

Fetuses can Listen, Learn, and Remember: We Need to be Cautious about What and How We Say It!

Adrianna Frydrysiak-Brzozowska^{1,2}, Kedar Jape³, Gayatri Athalye-Jape⁴, Kinga Piórkowska¹, Srijan Singh^{2,5}, Thierry AGM Huisman^{2,6}, Akhil Maheshwari^{2,7,8}

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ABSTRACT

The fetal auditory system becomes functional during mid-gestation or possibly even earlier. Existing data show that fetuses can respond to maternal voice and different types of music, both vocal and instrumental. The ability to receive and transmit sound waves, and then recognize and retain some memory of these auditory stimuli could possibly be one of the most important developmental sensory milestones that we need to learn about. Unfortunately, we still have limited evidence for the precise role and timing of prenatal sound simulation. There is a need for methodologically strong, randomized controlled trials with rigorously designed interventions and standardized reporting measures. We may need to compare different durations and types of musical (sound) intervention. At a minimum, these interventions can improve maternal–fetal bonding and family-centered outcomes. Any evidence of neurodevelopmental gains would be an important scientific/medical advancement. In certain conditions such as neonatal abstinence syndrome, emerging evidence suggests that early, *in utero* intervention with music therapy can be helpful; these findings bring hope for new therapeutic tools to enhance the neurological development of at-risk fetuses. Considering that prenatal music exposure might have positive effects on the fetus and newborn infant, we need carefully conducted studies of intrauterine neurosensory organization with long-term follow-up.

Keywords: Fetus, Mother's cardiocographic parameters, Music therapy, Neonatal behavior, Neonatal neurological system, Newborn, Pregnancy, Rhythm, Sound, Speech.

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KEY POINTS

- In terms of structure, the fetal hearing system is recognizable at early as 3–6 weeks of pregnancy. Functionally, the inner, middle, and outer ear are sufficiently developed at 24–25 weeks' gestation to identify/distinguish between vibroacoustic stimuli.
- Fetuses and newborn infants can respond to maternal voice across her abdominal wall. They also respond to different types of music, both vocal and instrumental.
- *In utero* exposure to music and speech induces stimulus-specific memory traces in the fetus. These stimuli may promote brain growth and possibly even short- and long-term cognitive gains.
- There is some evidence that music therapy can help in disorders such as neonatal abstinence syndrome. Possibly, timely *in utero* intervention may also enhance neurological development in other high-risk conditions.
- We need to carefully evaluate the immediate and long-term effects of music therapy on intrauterine neurosensory organization.

INTRODUCTION

The fetus is exposed to a variety of auditory stimuli from as maternal heart sounds, voice, respiratory and bowel sounds, and environmental stimuli.¹ Since hearing is acquired early during pregnancy, there is a possibility that planned antenatal exposure to music might have a role in fetal learning.^{2,3} Considering that music also has a calming effect on the mother with few known adverse effects,⁴ there is curiosity as to whether music could/should be a routine intervention during pregnancy.⁵ In this article, we have reviewed the information available on the impact of prenatal exposure to music on the growing fetus. We have assimilated

¹Faculty of Health Science, Collegium Medicum, The Masovian University in Płock, Płock, Poland

²Global Newborn Society, Clarksville Maryland, United States of America

³Department of Obstetrics and Gynaecology, Joondalup Health Campus, Perth, Western Australia

⁴Department of Neonatology, Child and Adolescent Health Service, King Edward Memorial Hospital, Perth, Western Australia, Australia

⁵Department of Neonatology, Kailash Hospital, Noida, Uttar Pradesh, India

⁶Edward B. Singleton Department of Radiology, Texas Children's Hospital and Baylor College of Medicine, Houston, United States of America

⁷Department of Neonatology/Pediatrics, Louisiana State University Health Sciences Center – Shreveport, Louisiana, United States of America

⁸Banaras Hindu University Institute of Eminence, Varanasi, Uttar Pradesh, India

Corresponding Author: Adrianna Frydrysiak-Brzozowska, Faculty of Health Sciences, Collegium Medicum, The Masovian University in Płock, Płock, Poland, Phone: +243665414 216, e-mail: a.frydrysiak-brzozowska@mazowiecka.edu.pl

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information from an extensive review of the literature utilizing key terms in multiple databases including PubMed, EMBASE, and Science Direct.^{6,7} To avoid bias in the identification of studies, keywords were short-listed *a priori* from anecdotal experience and PubMed's Medical Subject Heading (MeSH) thesaurus.^{8–12}

The Beginnings of Sound Reception in Fetal Development

At the time of birth, a newborn baby is largely equipped with all the senses available to an adult.¹³ The ability to receive and transmit stimuli in sound waves is important and matures rapidly in the second trimester.¹⁴ Training becomes possible in later pregnancy.¹⁵

The development of the auditory system begins between 3 and 6 weeks of pregnancy.¹⁶ At 24–25 weeks' gestation, the structures of the inner, middle, and outer ear are sufficiently developed to recognize vibroacoustic stimuli.¹⁷ In the developing central nervous system (CNS), auditory stimuli can promote the maturation of the temporal lobe regions that help recognize low-frequency sounds.¹⁸ The fetus hears sounds coming from the mother's body first, such as those emanating from her cardiac contractions, blood flow in larger vessels, intestinal peristalsis, or airflow in the respiratory tract. The hearing threshold between 27 and 28 weeks' gestation is about 40 dB.^{19,20} The fetuses then respond to sounds with a frequency of 250–500 Hz.^{21,22} Ongoing myelination of nerve fibers in the CNS continues to improve impulse conduction.²³ Between 29 and 30 weeks' gestation, the fetus can perceive higher-frequency sounds such as the mother's voice, an infant's cry, or an alarm.²⁴ During the 34–35 weeks period, fetuses no longer seem to get alarmed with sounds that previously caused arousal. They begin to respond to sounds with frequencies of 1000–3000 Hz. At term, the hearing threshold is 13.5 Hz.²⁵

During pregnancy, extraneous sounds are usually first transmitted from the air, through the abdominal wall, the uterus, and finally to the amniotic fluid and the head of the fetus.²⁶ Intra-uterine sounds might be more important. The sound of the mother's heartbeat is 25 dB above basic noise, dominating the fetal environment.²⁴ Her voice is heard almost 4 times louder in the womb than outside.²⁷ Most sounds in the fetal environment are dominated by lower frequencies.²⁸ Energy above 0.5 kHz is attenuated by 40–50 dB. The fetus easily detects vowels, whereas consonants, which are higher in frequency and less intense than vowels, are largely unavailable.²⁸

The Importance of Music in the Prenatal Period

Music stimulation of a growing fetus is plausibly a positive stimulus to promote a range of positive outcomes for both the mother and the fetus, the child.

Emotional Bonding and Stress Reduction

Long-standing cultural practices view music as a likely catalyst for emotional connection, allowing a mother to establish a deeper bond with her unborn child.²⁹ Soothing songs and singing lullabies have been encouraged as a means to decrease stress and promote relaxation and emotional well-being. The harmonious vibrations of music can help to create a serene environment, fostering a sense of peace and tranquility for both the mother and baby.

Language Development and Cognitive Stimulation

Prenatal music and speech form stimulus-specific memory traces during the fetal period,^{30,31} thereby stimulating brain growth, cognitive development, and long-term neural effects.³² Fetuses start

responding to music in the second trimester.³³ Singing by pregnant women provides auditory stimulation to their unborn child.^{33,34}

Language acquisition and development of hearing could be enhanced by rhythmic melodies.^{33,34} Cognitive abilities and habituation for music can be demonstrated in fetuses and young infants.³⁵ Arguably, the ability to discriminate speech stimuli *in utero* can promote acquisition of native language fundamentals during fetal life.³⁶ Neonates prefer their mother's voice and also the language their mothers used while pregnant with them.^{37–39}

Sleep Regulation and Relaxation

In newborn infants, soothing melodies and instrumental compositions can help establish healthy sleep patterns in both the mother and a newborn infant.⁴⁰ Similarly, music can promote natural sleep in both the pregnant mother and her fetus by reducing stress.⁴¹ As shown in late preterm infants, these effects could possibly be mediated by the release of endorphins in the CNS.^{40,42,43}

Bonding and Maternal Heartbeat

In the antenatal period, mother's heartbeat is a familiar and comforting rhythm for the baby.⁴⁴ Rhythmic melodies and instrumental compositions can simulate maternal heartbeat and promote mother–fetal bonding, which might extend into the postnatal period.^{29,45–47}

Enhanced Sensory Development

Music can promote the development of the fetal sensory pathways and sensory integration.³¹ Vibrations caused by music can be felt by the fetus, creating a multisensory experience that involves hearing, movement, and consequently, touch with the uterine walls, the umbilical cord, and even its own face.^{42,48,49} The movements indicate their engagement with the auditory stimuli, thereby preparing them for the postnatal sensory experiences.³⁰

Maternal–Fetal Interactions

Fetal activities can be observed and followed since early pregnancy.⁵⁰ The maturation of these activities is closely related to the development of the sense organs even *in utero*. Fetal motor activity can be identified as early as at 6–8 weeks' gestation.⁵¹ At 12 weeks, the movements can now be characterized as “kicking,” “fetal rotation,” “opening/closing of the fist,” “nodding,” “sucking-like oral activity,” and “reactions to touch.”⁵¹ As pregnancy progresses, these activities become more frequent, mature with larger amplitude and improved coordination of muscle groups.⁵² These movements are brief but the frequency progressively increases toward term gestation.⁵³

In 2015, Marx and Nagy showed that fetuses reacted with arm, head, and oral movements when the mother touched the belly.^{54,55} Her voice evoked head and arm movements. Third-trimester fetuses showed more yawning (regulatory), resting (crossed arms), and independent touch (hands touching body) responses to stimuli compared with those seen during the second trimester. Two years later, the same investigators used 3- and 4-dimensional ultrasound to examine fetal responses to tactile stimuli on maternal abdomen. Touching the belly by the mother, father, and a stranger was compared. In the control group, the fetal response was examined while at rest, without touching the abdomen. When the mother touched her abdominal wall, the third-trimester fetus touched the uterine wall for significantly longer periods than those in the second trimester. The third-trimester fetuses showed stronger/longer responses when touched by their mother. These differential

responses in older fetuses may reflect the maturation of the CNS and the emergence of proprioceptive self-awareness.⁵⁴

In 2021, Nagy et al.⁵⁶ examined fetal responses to maternal touch and voice. The sensorimotor competencies of the fetus clearly suggest that communication readiness begins to develop prior to birth. In response to maternal abdominal touch, fetuses show responses that have been variably perceived by different observers as cross-communication or self-stimulation. When the mother touched her abdominal wall and spoke in a gentle voice, fetal movements paused for some time. Similar responses were seen during conversations with a third person (interpersonal conversation). The fetuses showed a longer mouth-opening time during these periods. The results suggest that third trimester fetuses can distinguish between interactive and non-interactive external stimuli and also distinguish these from incidental, other interactions.

In 2012, Kisilevsky et al.⁵⁷ examined the impact of auditory stimuli on fetal heart rate (HR) and heart movements in diabetic and overweight pregnancies. Fetuses in overweight pregnancies responded to their mother's voice with increased HR. These altered reactions to the maternal voice in diabetic pregnancies may indicate relatively immature nervous or auditory system, increased sensorineural threshold, lower levels of thyroid hormones, or iron deficiency in the mother.

Lee et al.⁵⁸ compared fetuses in women with hypertension vs controls upon hearing maternal voice. In the maternal hypertension group, the fetuses showed increased HR upon listening to the mother's voice. However, if the voice was recreated, there was no increase in fetal HR. The differential responses of the fetus to the maternal voice in pregnancies complicated by maternal hypertension may reflect a functional increase in the sensorineural threshold or a delay in the maturation of the auditory system, which may indicate functional differences during fetal life or subtle differences in the development of the CNS. Recent observations of maternal voice recognition provide evidence for residual memory and learning in healthy fetuses. We still need to investigate higher-order auditory processing in maternal hypertension, which is associated with reduced and/or compromised uteroplacental blood flow.

Knowledge of memory is based largely on observing newborns who seem to recognize their mother's voice. A few studies have assessed the biological processes that facilitate fetal responses to the maternal voice *in utero*.⁵⁹ Fetal HR and motor activity have been recorded at 36 weeks' gestation while pregnant women read aloud from a neutral passage.⁶⁰ Compared with the resting period, fetuses showed reduced motor activity and slower HR within 10 seconds after the onset of maternal speech. Subsequent analyses showed that the fetal response was modified by both maternal and fetal factors. The behavior of fetuses whose mothers were previously awake and talking was analyzed; a period of fetal orientation can be measured when a mother starts reading aloud. Compared with women who rested and remained silent, the fetuses reacted with increased HR and movements. The fetal response was also dependent on the initial variability of the fetal HR; the greatest response was shown during periods of low variability, when the mothers had been resting and were silent. The results indicate that fetal response is influenced by both maternal and fetal conditions, which has an impact on understanding the fetus's learning of the mother's voice in neutral settings.⁶⁰ The reaction to sounds was also compared with those of premature infants. In high-risk infants, the cardiac and motor responses were recognizable at 30 weeks' gestation. At 33 weeks, high-risk infants showed increased HR compared with low-risk fetuses. The results indicate that both

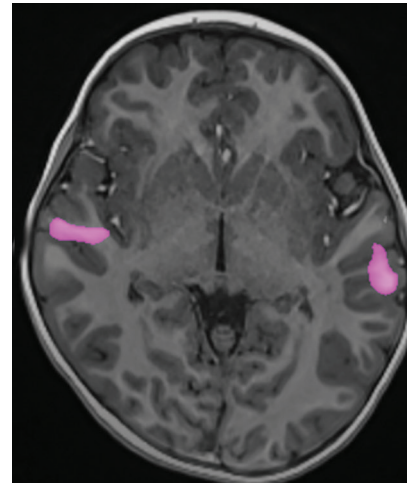


Fig. 1: Sounds activate specific temporal lobe regions in children: A passive-listening functional-magnetic resonance imaging (fMRI) task comprised of a series of words and non-language noise induced bilateral activation in the posterior superior temporal gyri. This particular fMRI image is from a 5-year old with a hypothalamic hamartoma who underwent fMRI for surgical planning; sounds were delivered using headphones. The image shows that we now have the technology to study the impact of sounds on specific areas of the brain and can study developmental changes in young infants

low- and high-risk premature infants begin to respond to sounds at the same gestational age. The varied responses observed during pregnancy in the high-risk group most likely indicate diversified functional development of the auditory system.⁶¹

The maturation of the fetal response to music was characterized in the last trimester of pregnancy using a 5-minute piano recording of Brahms' Lullaby played at an average volume level of 95, 100, 105, or 110 dB.⁶² Within 30 seconds of turning on the music, the youngest fetuses (28–32 weeks of gestation) showed an increase in HR limited to the two highest dB levels. During pregnancy, the threshold level decreased and a shift in response from acceleration to deceleration was observed at lower dB levels, indicating attention to the stimulus. During 5 minutes of music, fetuses over 33 weeks of gestation showed a sustained increase in HR. Changes in body movement occurred at 35 weeks of pregnancy. These findings suggest a change in the processing of complex sounds around the 33rd week of gestation, with response limited to the acoustic properties of the signal in younger fetuses, while episodes of focusing were observed in older fetuses.⁶²

An attempt was also made to examine the impact of lullabies sung by mothers on the bond and behavior of newborns. There was no significant effect on prenatal attachment. Only the postnatal bond was significantly greater in the group of singing mothers 3 months after birth. In the same group of singing mothers, episodes of newborn crying in the first month, as well as infantile colic, were significantly less common. At the same time, a reduction in the frequency of night awakenings was observed. It can be concluded that singing lullabies by mothers can improve the bond between mother and child. It may also have a positive impact on the behavior of newborns and thus reduce the stress level in the young mother (Fig. 1).⁶³

The Mozart Effect

There are two complementary communication systems in humans—language and music and functional imaging studies suggest these

are processed in nearby brain regions. Music affects many aspects of human behavior, encourages social interactions and promotes trust and cooperation within groups of culturally compatible individuals. Music acts on the limbic system, is rewarding and motivating, and can facilitate learning and memory.^{64–66}

Oxytocin is vital to the attachment between infants and parents through early contact and interaction can influence developing brain functions in infants culminating in behavioral changes in the child.^{67,68} Maternal behavior is mediated by oxytocin, a neuropeptide synthesized in the paraventricular nucleus (PVN).^{69–71} The central nucleus of amygdala (CeA) expresses oxytocin receptors and interacts with the reward circuit to motivate maternal behaviors.^{72–74} Experiments in rats have shown that maternal behaviors are influenced by environmental factors in pregnancy; being reduced by stress and enhanced by enriched environments. The “Mozart effect” influenced the licking of offspring by mother rats and such offsprings had decreased behavioral anxiety and stress responses as adults.^{75,76} Increased maternal oxytocin levels lead to greater affectionate contact behaviors in mothers following mother–infant contact.⁶⁷ The changes in rearing behaviors due to music are due to modulation of the oxytocinergic system by activation of oxytocin receptors and neuromodulation of the oxytocinergic system.^{72,74,77}

Placental Programming

Hereditary and environmental stimuli are assimilated as placental epigenetics which influence the fetal development. Fetal brain development in the third trimester is impacted by plasticity and hence neurobehavioral phenotype can be altered by prematurity. Prenatal music stimulation has a positive epigenetic effect on fetus by virtue of placental programming.⁷⁸

The auditory cortex is located in the posterior-medial part of Heschl’s gyrus. This region corresponds with the Brodmann’s area 41, a highly plastic epigenetic area vital to prenatal learning.^{79,80} Acoustic environments influence the functional organization and processing capabilities of the auditory cortex.^{81,82}

Music Therapy and Memory

Musical intervention has a therapeutic role in dementia or Alzheimer’s disease because music elicits feelings and memories.⁸³ Music therapy induces plastic changes in brain networks, thereby facilitating brain recovery processes and modulation of emotions and communication.⁸⁴ Hence, it is a propitious modality of rehabilitation.

It has been reported that listening to classical music, specifically selections from Mozart can result in a temporary improvement in cognitive functions like abstract/spatial reasoning tests.⁸⁵ The “Mozart Effect” is attributed to the arousal due to the pleasure of listening to music, rather than a direct impact on cognitive ability.^{86,87}

Influence of Music on Fetal Cardiocardiographic Parameters

Although music has long been recognized for its effect on human emotions, physiological responses, and overall well-being, the mechanisms remain largely unknown.⁸⁸ Machin and Dunbar proposed that activation of the endogenous opioid system, including β -endorphins and enkephalins, which are known to foster and maintain social bonds, improve mood and reduce the sensation of pain, are a possible mechanism.⁸⁹ In women with a

low- or high-risk pregnancy, and in non-stress tests (NSTs), music appeared to reduce maternal anxiety.^{90–92}

During pregnancy, maternal stress can affect fetal development, including altered fetal HR patterns.⁹³ In a recent systematic review, Shimada et al.⁹⁴ noted improved maternal relaxation, decreased anxiety, psychosocial stress and depression, reduced pain, increased maternal bonding with her fetus, improved sleep quality, improved blood pressure profiles, lower fetal HR, and lower need for medications after surgery. Music therapy during the pre- and perinatal periods may be benefit both the pregnant women and their newborns infants. Maul et al.⁹⁵ performed a systematic review to assess maternal mental health outcomes after musical intervention from 14 randomized controlled trials (RCTs) including 2,375 pregnancies. They showed reduced maternal stress, anxiety, and depression. These data are consistent with the beliefs that modulation of maternal physiological parameters through music may influence fetal well-being.⁵

Neural processing of music involves an extremely complex and extensive network of cortical and subcortical structures which integrate auditory, sensory motor, and cognitive functions as well as emotional changes.⁴ As expected from our understanding of the neurosensory development of the fetal auditory system, initial responsiveness to different sound frequencies begins around 23 weeks’ gestation. By 24 weeks, fetuses demonstrate a startle response to vibroacoustic stimulation. These responses are seen consistently at 28–30 weeks.^{33,96,97} Indeed, music may improve fetal autonomic responses and learning/memory formation.^{30,32,98}

EVIDENCE FROM RCTs

Effect of Music Exposure on FHR

In a prospective RCT, James et al.⁹⁹ examined whether prenatal exposure to a musical stimulus altered fetal behavior and whether these responses persisted after birth into the newborn period. Using an exposure-learning model, music was played to 10 fetuses via a headphone on the maternal abdomen and 10 controls had the headphone without sound. All fetal studies took place within 72 hour prior to elective delivery. After delivery, all 20 newborns were exposed to the same music on postnatal days 3–5. Computerized assessment of FHR and activity was documented and neonatal behavioral states were recorded. For the 1st hour of the study, neonates that had had *in utero* exposure showed higher mean HR and showed longer periods of high HR variation. These findings, however, were not statistically different from controls. However, by the 4th hour, the exposed fetuses began to show significantly more HR variation ($p = 0.04$) and more state transitions ($p = 0.01$) compared with unexposed fetuses. These effects persisted into the neonatal period with the same music stimulus evoking more state transitions ($p = 0.01$) and leading to longer awake states ($p = 0.05$). Thus, prenatal exposure to music altered the fetal behavior that persisted after birth.

Granier-Deferre et al.¹⁰⁰ showed that repetitive prenatal exposure to specific melodies influenced neonatal auditory perception and memory that was retained for 3–4 days to 6 weeks. In the test arm of the study, fetuses were given precisely controlled exposure to a descending piano melody twice a day during weeks 35–37 of gestation. After 6 weeks, cardiac responses of 25 exposed infants and 25 naive control infants to the descending melody and to an ascending control piano melody were examined during quiet sleep. The melodies had precisely inverse contours, but similar spectra, identical duration, tempo and rhythm, and thus, nearly identical

amplitude envelopes. All infants displayed a significant change in HR. In exposed infants, the descending melody evoked a cardiac deceleration that was twice larger than the decelerations elicited by the ascending melody and by both melodies in control infants.

Brillo et al.¹⁰¹ randomized 30 healthy mother–fetus dyads in a 1:1:1 ratio to one of three groups: (1) fetuses were submitted to pre-listening phase (33⁰–36³ week) and listening sessions during 4 NST; (2) fetuses were submitted to listening sessions during 4 NST only; and (3) 4 NST without any listening. Mean fetal HR, fetal HR accelerations/decelerations, fetal movements, and uterine contractility were assessed. The 1st group fetuses, who had heard a particular piece of music during previous sessions, showed significantly increased HR accelerations and movements during the music listening session of the last NST. Uterine contractions did not change in frequency. They concluded that fetuses respond to familiar but not to unknown music.

Catalgol and Ceber Turfan randomized 100 (50 intervention, 50 control) primipara women. The NST was applied in 36–38 weeks' gestation.¹⁰² During the test, music was played to the intervention group, while the control group received routine care. The music group showed lower mean scores of State Anxiety Inventory during NST. Acceleration, mean number of fetal movements and fetal HR reactivity were significantly higher in the intervention group. Thus, music therapy in pregnant women decreased maternal anxiety and had positive effects on NST findings.

Soylu et al.¹⁰³ showed that music affected vital signs, fetal movements, and lowered the state and trait anxiety levels during NST in pregnant women. In 74 (37 music and 37 control group) pregnant women, post-music exposure HRs were lower than the pre-procedure values ($p < 0.001$). The groups did not differ in baseline fetal HR, variability, fetal movement, presence/number of accelerations-decelerations, and NST parameters. The number of fetal movements was higher in the music group than controls ($p < 0.001$). The state anxiety inventory scores were lower in the music group than controls ($p < 0.001$).

In a cross-over RCT, Oh et al.¹⁰⁴ tested the effects of musical intervention on maternal anxiety, fetal HR, and testing time during NST. Sixty pregnant women in 28–40 weeks' gestation were randomly assigned to either an experimental ($n = 30$) or a control group ($n = 30$). The experimental group showed significantly lower scores in state anxiety than controls. There two groups showed no difference in systolic blood pressure and HR. Baseline fetal HR was significantly lower and frequency of acceleration was significantly increased in the experimental than in the control group.

Estrella-Juarez enrolled 343 full-term pregnant women in a RCT and divided them into three parallel groups: (1) music therapy intervention ($n = 104$), (2) virtual reality intervention ($n = 124$), and (3) controls ($n = 115$).¹⁰⁵ The interventions were delivered during NST in the third trimester and during labor. Measures included the Spielberger State-Trait Anxiety Inventory, maternal blood pressure, maternal and fetal HRs, and labor and birth outcomes. Women in the music therapy and virtual reality groups had less anxiety after NST ($p < 0.001$), and the women were more likely to have a reactive NST ($p < 0.001$) than controls. Following completion of NST and intervention, music therapy and virtual reality groups had lower systolic blood pressure ($p < 0.001$), diastolic blood pressure ($p < 0.001$), and maternal HR ($p = 0.003$) than controls. Furthermore, fetuses in the control group were more likely to show non-reassuring fetal HR tracings than those in the music therapy and virtual reality groups, respectively ($p = 0.004$).

Effect of Prenatal Music Exposure on Neonatal ECG and Neuro-behavioral Response

In a pilot study, Lang et al. showed that newborns displayed distinct reactions to maternal voice at 2 and 5 weeks after birth on a physiological level and identifiable with ECG and EEG changes.¹⁰⁶ Basic memory traces were formed *in utero* and shaped neonatal autonomic and neuronal reactions to speech and voice stimuli. Newborns exposed to nursery rhymes prenatally showed distinctly different reactions than those not exposed. The authors concluded that fetal brain is “programmed” for the predicted postnatal environment and maternal voice.

In an open-labeled RCT, Arya et al.³ evaluated the effects of antenatal music exposure in healthy primigravidae on the behavior of their term appropriate-for-date newborns, assessed using 7 clusters of the Brazelton Neonatal Behavioral Assessment Scale (BNBAS). Primigravida mothers aged 19–29 years who had a singleton pregnancy, at ≤ 20 weeks' gestation, and did not have any chronic medical diseases or significant hearing impairment, were randomized to listen to a pre-recorded music cassette for approximately 1 hour/day in addition to standard antenatal care (intervention arm) or standard care only (control arm). Perinatal factors with adverse effect on neonatal behavior were deemed as protocol violations. One hundred and twenty-six newborns (music group) and 134 (controls) were tested. Infants of mothers exposed to music during pregnancy performed significantly better on 5 of the 7 BNBAS clusters. The maximal beneficial effect was seen with respect to orientation (ES 1.13, 95% CI: 0.82–1.44, $p < 0.0001$) and habituation (ES 1.05, 95% CI: 0.53–1.57, $p = 0.0001$). Prenatal music exposure to mother significantly and favorably influenced neonatal behavior.

As seen above, many systematic reviews were of mixed methodological quality and showed ambiguous efficacy of auditory stimulation of preterm infants. Hence, a meta-analysis of several studies was performed.^{107–132} The authors evaluated the impact of parallel and cluster-RCTs on preterm infants <37 weeks' gestation during hospitalization, and also studied the effects on parents who were involved in the intervention. The evaluated interventions included any live/recorded music or vocal stimulation for >5 minutes, administered >3 times, by a music therapist, a parent, or a healthcare professional, and these were compared with standard care. Many studies had to be excluded from this analysis because of inadequate data.^{42,133–191}

The authors studied 25 trials including 1,532 infants born at 25–36 weeks' gestation and 691 parents (21 parallel-group RCTs, 4 cross-over RCTs). The intervention did vary in type, delivery, frequency, and duration; music and voice that were typified as calm, soft, musical parameters in lullaby style were often integrated with mother's voice. There was considerable variability in the risk of bias in included studies. Music/vocal interventions reduced HRs in infants during intervention (mean difference, MD, -1.38 , 95% CI: -2.63 to -0.12 ; $p = 0.03$; 1014 infants; 11 studies; moderate-certainty evidence) and after intervention (MD -3.8 , 95% CI: -5.05 to -2.55 ; $p < 0.00001$; 903 infants, 9 studies; high-certainty evidence). There were no reported adverse effects. There was no change in oxygen saturations during care. Similarly, there was no difference in infant development (Bayley Scales of Infant and Toddler Development with the cognitive composition score; motor composition score; and the language composition score). Parents showed no difference in the incidence of anxiety or depression.



Evidence from Non-RCTs

Kumar et al.⁴¹ examined the impact of music therapy for fetomaternal monitoring in nine pregnant women. They found that fetal HR increased from 146 to 169 beats per minute after exposure to music from headphones paced around the mother's abdomen. The increase in HR was consistent with a reactive NST. Music could serve as a means of communication with the fetus through sounds and voices. Caressing the fetus through the abdomen, producing soft and melodic sounds, using lights and vibrations were pleasant for the fetus.

In a prospective observational study in 100 uncomplicated pregnant women, between 32 and 41 weeks gestation, Erkun Dolker and Basar showed significant differences between the music-exposed and control groups in terms of mean numbers of acceleration, deceleration, and reactive NST results ($p = 0.001$).¹⁹²

Gebuzza et al.¹⁹³ showed a significant reduction in the number of uterine contractions, increased FHR accelerations and variability and increased fetal movements following exposure to classical music during NST in 48 pregnant women in their third trimester. Their recent study involving 30 (classical music during NST) and 60 (NST only) pregnant women between 27 and 41 weeks of gestation, confirmed their previous findings that classical music therapy could be a non-invasive, affordable method to improve fetal well-being.^{193,194}

A Polish study showed a positive correlation between uterine contractility, fetal movements, and increasing music rhythm. There was no influence on fetal HR.¹⁹⁵

Lee et al.¹⁹⁶ reported that musical intervention on fetal HR altered fetal Doppler between 30 and 38 weeks' gestation. Mean (\pm standard deviation) fetal movements during fixed singing activity (0.7 ± 0.79) were lower than during irregular singing (1.73 ± 1.37). This showed that fetal movement and HR could be stabilized by singing fixed songs to promote well-being and establish maternal-fetal bonding.

FUTURE DIRECTIONS

We still need more evidence to ascertain the precise role of timing of prenatal sound stimulation. Methodologically strong RCTs involving this concept with rigorously designed interventions, consistent outcomes, and standardized reporting measures are needed. Future studies may include comparisons of different durations and types of musical (sound) intervention. Future studies can focus on specific effects of different types of music including instrumental music compared with vocals, classical vs metal and maternal vs other voices. It has been shown to promote maternal-fetal bonding.^{49,79} Studies are needed for better understanding of the mechanisms by which music affects the growing brain, intrauterine neurosensory organization, and changes during development.¹⁹⁵ Changes to the physical environment, like darkening the room and decreasing visual and auditory stimuli in the infant's surroundings.

Music therapy may also be useful in specific maternal/fetal conditions such as drug dependence given the beneficial effects of music therapy in infants with neonatal abstinence syndrome postnatal conditions. Neonatal abstinence syndrome is potentially an important area for study; the numbers in the United States are staggering.^{197,198} The incidence of NAS in the US started increasing in 2000 and nearly doubled between 2010 and 2017.^{199,200} We need carefully conducted studies to determine the prenatal impact of maternal substance use on the fetus.²⁰¹

CONCLUSION

Prenatal sound stimulation including music and mother's speech influences the development and function of the fetal/neonatal neurological system. Fetuses seem to respond to various forms of music that they have been exposed to. Further studies can be beneficial.

ORCID

Adrianna Frydrysiak-Brzozowska  <https://orcid.org/0000-0003-2547-7748>

Srijan Singh  <https://orcid.org/0000-0002-2103-5232>

Akhil Maheshwari  <https://orcid.org/0000-0003-3613-4054>

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