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Psychiatry/Mental Health Section

Potential of Yoga Therapy in Modulating Neurotransmitters for Treatment and Prevention of Autism Spectrum Disorder: A Narrative Review

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ABSTRACT

Autism Spectrum Disorder (ASD) is a complex neurodevelopmental condition characterised by impairments in communication, social interaction, and behaviour. Increasing evidence suggests that maternal psychological stress during pregnancy can lead to neuroinflammation and disruption of neurotransmitter systems. This review aims to critically evaluate the potential role of yoga therapy in regulating neurotransmitter systems to mitigate ASD symptoms in children and reduce maternal psychological stress during pregnancy. A comprehensive literature review was conducted using PubMed and Google Scholar. The keywords used for the search included "Yoga," "pregnancy," "maternal psychological stress," "ASD," and "foetal neurodevelopment" in various permutations and combinations with "neurotransmitters." The focus was on neurotransmitters, including serotonin, dopamine, and Gamma-Aminobutyric Acid (GABA), emphasising their roles in mood regulation, behaviour, and cognitive functions. The review found that yoga therapy modulates neurotransmitter systems by reducing stress and enhancing emotional regulation and cognitive function in the treatment of ASD. It also helps prevent the adverse effects of maternal psychological stress on foetal neurodevelopment during pregnancy.

Keywords: Acetylcholine, Dopamine, Gamma-aminobutyric acid, Neuroplasticity, Serotonin

INTRODUCTION

ASD is a complex neurodevelopmental condition that affects social interaction, communication, and behaviour. Several studies have found that psychological imbalance, as well as sensory and cognitive dysfunctions, play major roles in abnormal brain development, contributing to memory, motor activity, and behavioural dysfunctions [1,2]. These factors indicate defects in nerve cell migration, differentiation, synaptogenesis, apoptosis, and synaptic pruning, which are caused by dysfunctions in neurotransmitter systems in children with ASD [2].

Given the broad distribution of neurotransmitters, the complex and diverse functions of monoamine neurotransmitters are crucial for social communication, as they coordinate sensory, motor, and limbic systems across different brain regions. Several studies have reported alterations in neurotransmitters, including GABA, glutamate, dopamine, norepinephrine, acetylcholine, serotonin, and opioid peptides in children with ASD [3,4]. Despite extensive research over the past three decades on the aetiology and pathophysiology of ASD, the mechanisms behind its pathogenesis remain elusive [5]. Since the root causes of this disorder are not yet fully understood, preventive strategies have yet to be developed.

With the growing prevalence of this condition, current estimates suggest that approximately 1 in 54 children are diagnosed with ASD [6]. Given this increasing incidence and the complex nature of ASD, there is a compelling need for effective interventions that address the multifaceted challenges associated with the disorder and its prevention strategies.

Several studies have found that regular yoga practice:

- 1. Improves motor coordination, academic-specific building actions, and imitation skills [7];
- Significantly reduces irritability, hyperactivity, aggression, and social withdrawal [8], as well as behavioural dysfunction [9], promoting a shift towards parasympathetic activity that fosters calmness, self-regulation, and resilience;
- 3. Helps ameliorate gastrointestinal distress and sleep disorders [10] in children with ASD [11].

Research indicates that yoga may help correct neuroinflammation caused by imbalanced neurotransmitters [12], such as GABA, which is linked to relaxation and anxiety [13], and may help increase serotonin levels, contributing to mood stabilisation [14]. This could potentially improve emotional regulation in children with ASD [15]. Preliminary findings suggest that yoga can positively influence neurotransmitter systems, promoting better mental health and social skills in individuals with ASD [2,16]. Most interventions target children already diagnosed with ASD, leaving a significant research gap regarding preventive measures [17,18].

The purpose of this review is to critically evaluate the potential of yoga therapy as an effective intervention for ASD, both as a therapeutic and preventive measure, with a particular focus on its influence on neurotransmitter alterations and associated neurodevelopmental processes.

REVIEW OF LITERATURE

A comprehensive literature review was performed using electronic databases, including PubMed and Google Scholar. This review examined relevant studies on the aetiopathogenesis of ASD and the effect of yoga therapy on the modulation of neurotransmitters in the treatment of ASD, as well as preventive measures for maternal psychological stress during pregnancy. Keywords used in the search included "Yoga Therapy," "Neurotransmitters," "ASD," "Maternal Psychological Stress," "Serotonin," "GABA," and "Dopamine." Data were extracted from each study, focusing on outcomes related to neurotransmitters or behavioural/emotional changes.

RESULTS

A summary of the effect of yoga interventions on neurotransmitters based on available literature for the treatment of ASD and the prevention of adverse effects of maternal psychological stress on foetal neurodevelopment during pregnancy is provided in [Table/Fig-1] [19-37].

Neurotransmitters and receptors	Imbalanced neurotransmitters	Area of dysfunction	Behavioural dysfunctions	Treatment: YT on behavioural dysfunctions in ASD	Prevention: YT on Maternal psychological stress during pregnancy
Acetylcholine [19-22] Nicotinic, Muscarinic	↓ α4β2 nAChRs in parietal and frontal cortex, ↓ 04, ↑ α7 nAChRs in cerebellum	CNS: Memory, cognition. PNS: Skeletal muscle and parasympathetic nervous system	Irritability, hyperactivity, poor verbal learning;poor thinking ability and memory, frequent mood swings; Inattention in patients with psychiatric disorders such as ASD and ADHD often associated with low levels of acetylcholine.	Improved motor coordination, static and dynamic balance; reduced irritability, aggression, social withdrawl, non compliance, and behavioural dysfunctions [23,24].	Reduced gastrointestinal, respiratory, and mental health symptoms in maternal psychological stress during pregnancy [25,26].
Catecholamine [27] Excites and Inhibits (α1β1; α2β2 β3) Noradrenaline Adrenaline	-	CNS: Opioid regulation. PNS: Sympathetic nervous system	Dysfunction of catecholamine leads to core symptoms of ASD.	Reduced core symptoms of ASD, particularly in latency-age children [8].	
Dopamine [27] DA _B ; DA _A	Imbalanced DA in prefrontal cortex	CNS: Motivation, motor control. PNS: BV, GIT, Renal system; Dysregulation of mesocortico-limbic and nigrostriatal circuit	Reduced motivation to pursue social interactions.	Increased ability to stay still without typical fidgeting, improved self-control, enhanced quality of life, and better social responsiveness, communication, cognition, and motivation [24].	
Serotonin [28] 5-HT2,3,4,5,6; 5-HT1-s1	↓ 5-HT2A, 5-HT1A in brain and blood	CNS: sleep, mood. PNS: GIT and skeletal system; Hippocampus	Poor sleep quality and social initiation.	Improved sleep [10].	
GABA [29,30] Inhibits: GABA _A , GABA _B	↓ GABA in motor, visual, auditory, somatosensory cortex	Inhibits action potential of neurons in superior frontal cortex, parietal cortex and cerebellum; and various brain regions.	Poor socio-emotional and cognitive processing, such as difficulty identifying faces and facial expressions, leads to functional deficits in individuals with ASD.	Developed positive social and emotional skills in youth with ASD [31].	
Glutamate [32,33] Excites: GluR _A , GluR _B	† glutamatergic activity	Excites neurons of anterior cingulated cortex and central nervous system	Poor social and communication skills.	Improved social communication and generalised joint attention skills [7].	
Neurotensin (NT) [34] Excites: NK1R	-	CNS and PNS: pain and inflammation	Stimulation of immune cells, especially mast cells, and/or have direct effects on brain inflammation	Movement-based experiences are multimodal in nature and activate similar "mirror" networks in the brain of participants, thereby forming the basis for social, emotional, and motor coordination in individuals with ASD [35].	
Substance P or Endorphins [36]	↑β endorphin in plasma		Hyperactivity and restlessness	Improvement in ADHD symptoms such as inattention and hyperactivity, as shown in parent reports [37].	

[Table/Fig-1]: Summary of the effect of YT on neurotransmitters in treatment and prevention of ASD [19-37]. α(4)β(2)nAChRs: alpha-4 beta-2 nicotinic acetylcholine receptors; CNS: Central nervous system; PNS: Peripheral nervous system; ANS: Autonomic nervous system; ADHD: Attention deficit hyperactivity disorder; DA(a or b): Dopamine (alpha or beta) hydroxylase; BV: Blood vessels; GIT: Gastro-intestinal tract; 5-HT: 5-hydroxytryptamine; GABA(a or b): G-protein coupled (alpha or beta) receptors for gamma aminobutyric acid; (i or m) GluRs: (ionotropic or metabotropic) glutamate receptors; GluR(a or b): subunits of glutamate receptorare functionally expressed on α and β cells; and NK1R: neurokinin 1 receptors.

DISCUSSION

Aetiopathogenesis of ASD: According to *Thirumanthiram*, a classical yoga text in Tamil dating back 3,000 years [38,39], Neurodevelopmental Disorders (NDDs) are described as congenital disorders in verse 481, stating:

"Maadha udhara mala migil mandanaam,

Maadha udhara jala migil moongaiyaam,

Maadha udhara irandokkil kannillai,

Maadha udharatthil vanda kuzhavikke."

It is explained that chronic psychological stress during pregnancy can lead to partial evacuation of the mother's bowel, bladder, or both. This may result in transient neuroinflammation in the foetus, potentially manifesting as NDDs such as Intellectual Disability (ID), Attention Deficit Hyperactivity Disorder (ADHD), or ASD [40].

Chronic stress can cause autoimmune diseases. When women with autoimmune diseases consume a high-fat diet during pregnancy and are exposed to pollution and infections, this leads to the development of inflammatory cytokines in plasma, the placenta, and amniotic fluid. Such a state is known as maternal psychological stress—maternal immune activation—which further leads to microglial activation,

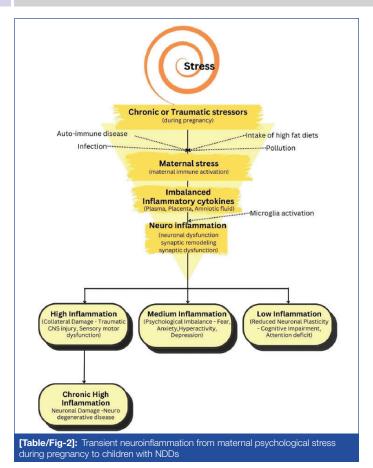
immune alteration, and neuronal and synaptic dysfunction in the brain of the offspring [17], as shown in [Table/Fig-2].

This neuroinflammation leads to:

Collateral Neuronal Damage: Sensorimotor dysfunction, especially when sensory processes from sound, touch, sight, smell, taste, and the vestibular system remain imbalanced. This can develop into challenging behaviours such as difficulties in catching and throwing, clumsiness, poor attention and perception, failure to perceive extreme odours, increased pica or overeating (spicy food), a high pain threshold, self-harm, and an inability to feel or chew food in the mouth, particularly an inability to acknowledge certain sounds. Overloaded sensory stimuli may increase sensitivity, cause hearing impairments, distortions, and indifference toward people (associated with distinctive perfumes, shampoos, etc.), leading to a restricted diet.

Psychological Imbalance: Anxiety, fear, hyperactivity, impulsivity, etc. **Cognitive Dysfunction:** Resulting from altered neurotransmission processes, which further lead to behavioural impairments in children with ASD, as illustrated in [Table/Fig-2] [18,41,42].

Therapeutic effect of yoga on treatment and prevention of ASD: According to Hatha Yoga Pradipika, a classical yoga text, it is stated



that regardless of age or condition, whether young, old, or afflicted, through regular yoga practice, one can attain health and wellbeing, as mentioned in verse 66 [43].

Role of Yoga Therapy in children with ASD

Yoga therapy includes:

Yogic counseling: Utilises psycho-socio-spiritual principles along with dietary and lifestyle modifications to integrate and restore fundamental bodily systems and psychological processes, thereby fostering improved self-regulation.

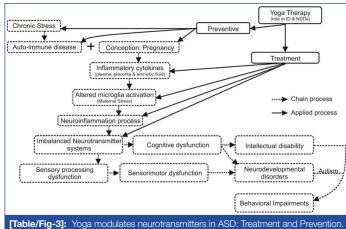
Yogasana: Body-breath coordinated movement enhances awareness, strength, and body stabilisation by stimulating the release of endorphins and mood enhancers.

Pranayama: Regulated breath prepares the body to handle anxiety and stress by modulating the autonomic nervous system and potentially influencing neurotransmitters like acetylcholine, leading to enhanced relaxation and reduced stress responses [44,45].

Yogic relaxation techniques promote deep respiration and calmness in the mind by increasing the production of serotonin and dopamine, neurotransmitters associated with feelings of wellbeing and pleasure, as shown in [Table/Fig-3] [16].

Yoga Therapy on Modulation of Neurotransmitters in Treatment of ASD (as presented in [Table/Fig-1])

1. Dopamine (DA) is a key neurotransmitter that plays a crucial role in regulating social cognition, involuntary movement control, and behaviour [46,47]. Several studies have found that regular yoga practice alleviates stress and anxiety, promoting a state of calmness and mindfulness, potentially due to dopamine release [9]. This activation may improve motivation to engage in social interactions, thereby addressing a core deficit in ASD. Moreover, a yogic diet—including avocados, bananas, pumpkin, sesame seeds, and soy products—may help improve tyrosine levels, while pranayama can increase the release of neurotransmitters associated with positive mood states. Yoga interventions have led to improvements in behavioural



outcomes for children with ASD, including increased social engagement and communication skills, potentially mediated by enhanced responsiveness of the dopaminergic system to social stimuli [8,48].

- 2. Serotonin is an important neurotransmitter that influences mood, sleep, and various brain functions. Multiple studies have shown that yoga interventions help achieve homeostasis by balancing sympathetic and parasympathetic activities, thereby maintaining equilibrium in plasma levels of serotonin, melatonin, and glutamate neurotransmitters [49]. Regular yoga practice is associated with higher serotonin levels, resulting in improved mood and reduced anxiety in individuals with ASD [16]. Additionally, following a yogic diet that includes vegetables, fruits, legumes, and whole grains helps increase tryptophan [50], which is crucial since poor sleep has been linked to decreased serotonin production [10]. Yoga therapy enhances Brain-Derived Neurotrophic Factor (BDNF) and serotonin levels, promoting better sleep and facilitating improved emotional regulation and social interaction in children with ASD [51].
- GABA is a significant inhibitory neurotransmitter that helps regulate brain activity. Several studies have found that yoga significantly reduces cortisol, increases GABA, and enhances the activity of GABAergic systems and peripheral oxytocin, thereby promoting relaxation and reducing anxiety [52-55]. Furthermore, incorporating a yogic diet, particularly with fermented foods, germinated wheat, barley, fresh vegetables (like broccoli and spinach), fruits (such as bananas and berries), nuts (almonds and walnuts), buttermilk, lentils, brown rice, and whole grains, helps increase the enzyme glutamic acid decarboxylase. Additionally, pranayama in yoga practice has been shown to enhance the regulation of the autonomic nervous system, often linked to lowered GABA levels [45]. Thus, yoga therapy is established as an effective intervention for modulating GABA levels, improving socio-emotional regulation, and mitigating anxiety-related behaviours in children with ASD [53].
- 4. Glutamate is the most common excitatory neurotransmitter that plays a key role in brain function, including memory, cognition, and mood regulation. Numerous studies have found that yoga enhances mood and emotional expression, increases empathy towards others, and improves social skills, which can be correlated with better regulation of glutamate, thereby increasing neuroplasticity in individuals with ASD [31,45,55,56].
- 5. Acetylcholine (ACh) plays a significant role in memory, learning, attention, and involuntary muscle movement. Multiple studies have shown that yoga enhances parasympathetic nervous system activity, potentially leading to increased ACh and its receptor sensitivity, which are associated with improved neuroplastic effects and neuropsychological functions such

as learning, attention, and neuromuscular coordination in individuals with ASD [8,23]. Furthermore, a yogic diet that includes legumes and seeds increases choline levels, facilitating better engagement in social situations and improving cognitive function.

Role of Yoga Therapy in prevention of ASD

Pregnancy can be impacted by various psychosocial and lifestyle factors that contribute to maternal psychological stress [57]. Yoga intervention reduces stress, cardiometabolic risks, and gestational hypertension, improves foetomaternal outcomes, and shows promise for enhancing newborn outcomes [58,59]. Several studies have demonstrated that yoga therapy significantly improves autonomic nervous system function, GABA activity, and allostatic load [55], as well as serum BDNF levels [60], while reducing serum cortisol [61]. These effects enhance neurotransmitter regulation, helping alleviate maternal psychological stress during pregnancy [59].

This review highlights the significant impact of yoga therapy, emerging as a multifaceted intervention capable of fostering positive behavioural outcomes through the systematic modulation of neurotransmitters, specifically focusing on GABA, glutamate, serotonin, and acetylcholine in children with ASD. Yoga therapy also enhances neurotransmitter regulation during pregnancy, offering a promising approach in reducing the risk of ASD.

Limitation(s)

The study identifies the following specific limitations: a lack of rigorous research (existing studies may have methodological weaknesses, such as small sample sizes, lack of control groups, or potential biases) and standardised research protocols (the lack of uniformity in research protocols makes it difficult to compare results across different studies and draw definitive conclusions). These limitations impact the strength and generalisability of the findings regarding the effect of yoga therapy on neurotransmitters in treating and preventing ASD.

CONCLUSION(S)

Yoga therapy modulates key neurotransmitters in the treatment of ASD, thereby enhancing emotional regulation and social skills. Additionally, it shows potential as a preventive measure for ASD by alleviating maternal psychological stress during pregnancy. While preliminary findings are promising, further research is required to fully validate its therapeutic benefits.

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