar;10(3):290.
- [71] Algarin C, Karunakaran KD, Reyes S, Morales C, Lozoff B, Peirano P, Biswal B. Differences on brain connectivity in adulthood are present in subjects with iron deficiency anemia in infancy. *Frontiers in aging neuroscience*. 2017 Mar 7;9:54.
- [72] Oosting A, Verkade HJ, Kegler D, van de Heijning BJ, van der Beek EM. Rapid and selective manipulation of milk fatty acid composition in mice through the maternal diet during lactation. *Journal of nutritional science*. 2015;4.
- [73] Freedman R, Hunter SK, Hoffman MC. Prenatal primary prevention of mental illness by micronutrient supplements in pregnancy. *American Journal of Psychiatry*. 2018 Jul 1;175(7):607-19.
- [74] Schipper L, Oosting A, Scheurink AJ, van Dijk G, van der Beek EM. Reducing dietary intake of linoleic acid of mouse dams during lactation increases offspring brain n-3 LCPUFA content. *Prostaglandins, Leukotrienes and Essential Fatty Acids*. 2016 Jul 1;110:8-15.
- [75] Yam KY, Schipper L, Reemst K, Ruigrok SR, Abbink MR, Hoeijmakers L, Naninck EF, Zarekiani P, Oosting A, Van Der Beek EM, Lucassen PJ. Increasing availability of ω -3 fatty acid in the early-life diet prevents the early-life stress-induced cognitive impairments without affecting metabolic alterations. *The FASEB Journal*. 2019 Apr;33(4):5729-40.
- [76] Provensi G, Schmidt SD, Boehme M, Bastiaanssen TF, Rani B, Costa A, Fouhy F, Strain C, Stanton C, Blandina P, Izquierdo I. Preventing adolescent stress-induced cognitive and microbiome changes by diet. *Proceedings of the National Academy of Sciences*. 2019 May 7;116(19):9644-51.
- [77] Donoso F, Egerton S, Bastiaanssen TF, Fitzgerald P, Gite S, Fouhy F, Ross RP, Stanton C, Dinan TG, Cryan JF. Polyphenols selectively reverse early-life stress-induced behavioural, neurochemical and microbiota changes in the rat. *Psychoneuroendocrinology*. 2020 Jun 1;116:104673.
- [78] Hu FB. Dietary pattern analysis: a new direction in nutritional epidemiology. *Current opinion in lipidology*. 2002 Feb 1;13(1):3-9.
- [79] Sarris J. Nutritional psychiatry: from concept to the clinic. *Drugs*. 2019 Jun;79(9):929-34.