

# Free will – an approach from the perspective of neuroscience

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## ABSTRACT

*Free will designates the possibility of conscious and free choice of the subject, as well as his real possibility to act. The topic has a special medical and social relevance as it is directly connected to how the subject's responsibility (moral, legal, medical) for his own actions are understood, but also the possibility (real or illusory) of intervention in his own life.*

*Free will has received various philosophical, psychological, legal and interdisciplinary approaches in recent decades. In this article we shall inventory some results from the area of neuroscience that we consider relevant in the analysis of free will, research and clinical studies that highlight various effects caused by voluntary actions, consciously repeated by the subject. Some psychotherapeutic interventions, the process of awareness, learning, attention monitoring, visualization, some forms of meditation, and the changes that these practices produce in the processes of neurogenesis, neuroplasticity and epigenetics, in terms of health and quality of life are analysed.*

*Since these interventions involve the conscious participation of the subject and his voluntary action, we consider that the discussed results are relevant in the debate on free will. The study also highlights the importance of education and social knowledge, as well as the need to promote participatory prevention and therapy, involving the subject in increasing the quality of life and health, through public messages that emphasize the responsibility of each person for their own choices and actions and the impact they have on his life.*

**Keywords:** free will, conscious participation, neuroplasticity, neurogenesis, epigenetics, placebo, mindfulness, visualization (mental imaging), the impact of spiritual life on health

## FREE WILL – SHORT DISCIPLINARY AND INTERDISCIPLINARY THEMATIC INVENTORY

Since the way we understand free will corresponds to real or illusory capacities to decide and act, the topic is of particular importance from an epistemological and practical perspective. The concept has a long philosophical history, starting with Greek antiquity [1, 2]. In

recent decades, the concept of free will has been analysed in relation to various concepts, such as the moral [3] and legal [4] responsibility of the subject, but also in connection to educational policies [5] and character cultivation [6], with the performed work [7] and academic excellence [8], being also approached in the evaluation of cultural creations [9]. There have also been approaches to free will as a decision-making process [10,11], in close connection with intentionality

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Article History:

Received: 9 September 2021

Accepted: 15 September 2021

[12] and the voluntary act [13], but also in connection with other concepts such as determinism, indeterminism and the possibility of choice [14] or with reference to issues concerning consciousness [15] and conscience [16].

In addition to the inevitable ethical discussions about free will [17], there are genetic [18], clinical [19] and interdisciplinary explorations, such as those that juxtapose psychological, neuro-ethical and moral theology radiographs [20]. Finally, there is no shortage of speculative, borderline discussions that place free will in the context of quantum mechanics [21, 22] or artificial intelligence [23], or those that analyse the conditions of possibility for the existence of free will among invertebrates [24].

The PhilPapers philosophical index is evidence of the multitude of perspectives on the analysis of free will in the field of science. Made by a digital community of philosophy specialists, PhilPapers monitors sources in the field of philosophy, including texts from periodicals, journals, books and open access archives, but also from personal web pages belonging to specialists, researchers or academics. PhilPapers represents the largest digital archive of texts in the field of philosophy, with free access to the general public. At the time this article was written (April 15, 2021), PhilPapers contained 2,570,856 classified entries, organized into 5,674 categories, with over 280,000 registered users [25]. In the subcategory "free will and science", this digital inventory offered no fewer than 1016 distinct titles; of these, 13 concern analyses of free will in connection with genetics, 351 in connection with neuroscience, 109 with physics, 426 with psychology, and 103 with other sciences.

Also, the topicality and complexity of the topic can be seen in the numerous attempts to systematize or evaluate free will, recorded in recent years [26-35].

## FREE WILL AND INDIVIDUAL AND SOCIAL RELEVANCE

In this vast territory of free will approaches, there are many perspectives. Some of them deny the reality of free will [36], others openly support it [37]. Beyond the many challenging and complex aspects it entails, the topic is important in terms of personal and social life [38]. There are many reasons that support this fact: the need to clarify the moral and legal responsibility for our actions, the real possibility to choose between various options, to be reasonably considered the authors of our own choices and deeds, conscious participation in decisions and actions which are assigned to us socially or in terms of lifestyle [39].

At an individual level, the way free will is understood provides a representation of the real or illusory

ability to change the patterns of one's life, to replace unhealthy habits with sanogenic practices. Debating one model or another of understanding free will, scientific information can increase the subject's responsibility for his own choices and actions or, on the contrary, it can relieve him of this responsibility.

One study highlights this influence of scientific discourse on free will in subjects' self-report of addiction [40]. The scientific information supporting a diminution of free will in the case of alcohol-dependent subjects is noted; by emphasizing the neurological imbalance and deterministic aspects of addiction, they indirectly support a decrease in the possibilities available to addicted subjects of the choices they make and of their actions. On the other hand, ethical or moral approaches to alcoholism present the behaviours of alcoholics as the result of their choices, the evolution of the situation still depending on the decisions of the subjects. After noting these differences at the level of public discourse, the study emphasizes that scientific information about the real or illusory possibility of choosing behaviours can change the self-perception of addictive subjects. A discourse that minimizes the possibilities of conscious and free choices and voluntary actions can intensify the self-deprecation and stigmatization of addictive people, fuelled by the awareness of their own deficiencies in decision-making, by the acceptance of the lack of self-control [40].

In our opinion, in addition to the research aiming at conceptual aspects or theoretical, legal, psychological or interdisciplinary models, free will can also be evaluated through a series of results and discoveries in the field of clinical neuroscience. We refer here to those that concern neuroplasticity, neurogenesis or epigenetics, but also to those that take into account the placebo phenomenon, the changes induced by cognitive-behavioural psychotherapy and the experience of spiritual life.

## SELF-INDUCED NEUROPLASTICITY

Neuroplasticity targets a wide spectrum of processes, being highlighted in some mammalian species [41] and in humans [42]. It accompanies the usual developmental stages of a healthy brain, but it also includes the adaptive or maladaptive structural and functional changes in brain tissue that occur throughout life as a result of experiences (sensory input), being also responsible for changes in reception and processing of experience data [43].

Neuroplasticity is present today in a wide spectrum of therapeutic interventions, from those highlighted by Edward Taub, in post stroke recovery [44], to those aiming to identify molecular targets to improve plasticity in neurodegenerative diseases [45]. Regarding

synaptic plasticity, for example, PubMed indicates a total of more than 66,000 entries since 2009, with more than 3000 articles registered annually [46]. Research and clinical studies that have explored neuroplasticity show encouraging results in Alzheimer's disease, in the case of spinal cord injuries, in apraxia or in post-stroke motor rehabilitation [47]. In another register, the adoption of sanogenic behaviours, such as regular physical exercise and healthy eating, improves neural adaptation processes [48].

In various forms, in situations of this kind, at the core of the intervention, there is the conscious action of the subject, the observable neural effects being, most often, the result of a voluntarily repeated exercise. For example, adopting healthy behaviours helps increase the production of brain-derived neurotrophic factor (BDNF). It is well known that BDNF levels influence cognitive function and that in some conditions, such as severe depression [49,50], Alzheimer's [51] or type II diabetes [52], BDNF levels are low. The impact of daily practice is evident in sports, as moderate physical exercise, even for as little as 30 minutes, has effects on BDNF levels [53], contributing to increased selective attention [54] and learning abilities [55], also causing structural brain changes [56].

## INTERVENTIONS FOR NEUROGENESIS STIMULATION

In the matter of free will, the findings that highlight the influence of voluntary actions and behaviours on neurogenesis are also relevant. Located in the hippocampus and dentate gyrus [57], the processes of new neuron formation by differentiation of neural stem cells have also been highlighted in adults [58]. Neurogenesis is of particular importance, having a regenerative potential [59]. And in the case of these processes, there are situations where the will, involvement and conscious action of the subjects prove decisive.

Some studies show that choosing a social environment [60] or diversifying experiences and enriching the living environment can stimulate neurogenesis [61]. Other examples are provided by studies targeting cognitive-behavioural interventions to improve the stress response. Effects of stress on cellular processes, dendritic architecture and growth factor levels are known [62]. The impact of stress is extensive, affecting the transmission of signals in the nervous system and its connections, the processes of neurogenesis [63] and neuroplasticity [64], but also the general clinical picture [65]. Cognitive-behavioural interventions to improve the stress response [66] are relevant in the discussion about free will, because they produce a change through a set of actions in which patients participate consciously. For example, intermittent fasting [67], ex-

ercise [68, 69] and diet [70] can stimulate processes of neurogenesis. Also, there are some studies that announce the beneficial impact of coping strategies in the regulation of neurogenesis processes, a fact first highlighted in monkeys [71], but also discussed in humans [72].

## SELF-DIRECTED EPIGENETICS

The field of epigenetics provides results worth considering in the free will debate for the same reason. It is about the methylation/demethylation processes of DNA, the acetylation/deacetylation of proteins (mainly histones), the activity of microRNAs (miRNAs) that accompany the modification of gene expressions and the functions that these genes encode. In these cases as well, there is evidence regarding self-induced plasticities, through conscious effort, through voluntary actions.

An example in this case is provided by the studies evaluating the improvement of the epigenetic effects of stress through cognitive-behavioural interventions. It is known that exposure to stress in the early period of life can produce changes in gene expression, at the level of methylation processes [73, 74]. Some studies highlight how maternal behaviour and the social environment influence the stress response in mice [75] and how attachment can ameliorate the epigenetics of the stress response in humans, in young adults [76]. Cognitive reframing (cognitive reappraisal), cognitive-behavioural psychotherapy [77] or other psychotherapeutic interventions [78], but also simple behavioural changes, such as choosing a sanogenic living environment and a healthy diet [79] or regular physical exercise [80], can significantly influence gene expression. It is about the regulation of genetic expressions (down-shifting or up-shifting) that modify correlative physiological mechanisms, improving the quality of life [81]. In particular, changes in life patterns can influence the decrease in pro-inflammatory expression and even the conserved transcriptional response to adversity – which includes the expression of the genes involved in the inflammatory and infectious response [82], a fact that contributes to improving health [83].

In the discussion of free will, studies that signal the influence of chronic stress [84] on telomere plasticity [85] and thus the impact on lifespan are also significant. And in this case, therapeutic interventions to improve the stress response prove effective. It is not only about the early periods of life, when telomere plasticity seems to be under the influence of maternal attachment [86], but also about conscious actions, voluntarily repeated by adult patients, actions that can produce significant changes. The method of multisystem resilience (multisystem resiliency) [87], for example, aims

at emotional regulation, strengthening social connections and adopting healthy behaviours, these beneficially influencing telomere plasticity. One study showed that the intervention applied to a group of adults, aged 45 - 90 years, caused a slight increase in telomere length and a decrease in the intensity of the depressive episodes [88]. Other studies show a telomere plasticity in adulthood through healthy routines [89-92].

Along with epigenetics, new processes of epigenetic-transgenerational nature, which can interfere with the decision-making process [93], feeding deterministic perspectives at the scientific level and at the level of the general public, are also under discussion [94]. However, we believe that the mentioned studies convincingly show that we can increase the conscious and active involvement in the preservation of one's own health and that there are ways of prevention and possibilities of therapeutic intervention directed to epigenetic responses [95].

## **SPONTANEOUS REMISSIONS, PLACEBOS AND PERSONAL FACTORS**

The study of the placebo phenomenon and those that follow spontaneous remissions provide arguments in favour of free will, arguments which claim that our own choices and actions influence our lives and health.

The placebo phenomenon [96] has registered an increased presence in the scientific approaches of the last decades [97]. Studies show varying degrees of effectiveness of this effect in improving stress response and post-surgery recovery [98], in migraines [99] and anxiety [100], in rheumatoid arthritis [101] and in carpal tunnel syndrome [102], in fibromyalgia [103], in cases of allergies [104], chronic pain [105] and duodenal ulcer [106], but also in depression [107] and cancer [108]. For example, a meta-analysis indicates a placebo efficacy rate of 29%, compared to 36% in the case of classical medical interventions [109].

In recent years, approaches that propose genetic investigations of placebo-responders have emerged. It is about the area called *placebome*, which inventories the genomic particularities of the patients who respond to placebo and the therapeutic possibilities that arise [110]. One of the acknowledged authors in this field warned, at the end of an older study, that the existence of the placebo effect suggests the expansion of "the conception of human capabilities" [111].

We include here the growing scientific interest in spontaneous remissions as well. It is about some case studies [112-115] and meta-analyses [116-119], which affirm the need for scientific clarification studies regarding the role of subjective aspects, such as prayer [120] or other psychological factors [121] in the process of spontaneous remission.

Studies on the effectiveness of the placebo effect or on the conditions of possibility for spontaneous remission are worth considering in the evaluation of free will, since we are dealing with a process in which compliance, belief in the beneficial effect of the treatment [122], higher organized beliefs are decisive, data of knowledge [123], faith and religious practice [124].

## **ATTENTION SELF-CONTROL, LEARNING AND AWARENESS**

Those studies (evidence-based medicine) regarding cerebral changes determined by the conscious direction of thoughts are also relevant in the inventory of therapeutic possibilities capable of producing self-induced plasticity. For example, proper mental training through deliberate effort can affect pathological brain circuits, causing adaptive alteration. This is the case of some successes obtained through therapeutic interventions in the treatment of OCD (obsessive compulsive disorder) [125], in phobias, anxieties or in post-traumatic syndrome [126].

Therapeutic interventions aimed at controlling conscious mental activity consist in self- or refocusing techniques, methods of increasing awareness [127] or acquiring coping strategies [128], the learning effort being the main vector in increasing resilience and functional neuroadaptation. Basically, the meaning associated with an action reconfigures the effects of that action on the acting subject, to the extent that he is aware of that meaning. For example, one study showed that informing some housekeepers about the beneficial effects of their activities increased the beneficial impact of their work on their health, compared to another group of housekeepers who performed similar activities but were not informed about the sanogenic impact of their work [129]. The subject's conscious participation in the activities they carry out, a mental way to relate to his own actions (mind-set), comes into question in this case as well.

Other studies, in which subjects are asked to increase awareness of an action or learn a new movement, also have relevance to the discussion of free will. For example, the neural map correlative to a visual awareness task has been found to be more extensive, including higher brain centres, in contrast to the brain footprint corresponding to a visual attention task [130,131]. In another study, we find that the brain imprint corresponding to the learning efforts to learn a new movement (accompanied by a conscious effort) differs from the active one in the automatic performance of the same movement [132]. These results bring into discussion learning as a conscious effort to retain some information or to acquire some movements and the qualitative difference that the aware-

ness effort produces at the level of brain networks [133]. Thus, some particularities have also been identified in the case of mathematicians [134] or those who practice meditation [135].

On the other hand, by consciously directing attention to certain stimuli, the subject influences his own perceptions and neural processing of signals. Intentional focusing of attention on certain stimuli in the sensory field causes differential processing of selected signals (event), relative to all other signals (background noise). Voluntary attention modulates neural communication (top-down), causing an increase in the efficiency of processing presynaptic input and postsynaptic response, in the case of the tracked signal, and a decrease in efficiency in the processing of redundant signals, a fact that allows improved reception of selected stimuli (event) relative to background noise [136].

Furthermore, attentively following images can produce measurable physiological effects, with mirror neurons as a possible explanation [137]. Some studies have reported an increase in finger length through training to view suggestive materials [138], but also an improvement in the condition of stroke patients [139] when they watched video materials with specially adapted content.

### **SELF-INDUCED PLASTICITY THROUGH MENTAL IMAGING**

The studies highlighting the impact of thinking and mental imagery on brain functioning hold an important place in the discussion of self-induced plasticity. One of the first studies that inaugurated this field of work proved that merely practising an action mentally produces neural changes similar to those caused by actually performing the action [140,141]. Following this result, other studies have shown the importance of thoughts and mental images on physiology. The mental representation of a movement of the fingers and toes, as well as the tongue, produces activity in the brain regions that govern them [142], and directed imaginal training produces an increase in physical strength [143,144].

Mental visualization (or mental imaging) has been tested with encouraging results in various conditions, with notable results in the healing of post-surgery wounds [145] and the reduction of pain in fibromyalgia [146], in post stroke recovery [147] and in relieving pain caused by cystitis [148], in rehabilitation after a spinal cord injury [149]. There are notable results of mental imaging therapy, showing improvements in the condition of patients with asthma [150] and Parkinson's [151], decreases in stress levels and recurrence conditions in breast cancer [152], improvement in the quality of life of patients with osteoarthritis [153], but

also strengthening the state of calm in patients with lung diseases [154].

In these cases, we are dealing with patients who cultivate specific mental images, obtaining significant improvements, a fact that indicates the important impact of thoughts and mental representations on health and life.

### **MINDFULNESS, RELIGIOUS AND SPIRITUAL EXPERIENCES**

The possibility of real interventions likely to produce significant changes in health and life is also supported by studies evaluating mental practices and spiritual experiences.

In the last decade, there have been neuroscientific investigations on various forms of mental self-control, meditative practices such as relaxation response and mindfulness [155], mindfulness based stress reduction – MBSR [156], mind-body therapy [157] and their possible health effects. The results are significant in the treatment of post-traumatic syndrome [158], anxiety [159] and depression [160], in the regulation of cortisol levels [161], in the regulation of blood glucose levels, in patients with type 2 diabetes [162], in the potentiation of neurogenesis [163], but also post transplantation, reducing depression and anxiety and improving the quality of patients' sleep [164].

A study showed the effectiveness of an 8-week mindfulness-based stress reduction program (MBSR) in 49 breast cancer patients and 10 prostate cancer patients, all at an early stage. The program included relaxation periods, meditation techniques practiced daily at home. Measurements, made before and after surgery, revealed changes in endocrine, immune and autonomic parameters, salivary cortisol levels, immune cell counts, blood pressure and heart rate [165].

Other results point out that the MBSR program can contribute to the improvement of the quality of life [166], causing a reduction in the level of stress in patients with various forms of cancer [167]. In some cases, the effectiveness of the intervention is rapid, if we take into account the conclusions of another study, which highlighted, after only 8 hours of meditation practice, changes in the inflammatory mechanisms (decrease in the level of interleukin-8, a pro-inflammatory cytokine) [168].

Numerous other results refer to other spiritual practices. We have in mind here the “Mother Teresa Effect”, highlighted by David McClelland [169], which signals the change of the immune response as a result of an experience which has a spiritual content, but also the therapeutic effects of compassion [170-172]. In this case, the studies that identify distinct brain imprints and psycho-emotional effects from various

spiritual or religious practices, highlighted by Andrew Newberg in neurotheology research [173,174], are also relevant.

The conclusions of the meta-analyses carried out by Harold Koenig, in the studies regarding the therapeutic effects of religious or spiritual life, are also remarkable. In one of these large meta-analyses, evaluating 1200 medical studies published between 1872 and 2000, and 2100 studies published between 2000 and 2010, strong correlations between religion/spirituality and various aspects of health are highlighted [175]. Operating a rigorous selection that took into account criteria of scientific probity, experimental design and correctness of data analysis, the meta-analysis provided a scientific x-ray of the relationship between religious life and health, at the level of 2010. The conclusions of the meta-analysis indicate a significant beneficial impact of spiritual and religious life practices in coronary diseases, arterial hypertension, cerebral-vascular diseases, Alzheimer's disease and dementia, in the regulation of immune function and endocrine function, in the evolution of cancer, in the quality of mobility (mobility and capacities physical) and health and life expectancy [175]. In addition, spiritual life has a beneficial impact in shaping the response to stress, in cultivating positive emotions, in well-being, in cultivating hope and optimism, in strengthening the sense of a meaningful life, in increasing self-esteem, in positive character traits, at the same time reducing the incidence of depression and suicide, anxiety and substance abuse, delinquency and marital instability [175].

Another meta-analysis, led by the same physician [176], assessed the link between spiritual/religious life (R/S) and mental health captured by scientific studies published in specialized journals over a 20-year period between 1990 and 2010. Original research on religion, religiosity, spirituality and related terms published in the top 25% of psychiatry and neurology journals, according to the 2010 ISI index rankings, was examined. In percentage terms, 72.1% of the analysed research showed a positive relationship between the level of re-

ligious/spiritual involvement and a lower presence of mental disorders. All studies on dementia, suicide and stress disorders found a positive association of R/S, 79% of them found that R/S correlated with lower depression and 67% of them with lower substance abuse [177].

Other approaches go even further, forcing new territories, targeting possible effects of mental activity on neurogenesis and DNA expression [178] or the epigenetic impact of the relaxation response [179]. A significant example is a study of the changes brought about by a meditative practice that focused on the meaning of life. The practitioners recorded - after 6 hours of daily meditation, during 3 months spent in retreat - important changes in immune cells, a decrease in negative affectivity, increased telomerase activity, increased telomere length and an increased longevity of immune cells [180].

Even though the detailed analysis of such results is just beginning [181-183], we believe that studies of this kind remain significant for discussions of the real possibilities of choice and life change, discussed with reference to free will.

## CONCLUSIONS

Debates about free will bring into question various philosophical and psychological concepts and nuances. In addition to these considerations, clinical studies and experimental research in the field of neuroscience provide significant results that deserve attention in the evaluation of free will. The stake is precious, as we are dealing with the possibility of deliberate, conscious intervention in the plan of personal life. The briefly inventoried research indicates various possibilities of involving the subject, which can alleviate suffering, contribute to the recovery or development of some skills, can stimulate positive emotionality and can change patterns of thought and action, contributing considerably to the improvement of the quality of life.

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## REFERENCES

1. Frede M. A Free Will: Origins of the Notion in Ancient Thought, Long AA (ed) series: Sather Classical Lectures. University of California Press; 2011.
2. Cary P. A brief history of the concept of free will: issues that are and are not germane to legal reasoning. *Behav Sci Law*. 2007;25(2):165-81.
3. Levy N, McKenna M. Recent Work on Free Will and Moral Responsibility. *Philos Compass*. 2009 Jan 21;4(1):96-133.
4. Kolber AJ. Free Will as a Matter of Law in Philosophical Foundations of Law and Neuroscience, Patterson D, Pardo MS (eds.). Oxford Scholarship Online; 2016.
5. Giesinger J. Free Will and Education. *J Philos Educ*. 2010 Nov;44(4):515-28.
6. Danto AC, Morgenbesser S. Character and Free will. *J Philos*. 1957 Aug; 54(16):493-505.
7. Stillman TF, Baumeister RF, Vohs KD, Lambert NM, Fincham FD, Brewer LE. Personal Philosophy and Personnel Achievement: Belief in Free Will Predicts Better Job Performance. *Soc Psychol Personal Sci*. 2010;1(1):43-50.
8. Feldman G, Chandrashekar P, Wong KFE. The Freedom to Excel: Belief in Free Will Predicts Better Academic Performance. *Pers Individ Dif*. 2016 Feb 90:377-83.
9. Bartra R. Anthropology of the Brain: Consciousness, Culture, and Free Will, Cambridge University Press; 2014.

10. Yoder KJ, Decety J. The neuroscience of morality and social decision-making. *Psychol Crime Law*. 2018;24(3):279-295.
11. Burns K, Bechara A. Decision making and free will: a neuroscience perspective. *Behav Sci Law*. 2007;25(2):263-80.
12. Leisman G, Macahdo C, Melillo R, Mualem R. Intentionality and „free-will“ from a neurodevelopmental perspective. *Frontiers in Integrative Neuroscience*. 2012;6:36.
13. Libet B. Unconscious cerebral initiative and the role of conscious will in voluntary action. *Behav Brain Sci*. 1985;8(4):529-539.
14. Atmanspacher H, Bishop R. Between Chance and Choice. *Interdisciplinary Perspectives on Determinism*. Imprint Academic Thorverton; 2002.
15. Shepherd J. Free will and consciousness: experimental studies. *Conscious Cogn*. 2012 Jun;21(2):915-27.
16. Lavazza A. Free Will and Neuroscience: From Explaining Freedom Away to New Ways of Operationalizing and Measuring It. *Front Hum Neurosci*. 2016 Jun 1;10:262.
17. Zürcher T, Elger B, Trachsel M. The notion of free will and its ethical relevance for decision-making capacity. *BMC Med Ethics*. 2019 May 8;20(1):31.
18. Alper JS. Genes, free will, and criminal responsibility. *Soc Sci Med*. 1998 Jun;46(12):1599-611.
19. Meynen G. Free will and mental disorder: exploring the relationship. *Theor Med Bioeth*. 2010 Dec;31(6):429-43.
20. Dodson GF. Free Will. *Neuroethics, Psychology and Theology*. *Vernon Series in Philosophy*. Hardcover; 2017.
21. Stapp HP. Philosophy of Mind and the Problem of Free Will in the Light of Quantum Mechanics. internet], citat 2021 Sep 8]. Available at: <https://arxiv.org/ftp/arxiv/papers/0805/0805.0116.pdf>.
22. Stapp HP. Quantum Theory and Free Will: How Mental Intentions Translate into Bodily Actions. Springer International; 2017.
23. Krausová A, Hazan H. Creating Free Will in Artificial Intelligence. Beyond AI: Artificial Golem Intelligence. *Proc Int Conf Beyond AI*. Pilsen. Czech Republic. *University of Bohemia*; 2013 November 12-14: 96-109.
24. Brembs B. Towards a scientific concept of free will as a biological trait: spontaneous actions and decision-making in invertebrates. *Proc Biol Sci*. 2011 Mar 22;278(1707):930-9.
25. PhilPapers. Available at: <https://philpapers.org/browse/free-will-and-science>.
26. Dennett DC, E Room. *The Varieties of the Free Will Worth Wanting*. MIT Press; 1984.
27. Kane R. *The Significance of Free Will*. Oxford University Press; 1996.
28. *Responsibility and the Moral Sentiments*. Harvard University Press; 1998.
29. Dilman I. *Free Will: An Historical and Philosophical Introduction*. Routledge; 1999.
30. Kane R. *A Contemporary Introduction to Free Will*. Oxford University Press; 2005.
31. Fischer JM, Kane R, Pereboom D, Vargas M. Four Views on Free Will. (Great Debates in Philosophy). *Blackwell*; 2014.
32. Waller BN. *Against Moral Responsibility*. MIT Press; 2011.
33. Kane R. *The Oxford Handbook of Free Will* (second edition). Oxford University Press; 2011.
34. Iredale M. *The Problem of Free Will: A Contemporary Introduction*. Routledge; 2012.
35. McKenna M, Pereboom D, *Free Will: A Contemporary Introduction*. Routledge; 2014.
36. Wegner DM. *The Illusion of Conscious Will*. The MIT Press; 2002.
37. List C. *Why Free Will Is Real*. Harvard University Press; 2019.
38. Kane R. *A Contemporary Introduction to Free Will*. New York. NY: Oxford University Press; 2005.
39. Lavazza A. Free Will and Neuroscience: From Explaining Freedom Away to New Ways of Operationalizing and Measuring It. *Front Hum Neurosci*. 2016 Jun 1;10:262.
40. Racine, E, Bell, E, Zizzo, N, Green, C. Public discourse on the biology of alcohol addiction: implications for stigma, self-control, essentialism, and coercive policies in pregnancy. *Neuroethics*. 2015;8:177-86.
41. La Rosa C, Bonfanti L. Brain Plasticity in Mammals: An Example for the Role of Comparative Medicine in the Neurosciences. *Frontiers in veterinary science*. 2018 Nov 1;5:274.
42. Zoladz JA, Pilc A. The effect of physical activity on the brain derived neurotrophic factor: from animal to human studies. *J Physiol Pharmacol*. 2010 Oct;61(5):533-41.
43. Rivera SM, Carlson SM, Zelazo P. Introduction to Special Issue: Current Perspectives on Neuroplasticity, *Cogn Dev*. 2017; 42:1-3.
44. Taub E, Miller NE, Novack TA, Cook EW 3rd, Fleming WC, Nemonuceno CS, Connell JS, Crago JE. Technique to improve chronic motor deficit after stroke. *Arch Phys Med Rehabil*. 1993 Apr; 74(4):347-54.
45. Kozubski W, Ong K, Waleszczyk W, Zabel M, Dorszewska J. Molecular Factors Mediating Neural Cell Plasticity Changes in Dementia Brain Diseases. *Neural Plast*. 2021 Mar 29;2021:8834645.
46. National Library of Medicine. Synaptic plasticity. Available at: <https://pubmed.ncbi.nlm.nih.gov/?term=synaptic%20plasticity&sort=date&timeline=expanded>.
47. Baudry M, Bi X., Schreiber SS. Synaptic Plasticity. *Basic Mechanisms to Clinical Applications*. CRC Press, 2005:229-80.
48. Peterson JC. The adaptive neuroplasticity hypothesis of behavioral maintenance. *Neural Plast*. 2012;2012:516364.
49. Lang UE, Hellweg R, Gallinat J. BDNF serum concentrations in healthy volunteers are associated with depression-related personality traits. *Neuropsychopharmacology*. 2004 Apr;29(4):795-8.
50. Emon MPZ, Das R, Nishuty NL, Shalahuiddin Qusar MMA, Bhuiyan MA, Islam MR. Reduced serum BDNF levels are associated with the increased risk for developing MDD: a case-control study with or without antidepressant therapy. *BMC Res Notes*. 2020 Feb 21;13(1):83.
51. Connor B, Young D, Yan Q, Faull RL, Synek B, Dragunow M. Brain-derived neurotrophic factor is reduced in Alzheimer's disease. *Brain Res Mol Brain Res*. 1997 Oct 3;49(1-2):71-81.
52. Krabbe KS, Nielsen AR, Krogh-Madsen R, Plomgaard P, Rasmussen P, Erikstrup C, Fischer CP, Lindegaard B, Petersen AM, Taudorf S, Secher NH, Pilegaard H, Bruunsgaard H, Pedersen BK. Brain-derived neurotrophic factor (BDNF) and type 2 diabetes. *Diabetologia*. 2007 Feb;50(2):431-8.
53. Gold SM, Schulz KH, Hartmann S, Mladek M, Lang UE, Hellweg R, Reer R, Braumann KM, Heesen C. Basal serum levels and reactivity of nerve growth factor and brain-derived neurotrophic factor to standardized acute exercise in multiple sclerosis and controls. *J Neuroimmunol*. 2003 May;138(1-2):99-105.
54. Isbell E, Stevens C, Pakulak E, Hampton Wray A, Bell TA, Neville HJ. Neuroplasticity of selective attention: Research foundations and preliminary evidence for a gene by intervention interaction. *Proc Natl Acad Sci U S A*. 2017 Aug 29;114(35):9247-9254.
55. Green CS, Bavelier D. Exercising your brain: a review of human brain plasticity and training-induced learning. *Psychol Aging*. 2008 Dec;23(4):692-701.
56. Rogge AK, Röder B, Zech A, Hötting K. Exercise-induced neuroplasticity: Balance training increases cortical thickness in visual and vestibular cortical regions. *Neuroimage*. 2018 Oct 1;179:471-479.
57. Eriksson PS, Perfilieva E, Björk-Eriksson T, Alborn AM, Nordborg C, Peterson DA, Gage FH. Neurogenesis in the adult human hippocampus. *Nat Med*. 1998 Nov;4(11):1313-7.
58. Kempermann G, Song H, Gage FH. Neurogenesis in the Adult Hippocampus.

- Cold Spring Harb Perspect Biol.* 2015 Sep 1;7(9):a018812.
59. Kumar A, Pareek V, Faiq MA, Ghosh SK, Kumari C. Adult neurogenesis in humans: A Review of Basic Concepts, History, Current Research, and Clinical Implications. *Innov Clin Neurosci.* 2019 May 1;16(5-6):30-37.
  60. Lu L, Bao G, Chen H, Xia P, Fan X, Zhang J, Pei G, Ma L. Modification of hippocampal neurogenesis and neuroplasticity by social environments. *Exp Neurol.* 2003 Oct;183(2):600-9.
  61. Kempermann G, Gast D, Gage FH. Neuroplasticity in old age: sustained fivefold induction of hippocampal neurogenesis by long-term environmental enrichment. *Ann Neurol.* 2002 Aug;52(2):135-43.
  62. McEwen BS. Stress and hippocampal plasticity. *Annu Rev Neurosci.* 1999; 22:105-22.
  63. Mirescu C, Gould E. Stress and adult neurogenesis. *Hippocampus;* 2006;16(3):233-8.
  64. Pittenger C, Duman RS. Stress, depression, and neuroplasticity: a convergence of mechanisms. *Neuropsychopharmacology.* 2008 Jan;33(1):88-109.
  65. Yaribeygi H, Panahi Y, Sahraei H, Johnston TP, Sahebkar A. The impact of stress on body function: A review. *EXCLI J.* 2017 Jul 21;16:1057-1072.
  66. Davidson RJ, McEwen BS. Social influences on neuroplasticity: stress and interventions to promote well-being. *Nat Neurosci.* 2012 Apr 15;15(5):689-95.
  67. Baik SH, Rajeev V, Fann DY, Jo DG, Arumugam TV. Intermittent fasting increases adult hippocampal neurogenesis. *Brain Behav.* 2020 Jan;10(1):e01444.
  68. Lei X, Wu Y, Xu M, Jones OD, Ma J, Xu X. Physical exercise: bulking up neurogenesis in human adults. *Cell Biosci.* 2019 Sep 3;9:74.
  69. Yau SY, Gil-Mohapel J, Christie BR, So KF. Physical exercise-induced adult neurogenesis: a good strategy to prevent cognitive decline in neurodegenerative diseases? *Biomed Res Int.* 2014;2014:403120.
  70. Poulouse SM, Miller MG, Scott T, Shukitt-Hale B. Nutritional Factors Affecting Adult Neurogenesis and Cognitive Function. *Adv Nutr.* 2017 Nov 15;8(6):804-811.
  71. Øverli Ø, Sørensen C. On the Role of Neurogenesis and Neural Plasticity in the Evolution of Animal Personalities and Stress Coping Styles. *Brain Behav Evol.* 2016;87(3):167-74.
  72. Lyons DM, Buckmaster PS, Lee AG, Wu C, Mitra R, Duffey LM, Buckmaster CL, Her S, Patel PD, Schatzberg AF. Stress coping stimulates hippocampal neurogenesis in adult monkeys. *Proc Natl Acad Sci U S A.* 2010 Aug 17;107(33):14823-7.
  73. Papale LA, Seltzer LJ, Madrid A, Pollak SD, Alisch RS. Differentially Methylated Genes in Saliva are linked to Childhood Stress. *Sci Rep.* 2018 Jul 17;8(1):10785.
  74. Jiang S, Postovit L, Cattaneo A, Binder EB, Aitchison KJ. Epigenetic Modifications in Stress Response Genes Associated With Childhood Trauma. *Front Psychiatry.* 2019 Nov 8;10:808.
  75. Gudsnuk K, Champagne FA. Epigenetic influence of stress and the social environment. *ILAR J.* 2012;53(3-4):279-88.
  76. Ein-Dor T, Verbeke WJMI, Mokry M, Vrtička P. Epigenetic modification of the oxytocin and glucocorticoid receptor genes is linked to attachment avoidance in young adults. *Attach Hum Dev.* 2018 Aug;20(4):439-454.
  77. Roberts S, Lester KJ, Hudson JL, Rapee RM, Creswell C, Cooper PJ, Thirlwall KJ, Coleman JR, Breen G, Wong CC, Eley TC. Serotonin transporter corrected. methylation and response to cognitive behaviour therapy in children with anxiety disorders. *Transl Psychiatry.* 2014 Sep 16;4(9):e444. Erratum in: *Transl Psychiatry.* 2014;4:e467.
  78. Miller CWT. Epigenetic and Neural Circuitry Landscape of Psychotherapeutic Interventions. *Psychiatry J.* 2017; 2017:5491812.
  79. Tiffon C. The Impact of Nutrition and Environmental Epigenetics on Human Health and Disease. *Int J Mol Sci.* 2018 Nov 1;19(11):3425
  80. Barrón-Cabrera E, Ramos-Lopez O, González-Becerra K, Riezu-Boj JI, Milagro FI, Martínez-López E, Martínez JA. Epigenetic Modifications as Outcomes of Exercise Interventions Related to Specific Metabolic Alterations: A Systematic Review. *Lifestyle Genom.* 2019;12(1-6):25-44.
  81. Alegría-Torres JA, Baccarelli A, Bollati V. Epigenetics and lifestyle. *Epigenomics.* 2011 Jun;3(3):267-77.
  82. Cole SW. The Conserved Transcriptional Response to Adversity. *Curr Opin Behav Sci.* 2019 Aug;28:31-37.
  83. Boyle CC, Cole SW, Dutcher JM, Eisenberger NI, Bower JE. Changes in eudaimonic well-being and the conserved transcriptional response to adversity in younger breast cancer survivors. *Psychoneuroendocrinology.* 2019 May;103:173-79.
  84. Epel ES. Telomeres in a life-span perspective: a new “psychobiomarker”? *Curr Dir Psychol Sci.* 2009;18:6-10.
  85. Sibille KT, Witek-Janusek L, Mathews HL, Fillingim RB. Telomeres and epigenetics: potential relevance to chronic pain. *Pain.* 2012 Sep;153(9):1789-93.
  86. Asok A, Bernard K, Rosen JB, Dozier M, Roth TL. Infant-caregiver experiences alter telomere length in the brain. *PLoS One.* 2014 Jul 1;9(7):e101437.
  87. Bartley EJ, Palit S, Fillingim RB, Robinson ME. Multisystem Resiliency as a Predictor of Physical and Psychological Functioning in Older Adults With Chronic Low Back Pain. *Front Psychol.* 2019 Aug 22;10:1932.
  88. Puterman E, Epel ES, Lin J, Blackburn EH, Gross JJ, Whooley MA, Cohen BE. Multisystem resiliency moderates the major depression-telomere length association: findings from the Heart and Soul Study. *Brain Behav Immun.* 2013 Oct;33:65-73.
  89. Boccardi V, Paolisso G, Mecocci P. Nutrition and lifestyle in healthy aging: the telomerase challenge. *Aging (Albany NY).* 2016 Jan;8(1):12-5.
  90. Ling C, Rönn T. Epigenetics in Human Obesity and Type 2 Diabetes. *Cell Metab.* 2019 May 7;29(5):1028-1044.
  91. Fitzgerald KN, Hodges, R, Hanes D, Stack E, Cheishvili, D, Szyf M, Henkel J, Twedt MW, Giannopoulou, D, Herdell, J, Logan, S, Bradley, R. Potential reversal of epigenetic age using a diet and lifestyle intervention: a pilot randomized clinical trial. *Aging.* 2021;13(7):9419-32.
  92. Alegría-Torres JA, Baccarelli A, Bollati V. Epigenetics and lifestyle. *Epigenomics.* 2011 Jun;3(3):267-77.
  93. Bókkon I, Vas JP, Császár N, Lukács T. Challenges to free will: transgenerational epigenetic information, unconscious processes, and vanishing twin syndrome. *Rev Neurosci.* 2014;25(1):163-75.
  94. Waggoner MR, Uller T. Epigenetic Determinism in Science and Society. *New Genet Soc.* 2015 Apr 3;34(2):177-195.
  95. McBride CM, Koehly LM. Imagining roles for epigenetics in health promotion research. *J Behav Med.* 2017 Apr;40(2):229-38.
  96. Meissner K, Kohls N, Colloca L. Introduction to placebo effects in medicine: mechanisms and clinical implications. *Philos Trans R Soc Lond B Biol Sci.* 2011 Jun 7;366(1572):1783-9.
  97. Finniss DG. Placebo Effects: Historical and Modern Evaluation. *Int Rev Neurobiol.* 2018;139:1-27.
  98. Turner JA, Deyo RA, Loesser JD, Von Korff M, Fordyce WE. The importance of placebo effects in pain treatment and research. *JAMA.* 1994 May 25;271(20):1609-14.
  99. Diener HC. Placebo effects in treating migraine and other headaches. *Curr Opin Investig Drugs.* 2010 Jul;11(7):735-9.
  100. Darragh M, Yow B, Kieser A, Booth RJ, Kydd RR, Consedine NS. A take-home

- placebo treatment can reduce stress, anxiety and symptoms of depression in a non-patient population. *Aust N Z J Psychiatry*. 2016 Sep;50(9):858-65.
101. Abdullah N. THU0189 Placebo Effect in the Treatment of Rheumatoid Arthritis: A Systematic Review and Meta-Analysis of Randomized Controlled Trials. *Ann Rheum Dis*. 2015;74:263.
  102. Faig-Martí J, Martínez-Catassús A. Measuring the placebo effect in carpal tunnel syndrome. *J Orthop Traumatol*. 2020 Jan 28;21(1):1.
  103. Chen X, Zou K, Abdullah N, Whiteside N, Sarmanova A, Doherty M, Zhang W. The placebo effect and its determinants in fibromyalgia: meta-analysis of randomised controlled trials. *Clin Rheumatol*. 2017 Jul;36(7):1623-30.
  104. Narkus A, Lehnigk U, Haefner D, Klingler R, Pfaar O, Worm M. The placebo effect in allergen-specific immunotherapy trials. *Clin Transl Allergy*. 2013 Dec 21;3(1):42.
  105. Kaptchuk TJ, Hemond CC, Miller FG. Placebos in chronic pain: evidence, theory, ethics, and use in clinical practice. *BMJ*. 2020 Jul 20;370:m1668.
  106. de Craen AJ, Moerman DE, Heisterkamp SH, Tytgat GN, Tijssen JG, Kleijnen J. Placebo effect in the treatment of duodenal ulcer. *Br J Clin Pharmacol*. 1999 Dec;48(6):853-60.
  107. Peciña M, Bohnert AS, Sikora M, Avery ET, Langenecker SA, Mickey BJ, Zubieta JK. Association Between Placebo-Activated Neural Systems and Antidepressant Responses: Neurochemistry of Placebo Effects in Major Depression. *JAMA Psychiatry*. 2015 Nov;72(11):1087-94.
  108. Chvetzoff G, Tannock IF. Placebo effects in oncology. *J Natl Cancer Inst*. 2003 Jan 1;95(1):19-29.
  109. Junior PNA, Barreto CMN, de Iracema Gomes Cubero D, Del Giglio A. The efficacy of placebo for the treatment of cancer-related fatigue: a systematic review and meta-analysis. *Support Care Cancer*. 2020 Apr;28(4):1755-64.
  110. Hall KT, Loscalzo J, Kaptchuk TJ. Genetics and the placebo effect: the placebo. *Trends Mol Med*. 2015 May;21(5):285-94.
  111. Benedetti F, Mayberg HS, Wager TD, Stohler CS, Zubieta JK. Neurobiological mechanisms of the placebo effect. *J Neurosci*. 2005 Nov 9;25(45):10390-402.
  112. Chang WY. Complete spontaneous regression of cancer: four case reports, review of literature, and discussion of possible mechanisms involved. *Hawaii Med J*. 2000 Oct;59(10):379-87.
  113. Kalialis LV, Drzewiecki KT, Mohammadi M, Mehlsen AB, Klyver H. Spontaneous regression of metastases from malignant melanoma: a case report. *Melanoma Res*. 2008;18(4):279-83.
  114. Horino T, Takao T, Yamamoto M, Geshi T, Hashimoto K. Spontaneous remission of small cell lung cancer: a case report and review in the literature. *Lung Cancer*. 2006 Aug;53(2):249-52.
  115. Heianna J, Miyauchi T, Suzuki T, Ishida H, Hashimoto M, Watarai J. Spontaneous regression of multiple lung metastases following regression of hepatocellular carcinoma after transcatheter arterial embolization. A case report. *Hepatogastroenterology*. 2007 Jul-Aug;54(77):1560-2.
  116. Abdelrazeq AS. Spontaneous regression of colorectal cancer: a review of cases from 1900 to 2005. *Int J Colorectal Dis*. 2007 Jul;22(7):727-36.
  117. Papac RJ. Spontaneous regression of cancer. *Cancer Treat Rev*. 1996 Nov;22(6):395-423.
  118. Chohan MBY, Taylor N, Coffin C, Burak KW, Bathe OF. Spontaneous Regression of Hepatocellular Carcinoma and Review of Reports in the Published English Literature. *Case Rep Med*. 2019 Mar 31;2019:9756758.
  119. Kuwal A, Chauhan N, Dutt N, Elhence P, Advani M, Kumar S. Spontaneous Partial Regression of a Carcinoid Tumor: Radiology May Not Capture the Real Picture. *Turk Thorax J*. 2019 Jan 31;20(2):153-6.
  120. Harris WS, Gowda M, Kolb JW, Strychacz CP, Vacek JL, Jones PG, Forker A, O'Keefe JH, McCallister BD. A randomized, controlled trial of the effects of remote, intercessory prayer on outcomes in patients admitted to the coronary care unit. *Arch Intern Med*. 1999 Oct 25;159(19):2273-8.
  121. Schilder JN, de Vries MJ, Goodkin K, Antoni M. Psychological Changes Preceding Spontaneous Remission of Cancer. *Clin Case Stud* 2004;3(4):288-312.
  122. Eccles R. The power of the placebo. *Curr Allergy Asthma Rep*. 2007 May;7(2):100-4.
  123. Chiffi D, Zanotti R. Knowledge and Belief in Placebo Effect. *J Med Philos*. 2017 Feb;42(1):70-85.
  124. Schienle A, Gremsl A, Wabnegger A. Placebo Effects in the Context of Religious Beliefs and Practices: A Resting-State Functional Connectivity Study. *Front Behav Neurosci*. 2021 May 6;15:653359.
  125. Schwartz JM, Stoessel PW, Baxter LR Jr, Martin KM, Phelps ME. Systematic changes in cerebral glucose metabolic rate after successful behavior modification treatment of obsessive-compulsive disorder. *Arch Gen Psychiatry*. 1996 Feb;53(2):109-13.
  126. Nechvatal JM, Lyons DM. Coping changes the brain. *Front Behav Neurosci*. 2013 Feb 22;7:13.
  127. Schwartz JM. A role for volition and attention in the generation of new brain circuitry. Toward a neurobiology of mental force. *J Conscious Stud*. 1999;6(8-9):115-42.
  128. Nechvatal JM, Lyons DM. Coping changes the brain. *Front Behav Neurosci*. 2013 Feb 22;7:13.
  129. Crum AJ, Langer EJ. Mind-set matters: exercise and the placebo effect. *Psychol Sci*. 2007 Feb;18(2):165-71.
  130. Lamme VA. Separate neural definitions of visual consciousness and visual attention; a case for phenomenal awareness. *Neural Netw*. 2004 Jun-Jul;17(5-6):861-72.
  131. Lamme VA. Why visual attention and awareness are different. *Trends Cogn Sci*. 2003 Jan;7(1):12-18.
  132. Jueptner M, Stephan KM, Frith CD, Brooks DJ, Frackowiak RS, Passingham RE. Anatomy of motor learning. I. Frontal cortex and attention to action. *J Neurophysiol*. 1997 Mar;77(3):1313-24.
  133. Draganski B, Gaser C, Kempermann G, Kuhn HG, Winkler J, Büchel C, May A. Temporal and spatial dynamics of brain structure changes during extensive learning. *J Neurosci*. 2006 Jun 7;26(23):6314-7.
  134. Aydin K, Ucar A, Oguz KK, Okur OO, Agayev A, Unal Z, Yilmaz S, Ozturk C. Increased gray matter density in the parietal cortex of mathematicians: a voxel-based morphometry study. *AJNR Am J Neuroradiol*. 2007 Nov-Dec;28(10):1859-64.
  135. Lazar SW, Kerr CE, Wasserman RH, Gray JR, et al. Meditation experience is associated with increased cortical thickness. *Neuroreport*. 2005 Nov 28;16(17):1893-7.
  136. Cf. Briggs F, Mangun GR, Usrey WM. Attention enhances synaptic efficacy and the signal-to-noise ratio in neural circuits. *Nature*. 2013 Jul 25;499(7459):476-80.
  137. Buccino G, Binkofski F, Fink GR, Fadiga L, et al. Action observation activates premotor and parietal areas in a somatotopic manner: an fMRI study. *Eur J Neurosci*. 2001 Jan;13(2):400-4.
  138. Porro CA, Facchin P, Fusi S, Dri G, Fadiga L. Enhancement of force after action observation: behavioural and neurophysiological studies. *Neuropsychologia*. 2007 Oct 1;45(13):3114-21.
  139. Ertelt D, Small S, Solodkin A, Dettmers C, McNamara A, Binkofski F, Buccino G. Action observation has a positive impact on rehabilitation of motor deficits after stroke. *Neuroimage*. 2007;36 Suppl 2:T164-73.
  140. Pascual-Leone A, Nguyet D, Cohen LG, Brasil-Neto JP, Cammarota A, Hallett M. Modulation of muscle responses evoked

- by transcranial magnetic stimulation during the acquisition of new fine motor skills. *J Neurophysiol.* 1995 Sep;74(3):1037-45.
141. Pascual-Leone A, Amedi A, Fregni F, Merabet LB. The plastic human brain cortex. *Annu Rev Neurosci.* 2005; 28:377-401.
142. Ehrsson HH, Geyer S, Naito E. Imagery of voluntary movement of fingers, toes, and tongue activates corresponding body-part-specific motor representations. *J Neurophysiol.* 2003 Nov;90(5):3304-16.
143. Ranganathan VK, Siemionow V, Liu JZ, Sahgal V, Yue GH. From mental power to muscle power-gaining strength by using the mind. *Neuropsychologia.* 2004; 42(7):944-56.
144. Yue G, Cole KJ. Strength increases from the motor program: comparison of training with maximal voluntary and imagined muscle contractions. *J Neurophysiol.* 1992 May;67(5):1114-23.
145. Holden-Lund C. Effects of relaxation with guided imagery on surgical stress and wound healing. *Res Nurs Health.* 1988 Aug;11(4):235-44.
146. Fors EA, Sexton H, Göttestam KG. The effect of guided imagery and amitriptyline on daily fibromyalgia pain: a prospective, randomized, controlled trial. *J Psychiatr Res.* 2002 May-Jun;36(3):179-87.
147. Page SJ, Levine P, Leonard AC. Effects of mental practice on affected limb use and function in chronic stroke. *Arch Phys Med Rehabil.* 2005 Mar;86(3):399-402.
148. Carrico DJ, Peters KM, Diokno AC. Guided imagery for women with interstitial cystitis: results of a prospective, randomized controlled pilot study. *J Altern Complement Med.* 2008 Jan-Feb;14(1):53-60.
149. Cramer SC, Orr EL, Cohen MJ, Lacourse MG. Effects of motor imagery training after chronic, complete spinal cord injury. *Exp Brain Res.* 2007 Feb;177(2):233-42.
150. Freeman LW, Welton D. Effects of imagery, critical thinking, and asthma education on symptoms and mood state in adult asthma patients: a pilot study. *J Altern Complement Med.* 2005 Feb;11(1):57-68.
151. Tamir R, Dickstein R, Huberman M. Integration of motor imagery and physical practice in group treatment applied to subjects with Parkinson's disease. *Neurorehabil Neural Repair.* 2007 Jan-Feb;21(1):68-75.
152. Freeman L, Cohen L, Stewart M, White R, Link J, Palmer JL, Welton D. Imagery intervention for recovering breast cancer patients: clinical trial of safety and efficacy. *J Soc Integr Oncol.* 2008 Spring;6(2):67-75.
153. Baird CL, Sands LP. Effect of guided imagery with relaxation on health-related quality of life in older women with osteoarthritis. *Res Nurs Health.* 2006 Oct;29(5):442-51.
154. Louie SW. The effects of guided imagery relaxation in people with COPD. *Occup Ther Int.* 2004;11(3):145-59.
155. Luberto CM, Hall DL, Park ER, Haramati A, Cotton S. A Perspective on the Similarities and Differences Between Mindfulness and Relaxation. *Glob Adv Health Med.* 2020 Feb 5;9:2164956120905597.
156. Goldin PR, Gross JJ. Effects of mindfulness-based stress reduction (MBSR) on emotion regulation in social anxiety disorder. *Emotion.* 2010 Feb;10(1):83-91.
157. Brower V. Mind-body research moves towards the mainstream. *EMBO Rep.* 2006 Apr;7(4):358-61.
158. Cushing RE, Braun KL. Mind-Body Therapy for Military Veterans with Post-Traumatic Stress Disorder: A Systematic Review. *J Altern Complement Med.* 2018 Feb;24(2):106-114.
159. Khusid MA, Vythilingam M. The Emerging Role of Mindfulness Meditation as Effective Self-Management Strategy, Part 1: Clinical Implications for Depression, Post-Traumatic Stress Disorder, and Anxiety. *Mil Med.* 2016 Sep;181(9):961-8.
160. Scott J, Teasdale JD, Paykel ES, Johnson AL, Abbott R, Hayhurst H, Moore R, Garland A. Effects of cognitive therapy on psychological symptoms and social functioning in residual depression. *Br J Psychiatry.* 2000 Nov;177:440-6.
161. Jacobs TL, Shaver PR, Epel ES, Zanesco AP, et al. Self-reported mindfulness and cortisol during a Shamatha meditation retreat. *Health Psychol.* 2013 Oct;32(10):1104-9.
162. Rosenzweig S, Reibel DK, Greeson JM, Edman JS, Jasser SA, McMearty KD, Goldstein BJ. Mindfulness-based stress reduction is associated with improved glycemic control in type 2 diabetes mellitus: a pilot study. *Altern Ther Health Med.* 2007 Sep-Oct;13(5):36-8.
163. Hölzel BK, Carmody J, Vangel M, Congleton C, Yerramsetti SM, Gard T, Lazar SW. Mindfulness practice leads to increases in regional brain gray matter density. *Psychiatry Res.* 2011 Jan 30;191(1):36-43.
164. Gross CR, Kreitzer MJ, Russas V, Treesak C, Frazier PA, Hertz MI. Mindfulness meditation to reduce symptoms after organ transplant: a pilot study. *Adv Mind Body Med.* 2004 Summer;20(2):20-9.
165. Carlson LE, Speca M, Patel KD, Goodey E. Mindfulness-based stress reduction in relation to quality of life, mood, symptoms of stress, and immune parameters in breast and prostate cancer outpatients. *Psychosom Med.* 2003 Jul-Aug; 65(4):571-81.
166. Carlson LE, Speca M, Faris P, Patel KD. One year pre-post intervention follow-up of psychological, immune, endocrine and blood pressure outcomes of mindfulness-based stress reduction (MBSR) in breast and prostate cancer outpatients. *Brain Behav Immun.* 2007 Nov;21(8):1038-49.
167. Speca M, Carlson LE, Goodey E, Angen M. A randomized, wait-list controlled clinical trial: the effect of a mindfulness meditation-based stress reduction program on mood and symptoms of stress in cancer outpatients. *Psychosom Med.* 2000 Sep-Oct;62(5):613-22.
168. Rosenkranz MA, Davidson RJ, Maccoon DG, Sheridan JF, Kalin NH, Lutz A. A comparison of mindfulness-based stress reduction and an active control in modulation of neurogenic inflammation. *Brain Behav Immun.* 2013 Jan; 27(1):174-84.
169. Cf. McClelland DC, Kirshnit. C. The effect of motivational arousal through films on salivary immunoglobulin A. *Psychology and Health.* 1998;2:31-52.
170. Davidson RJ. The Emotional Life of Your Brain: How Its Unique Patterns Affect the Way You Think, Feel, and Live – and How You Can Change Them, Avery; 2012.
171. Desbordes G, Negi LT, Pace TW, Wallace BA, Raison CL, Schwartz EL. Effects of mindful-attention and compassion meditation training on amygdala response to emotional stimuli in an ordinary, non-meditative state. *Front Hum Neurosci.* 2012 Nov 1;6:292.
172. Klimecki OM, Leiberg S, Lamm C, Singer T. Functional neural plasticity and associated changes in positive affect after compassion training. *Cereb Cortex.* 2013 Jul;23(7):1552-61.
173. Newberg A. The brain and the biology of belief: An interview with Andrew Newberg, MD. Interview by Nancy Nachman-Hunt. *Adv Mind Body Med.* 2009 Spring;24(1):32-6.
174. Werk RS, Steinhorn DM, Newberg A. The Relationship Between Spirituality and the Developing Brain: A Framework for Pediatric Oncology. *J Relig Health.* 2021 Feb;60(1):389-405.
175. Koenig HG. Religion, Spirituality, and Health: The Research and Clinical Implications. *International Scholarly Research Network ISRN Psychiatry.* 2012.
176. Bonelli RM, Koenig HG. Mental disorders, religion and spirituality 1990 to 2010: a systematic evidence-based review. *J Relig Health.* 2013 Jun;52(2):657-73.
177. Koenig HG, Al Zaben FN, Al Shohaib S. Religion and Health: Clinical Considerations and Applications, International Encyclopedia of the Social &

- Behavioral Sciences, 2nd edition, Volume 20:262-8.
178. Rossi EL. Psychobiology of Gene Expression, Neuroscience and Neurogenesis in Hypnosis and the Healing Arts, WW Norton, 2002.
179. Dusek JA, Otu HH, Wohlhueter AL, Bhasin M, Zerbini LF, Joseph MG, Benson H, Libermann TA. Correction: Genomic Counter-Stress Changes Induced by the Relaxation Response. *PLoS One*. 2017 Feb 21;12(2):e0172845.
180. Jacobs TL, Epel ES, Lin J, Blackburn EH, et al. Intensive meditation training, immune cell telomerase activity, and psychological mediators. *Psychoneuroendocrinology*. 2011 Jun;36(5):664-81.
181. Schutte NS, Malouff JM. A meta-analytic review of the effects of mindfulness meditation on telomerase activity. *Psychoneuroendocrinology*. 2014 Apr;42:45-8.
182. Conroy D, Hagger MS. Imagery interventions in health behavior: A meta-analysis. *Health Psychol*. 2018 Jul;37(7):668-679.
183. Jiménez JP, Botto A, Herrera L, Leighton C, Rossi JL, et al. Psychotherapy and Genetic Neuroscience: An Emerging Dialog. *Front Genet*. 2018 Jul 17;9:257.