# The Effect of Music Therapy Applied to Neonatales on their Pain: Systematic Review

Omer Terzi<sup>1</sup>, Gulay Manav<sup>1</sup>, Pelin Uymaz<sup>2</sup>

<sup>1</sup> Mugla Sıtkı Koçman University, Health Science Institute, Turkey <sup>2</sup> Keykubat University, Turkey

Pain is emotional and psychological emotional state that an individual feels as a result of a tissue injury in his/her body along with past experiences. Although individuals can express this emotional state, infants cannot. Instead, infants show pain with physiological or behavioural symptoms. Behavioural symptoms can appear as crying, facial expressions and motor movements, while physiological symptoms can appear as changes in heart rate, an increase in the number of breaths and change in the oxygen value. Various measurement tools have been developed to measure pain. In total, more than 40 scales have been developed for use in premature, new born and post mature babies. The treatment of the pain evaluated with these measurement tools is performed by pharmacological and nonpharmacological treatment methods. Pharmacologically, opioid, non-opioid and cholangitis are used; while breast milk, kangaroo care, sucrose application, massage and voice applications are used as nonpharmacological methods. Studies have recently been carried out where nonpharmacological techniques are more prominent. SES applications, which are one of the nonpharmacological methods, have become widespread in intensive care units in recent years. This study aimed to reveal the effect of this prevalence in the last 20 years (2002-2022) and determine the effect of sound applications on pain in premature infants. In the research conducted in 16 databases with keywords, a total of 9301 articles were accessed. Among them, a selection was made according to the inclusion criteria and 17 articles were selected for the study. Of these articles, 82.3% (n=14) yielded positive results in audio applications. In addition to pain assessment, positive results were obtained by increasing SpO2 value, stabilization of heart rate, decrease in respiratory rate and shortening of crying time in studies. However, metaanalysis studies need to be carried out for voice applications to be called positive or negative.

Key words: Neonatal nursing, neonate, pain, voice application

Address for correspondence:

Dr.Pelin Uymaz, Keykubat University, Turkey, pelin.uymaz@alanya.edu.tr

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### INTRODUCTION

Pain is an emotional and psychological mood, which occurs due to harmful stimuli. We can understand that infants feel pain through physiological symptoms such as change in their heart rate, increase in the number of breaths and change in the oxygen value or behavioral symptoms such as crying, facial expressions and motor movements. However, it is not easy to measure pain in neonates [1].

There are more than 40 scales used to measure the state of pain in infants, yet four of them are widely used. These are Premature Infant Pain Profile (PIPP), Neonatal Infant Pain Scale (NIPS), CRIES and FLACC pain scales [1,2].

Treatment of pain evaluated through these measurement tools is done with pharmacological and nonpharmacological treatment methods. Pharmacological methods are opioid, non-opioid and cholangitis; while nonpharmacological methods are breast milk, kangaroo care, sucrose application, massage and music therapy. Recent research has been conducted indicating that nonpharmacological methods outweigh [3].

Music therapy, one of nonpharmacological methods is diversion of the infant's attention by making it listen to the mother's voice, white noise, lullaby or musical instruments [4,5]. In such an application, infants showed recovery as decrease in pain, and increase in comfort and vital findings [6]. The mother's heartbeat sound and her own voice were used for the relief of pain in neonates and found to be effective [4]. In an RCT study conducted with 60 premature infants, the oxygen saturation value of 30 infants who listened to music during peripheral central venous catheter insertion increased, and heart rates and pain scores decreased compared to the control group. Another RCT study, using white noise, the PIPP score, heart rate and respiratory rate decreased, while oxygen saturation value increased [7]. In an RCT study that listened to heart rate, there was an increase in SpO<sub>2</sub> value and a decrease in PIPP pain score of the experimental group [8].

The literature review showed that Standley (2002) conducted a meta-analysis study by scanning studies from 1964 to 1999 to examine the effect of music therapy on pain in premature infants. After the year 2002, there was neither systematic review nor meta- analysis study. In recent years, the use of voice applications in intensive care units has become prevalent. To see how effective this prevalence has been, the present study is designed to evaluate the effect of voice applications on pain in premature infants over the last 20 years (2002-2022). The study included case-control

studies conducted in the last 20 years. This systematic review FLACC PAIN SCALE aimed both to determine the effectiveness of voice applications in premature infants and to be a guide to new research topics.

### PREMATURE INFANT PAIN PROFILE (PIPP)

It was developed by Steven et al. (1996) for premature infants aged 28 weeks-36 weeks. Its Turkish validity and reliability study Nursing practices and voice applications in pain was conducted by Akcan and Yigit (2015). It is applicable to preterm and term infants and is scored between 0 and 21. A score of 0 points-6 points indicate that there is no pain or it is minimal, 7 points-12 points indicate that there is moderate severe pain and 13-21 points indicate severe pain. It is used as an assessment 15 seconds before and 30 seconds after the painful procedure (Table 1) [9].

### Neonatal Infant Pain Scale(NIPS)

It was developed by Lawrence et al. (1993) and its Turkish validity and reliability study was conducted by Akdovan (1999) [10,11]. It is appropriate for the assessment of pain in premature and neonates. The obtainable score from the scale consisting of six deaths in total varies between 0 and 7. A higher score indicates more severe pain (Table 2).

### CRIES PAIN SCALE

This scale, developed by Krechel and Bildner (1995), evaluates postoperative physiological pain [12]. It works through a system similar to Apgar score. It is used for over 32 weeks of gestational age and postoperative period infants. The lowest obtainable score is 0, while the highest is 10 points. A score of at least 4 points and over indicates pain (Table 3).

Legs

Wakefulness

Pain measurement is carried out through evaluation of five behavioural categories developed by Merkel et al. in 1997 [13]. It is used for term, preterm and non-expressive children in the age range of 3 years-7 years (Table 4) [14].

## management of neonates

Music has been accepted as a way of therapy for centuries [15]. It is believed that music is used in therapy because it affects people in many aspects and it is easy to use [16].

Music therapy is defined as "the use of music and/or musical elements (rhythm, sound melody) designed and used by a trained music therapist to optimize and improve the quality of life of a person, family or group" by the World Federation of Music Therapy [17].

Pythagoras (585 BC-500 BC) was one of the first founders of music therapy. Research has been conducted on treatment of psychiatric patients by playing rhythmic sounds. Plato (400 BC) pointed that music ensures tolerance and comfort in people [18]. In his work Kitabu-s Sifa (The Book of Healing), Avicenna referred to music in medical field as "one of the most effective ways of treatment is to increase the mental and spiritual powers of the patient, to give him/her the courage to fight the disease better, to make him listen to the best music and to bring him together with the people he loves" [19].

Nightingale described music therapy in the 1800s as a nursing initiative that increased patient comfort by reducing pain and anxiety [16]. Today, it is frequently used in the treatment of pain and anxiety particularly among the symptoms that occur in other

flexion

Eager, uneasy and unappeasable

Tab. 1. Premature Infant Pain Profile-(PIPP)	Categ	ories	0		1	2	3
	Costatio	nal ago	nalage ≥ 36 weeks 32		eeks - 35	28 weeks - 31	. <28 weeks
	Gestational age		≥ 30 weeks	week	s 6 days	weeks 6 days	<20 weeks
			Active/awake,	Calm/a	wake, eyes	Active/dormar	it, Active/dormant,
	Behavior	al status	eyes open, facial	open,	no facial	eyes closed, fac	
			movements	mov	ements	movements	facial movements
			An increase of 0-		ease of 5-	An increase of 1	
	Max he	art rate	4 heart rate per		rt rate per	24 heart rate p	
			minute	m	inute	minute	per minute
	Min ownon	caturation	Decrease by 0-	Decrea	se by 2.5-	Decrease by 5	- Decrease by 7.5%
	Min oxygen saturation		2.40%	4.	.90%	7.40%	and more
	Wrinkling forehead Squinting		No	S	light	Moderate	Too much
			No	S	light	Moderate	Too much
	Enlargement wir		No	S	light	Moderate	Too much
Tab. 2. Neonatal Infant Pain Scale (NIPS)	Catagoni	I	0			1	2
	Category		0			1	2
	Facial expression	Calm face, natural expression			Tense facial muscles, wrinkled forehead and chin		
	Crying	Silent, no crying			Howling and intermittent Sci crying		Screaming, constant loud crying
	Respiration type	Regular routine respiration		on	Unstable and irregular respiration, sigh		ar respiration, sigh
	Arms	No muscular rigidity, frequently random arm gestures			om Stretched, straight arms, hard or fast extension/ flexion		
	legs	No muscular	rigidity, frequently	random	Stretched, straight legs, hard or fast extension/		

leg gestures

Silent, sleeping peacefully or calm

Tab. 3. Cries pain scale	Categories		1	2	3		
	Crying O, need		No	Loudly	Unstoppable		
			O <sub>2</sub> need		No	<%30	>%30
	Increase in vital signs		10%	%11-%20	more than x%21		
	Appearance Insomnia		Appearance	Good	Grimace	Grimace and howling	
			No	Wakes up frequently	Constantly awake		
Tab 4. Flacc pain scale	Categories	0 No special expression		1	2		
	Face			Frowning, souring face	Grimace, clenching		

Legs         In normal position         Stretched, disturbed         Kicking here and there           Activity         Calm         Roll back and forth         Twirling like a bow, tossing           Crying         No         Howling         Crying screaming aloud           Consolability         Comfort         Consoling by hugging and touching         Inconsolable	Face	No special expression	Frowning, souring face	Grimace, clenching
Crying         No         Howling         Crying screaming aloud	Legs	In normal position	Stretched, disturbed	Kicking here and there
	Activity	Calm	Roll back and forth	Twirling like a bow, tossing
Consolability Comfort Consoling by hugging and touching Inconsolable	Crying	No	Howling	Crying screaming aloud
	Consolability	Comfort	Consoling by hugging and touching	Inconsolable

interventional diagnosis-treatment phases applied to patients studies published between 2002 and 2022 that examined the in intensive care, psychiatry, surgery, pediatrics, obstetrics, and effect of voice application on pain in premature infants. oncology chemotherapy stage [20].

In a study that evaluated 29 research results, music was used to reduce pain and anxiety in patients, to eliminate the side effects We reviewed the electronic databases of Google Akademik, of treatment and to increase patient satisfaction [21]. Loewy et al. ULAKBİM, Clinical Key, DynaMed, Elsevier, Ebsco, Asos Index, (2013) examined the effect of music on 272 premature infants and HiperKitap, ProQuest, Sage Premier Journals, Science Direct, found that their heart rate decreased, while their sleep patterns and Scopus, Springer Nature, Turcademy, Web of Science for the milk absorption increased through music therapy [22]. Clacatera et al. (2014) played musical sounds to 42 infants hospitalized in the pediatric surgery unit and found a positive change in their Data collection process cardiac parameters with reduced perception of pain. The mother's heartbeat sound and voice were used for the pain relief in neonates and found to be effective [4]. In an RCT study conducted with 60 premature infants, the oxygen saturation value of 30 infants who listened to music during peripheral central venous catheter insertion increased, and heart rates and pain scores decreased compared to the control group. Another RCT study, using white noise, the PIPP score, heart rate and respiratory rate decreased, while oxygen saturation value increased [7]. In an RCT study that listened to heart rate, there was an increase in SpO<sub>2</sub> value and a decrease in PIPP pain score of the experimental group [8]. Nine RCT studies on heel lance and circumcised infants found that pain responses reduced in infants who listened to music [23]. In another RCT study, the study group listened to white noise while the control group did not. The study results showed that the The coding form was created by the researcher and consisted of level of pain in the infants in the control group was significantly 60 preterm infants in the neonatal intensive care unit, the study group had listened to music for 30 minutes twice a day for seven days. There was a significant difference in physiological parameters Ethics of the study group and the control group with an increase in SpO<sub>2</sub> level and decrease in respiratory rate and in heart rhythm [24].

This study aimed to determine the current nursing practices for the elimination of infants' pain and to update the future practices so that nurses can develop a professional care in the pain care of Limitations of the study neonates.

### METHOD

The research is a systematic review.

### Population and sample of the research

The population of our study consisted of randomized controlled

### Data collection

present study.

The MeSH (Medical Subjects Headings) system was used to determine the keywords. These keywords, prepared in Turkish and English were as follows: "premature", "pain", "music", "white noise", "lullaby", "agrı", "muzik", "beyaz gurultu", "ninni", "anne sesi'. The keywords were searched in combinations in the keywords, title and abstract parts. The articles accessed by such screening were included in the study. This process was ended on January 1, 2022 accessing 9301 articles and including 17 of them in the study. The elimination phases of the studies are indicated in the PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-Analyses) 2020 flow diagram (Figure 1).

### Coding the study

seven sections, including author/year, country/city, research aim, higher than in the study group [5]. In another study conducted on research method/randomization, sample, intervention and study results. The coding was done using Microsoft Office Word.

The literature was reviewed and studies were found as the research was a systematic review. Therefore, no ethics committee approval was obtained because it was not necessary.

The studies were selected through participants (P: Population), interventions (I: Interventions), comparison groups (C: Comparators), results (O: Outcomes) and research methods (S: Study Designs)

- Population: Infants born before the week 37.6.
- Interventions: Voice application to the infant.

- Comparator: Infants in the control group.
- Physical vital signs (pulse, saturation, respiratory rate, etc.)
   and pain assessment scales (PIPP, NIPP).
- Studies with randomized control groups.

### Inclusion Criteria:

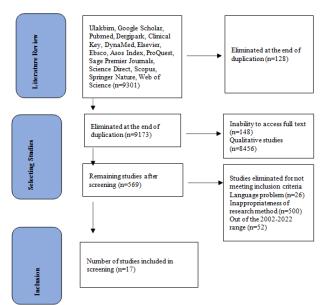


Fig. 1. PRISMA 2009 Flow Diagram

- Written in Turkish or English
- Being published between 2002 and 2022
- Inability to access full text of the studies
- Using the RCT methodology

### Exclusion criteria:

Congress declarations, non-experimental studies and qualitative studies.

### Reporting

The systematic review and meta-analysis control list of PRISMA published in 2020 was used in the study.

### FINDINGS

With the keywords determined, a total of 9301 studies were accessed between 2002 and 2022 through the screening in 16 databases. Of these studies, 128 were eliminated due to duplication, 148 were not in full text, 8456 were qualitative studies, 26 did not have language compatibility, 500 were not suitable for the method and 52 were out of the specified year range, thus the study group was created with the remaining 17 studies.

The sample group of 17 studies included in the systematic review consisted of preterm infants. The sample size ranged from a minimum of 25 to a maximum of 206. All studies selected were

<b>Tab. 5.</b> Coding tables of the studies included in the study	Author / Year	Country / City	Research Aim	Research Method / Randomiza- tion	Sample	Intervention	Study Results
	Ren et al., 2022	China	To examine the effects of white noise on cortical response due to pain, pain score, and be- havioral and physiological parameters in	A random- ized con- trolled trial	- Sample: 60 Sample: 60 Samp		During artery punc- ture, the mean rScO <sub>2</sub> , HR and SpO2 did not show difference between the groups. After the needle was taken, there was no significant difference in the rScO <sub>2</sub> , PIPP-R score, and trends for facial expression's tendency to return to the reference line. The white noise inter-
			newborns with procedural pain.			Cameras.	vention did not show a beneficial impact
				Double blind	Experimental group: 29		on pain.
					Control group: 31		
	Yu et al., 2022	Taiwan	To analyze the efficiency of maternal voice in relieving pain during heel lance of premature infants and in facilitating mother- infant bonding dur- ing hospitaliza- tion.	RCT	Sample: 64	Audio recordings of a mother reading a children's book were created and then played for the infant during the heel lance once a day for three days in a row. Heart rate, respiratory rate, oxygen saturation and pain response were evaluated before, during after the intervention.	One minute after the intervention, heart rate (p<.001) and Neonatal Infant Pain Scale score (p<.001) were lower in the intervention group than in the control group.

Sharara- Chami et al., 2022     To examine the effect of madionistration administration and signifi- cantly higher mean al- mean tate and in- creased crying time besides the analgesit to the infants to be circum- cised.     RCT     Sample: 206     The infants were vid- ectaged during the intervention to assess pain by double bill and and independent reviewers (face and body).     The infants were vid- ectaged during the intervention to assess pain by double bill and independent reviewers (face and body).     The infants were vid- ectaged during the intervention to assess pain by double bill and into reduce pain during the interven- room did not reduce infants were observed in their nest for 15 min- utes with a heart rate device and the control proma source during the infants were observed in their nest of 15 min- device.     During the interven- tion, the Spo, value infants were observed in their nest for 15 min- device.       Turkey 2021     To examine the effect of similation nest use on vital signs, pain level and correl and indice ent tates and on pain level and correl and lullables on pain perform.     RCT     Sample: 52     The texperimental group values during the effect of similates without any device.     During the interven- tion, the Spo, value infants during blook were the anal oxygen saturation rate infants during painful inter- ventions.       Turkey Buyuk, 2021     To examine the effect of white noise pain during the operation rate and lullables on pain per- mean heart rate and recorded before, during painful inter- ventions.     RCT     Sample: E6 Experimental group (White noise): 22     Premature infants ing the PIP.       Experimental group (white pain was evaluated us- ing the PIP.     Free main were feacthe the pain score was lowed in the white noise group wal					Experimental group: 32 Control group: 32		The intervention carried out with the mother's voice slowed the heart rate and relieved the pain response of the pre- mature infants.
Experimental group: 103     Experimental group: 103       Karadag et al., 2022     Turkey     To examine the effect of simulation heartbeat om vital signs, pain level and comfort in preterm.     RCT     Sample: 52     The experimental group was observed for 15 minutes without any device.     During the interven- tion, the SpO, value increased significantly (p<.003) while the PIPP score (p=.001) decreased in the ex- perimental group.25       Dora and and Tural Buyuk, 2021     Turkey     To examine white noise and lulables on pain per- ventions.     RCT     Sample: 52     Heart rate, respiratory rate and oxygen saturation were recorded.     Premature infants in the white noise group had the low- perimental group (white noise and lulables and fullables and fullables and fullables and fullables and fullables and fullables     RCT     Sample: 66     Heart rate, respiratory rate and oxygen saturation rate and proprocedure pain was evaluated us- infants during plood control group (white noise): 22     Premature infants in the white noise group than in the	Sharara- Chami et al.,	the effect of the adding music on pain besides the administration of combined analgesic to	the effect of the adding music on pain besides the administration of combined analgesic to		Sample: 206	eotaped during the intervention to assess pain by double blind and independent reviewers	group had a signifi- cantly higher mean heart rate and in- creased crying time than the control group (p=.00). Playing music sound from an lpad in the operation room did not reduce pain during circumci-
Karadag et al., 2022TurkeyTo examine the effect of wital signs, pain level and comfort in preterm.RCTSample: 52The experimental group infants were observed for 15 min- utes with a heart rate device and the control group was observed for 15 minutes without any device.During the interven- tion, the SpO, value (p<.003) while the pP score (p=.001) decreased significantly (p<.003) while the eprimental group: 25During the interven- tion, the SpO, value (p<.003) while the device.Dora and Tural and TurkeyTo examine the effect of white noise and lullables on pain per- ception and vital signs of premature infants during painful inter- ventions.RCTSample: 66Heart rate, respiratory rate and oxygen satura- tion and post- procedure ing the PIPP.Premature infants in the white noise group had the low- est mean PIPP score, mean heart rate and recorded before, during and after the interven- tion and post- procedure pain and the pain score was lower in the white noise group than in the uten onise i: 22 Experimental group (lullaby): 22Premature infants during pain and the pain score was lower in the white noise group than in theDora and tural buryuk, 2021TurkeyTo examine the effect of white noise premature infants during painful inter- ventions.RCTSample: 66Heart rate, respiratory rate and oxygen satura- infants during blood ing the PIPP.Premature infants infants during blood ing the PIPP.Lillables ing the PIPP.Experimental group (lullaby): 22Iullaby group.Lill				Double			
Karadag et al., 2022To examine the effect of simulation heartbeat nest use on vital signs, pain level and comfort in preterm.RCTSample: 52infants were observed in their nest for 15 group was observed for 15 minutes without any device.During the interven- tion, the \$p0, value (p<003) while the PIPP score (p=001) decreased in the ex- presend				blind			
Dora and and Turala 2021       To examine the effect of white noise and lullabies on pain per- ception and Suyuk, 2021       To examine the effect of white noise and lullabies on pain per- ception and surve painful inter- ventions.       RCT       Sample: 66       Heart rate, respiratory rate and oxygen satura- tion were measured and recorded before, during pain and after the interven- tion and post-procedure pain was evaluated us- ing the PIPP.       Premature infants in the white noise group had the low- est mean OXP- ore and lullabies played to premature infants during painful inter- ventions.         Experimental group (white noise): 22       Experimental group (ullaby): 22       Experimental group (ullaby): 22	et al.,	Turkey	the effect of simulation heartbeat nest use on vital signs, pain level and comfort in	RCT	Sample: 52	infants were observed in their nest for 15 min- utes with a heart rate device and the control group was observed for 15 minutes without any device. The heart rates and oxygen saturation were	tion, the SpO <sub>2</sub> value increased significantly (p<.003) while the PIPP score (p=.001) decreased in the ex-
Dora and TurkeyTo examine the effect of white noise and lullabies on pain per- vital signs of premature infants during painful inter- ventions.To examine the effect of white noise and lullabies on pain per- ception and vital signs of premature infants during painful inter- ventions.RCTSample: 66Heart rate, respiratory rate and oxygen satura- tion were measured and recorded before, during and after the interven- tion and post- procedure pain was evaluated us- ing the PIPP.Premature infants in the white noise group had the low- est mean PIPP score, mean heart rate and recorded before, during and after the interven- tion and post- procedure pain was evaluated us- ing the PIPP.Premature infants mean heart rate and recorded before, during and after the interven- tion and post- procedure pain was evaluated us- ing the PIPP.Iullaby group.Experimental group (White noise): 22Experimental group (lullaby): 22Iullaby group.		preterm.					
Dora and Tural Buyuk, 2021To examine the effect of white noise and lullabies on pain per- ception and vital signs of premature infants during painful inter- ventions.RCTSample: 66Heart rate, respiratory rate and oxygen satura- tion were measured and necorded before, during and after the interven- tion and post- procedure pain was evaluated us- ing the PIPP.Premature infants in the white noise group had the low- est mean NPP score, mean heart rate and respiratory rate, and highest mean oxy- gen saturation rate (p<.001). The white noise and lullabies played to premature infants during painful inter- ventions.RCTHeart rate, respiratory rate and oxygen satura- tion and post- procedure pain was evaluated us- ing the PIPP.Premature infants mean heart rate and respiratory rate, and highest mean oxy- gen saturation rate (p<.001). The white noise and lullabies played to premature infants during blood collection were ef- fective in reducing pain and the pain score was lower in the white noise group than in theExperimental group (lullaby): 22Experimental group (lullaby): 22Iullaby group.							
group (white noise): 22 Experimental group (lullaby): 22	and Tural Buyuk,	Turkey	the effect of white noise and lullabies on pain per- ception and vital signs of premature infants during painful inter-	RCT		rate and oxygen satura- tion were measured and recorded before, during and after the interven- tion and post- procedure pain was evaluated us-	in the white noise group had the low- est mean PIPP score, mean heart rate and respiratory rate, and highest mean oxy- gen saturation rate (p<.001). The white noise and lullabies played to premature infants during blood collection were ef- fective in reducing pain and the pain score was lower in the white noise group
group (lullaby): 22					group (white noise): 22		lullaby group.
					group (lullaby): 22		

						It had significant
Sarhangi et al., 2021	Iran	To examine the effect of the mother's heartbeat sound on physiological parameters and pain in- tensity after blood sam- pling in neo- nates in the intensive care unit.	RCT	Sample: 60	The experimental group listened to the mother's heartbeat sound until 10 minutes ago 10 min- utes after arterial blood samples were taken. The pain intensity was measured in three steps every 10 minutes using the NIPS 10 minutes be- fore, immediately after, and 10 minutes after the intervention.	moderate to major effects on oxygen sat- uration and respira- tory rate immediately after and 10 minutes after the interven- tion. Also, it had a major effect on heart rate immediately after the intervention (p<.05). Furthermore, the intervention had significant moderate to major effects on pain intensity im- mediately after blood sampling and 10 min- utes later (p<.05).
				group: 30		
		To examine		Control group: 30		
Sener		the effect of breast milk odor, white noise and		Sample: 86 Experimental group (breast milk odor): 22	Infants under mechani- cal ventilation were	White noise and fa- cilitated tucking were found to be more effective in relaxing
Taplak and Bayat,	Turkey	facilitated tucking on pain and	RCT	Experimental group (white noise): 21	given breast milk odor, white noise and facili- tated tucking and their	in pain reduction during EA procedure
2021		physiological parameters during the endotracheal		Experimental group [facilitated tucking]: 21	effect on pain was deter- mined.	
		aspiration (EA) procedure.		Control group: 22		(p>.05).
Baran- douzi et al., 2020	Iran	To investigate the analgesic effects of sucrose, mu- sic and their combination on pain when	RCT	Sample: 128	Two minutes before establishing vascular access,	Pain scores dur- ing vascular access procedure were significantly lower in the sucrose and com- bination groups com- pared to the control group, but not in the music group (p=.003, p<.001, respectively). Thirty seconds after the vascular ac- cess procedure, the pain score in three intervention groups (sucrose, music and combination groups p<.001, p=.009 and p<.001, respectively) was significantly
		opening vas- cular tracts			0.5 ml oral 24% sucrose was given to the sucrose	
		in preterm neonates.			and combination groups. The combination group listened to the same lul- laby as the music group.	lower than in the con- trol group.
			Double blind	Experimental group (sucrose) 33		
				Experimental group (music) 33		
				Experimental group		
				(sucrose-music): 31		
				Control group: 31		

Kahra- man et al., 2020	Turkey	To examine the effects of three auditory interventions (white noise, recorded ma- ternal voice and Mini- Muffs during heel lance) on pain and com- fort in prema- ture infants in neonatal intensive care units.	RCT	Sample: 64 Experimental group (white noise): 16 Experimental group (recorded mother's voice): 16 Experimental group (Mini- Muffs): 16 Control group: 16	During the heel lance, lancet was performed using a 19- gauge with incision depth of 1.1 mm in experimental groups. Sound interven- tions were performed five minutes before the intervention from a dis- tance of 30 cm.	White noise, recorded maternal voice and the mean oxygen saturation in the MiniMuffs group were higher than in the control group. The heart rate, cry- ing time, mean NIPS score and recorded maternal voice were significantly lower than the MiniMuffs groups control group (p<.001).
Tekgun- duz et al., 2019	Turkey	To examine the effect of oral glucose and lullaby to reduce pain in preterm in- fants support- ed with nasal continuous positive airway pressure	Random- ized Con- trolled trial Double blind	Sample: 106 Experimental group (lullaby): 35 Experimental group (glucose): 35 Control group: 37	The experimental groups were applied music and glucose during the inser- tion and removal of the tracheal tube. PIPP score was evaluated during and post-intervention.	Evaluation of the pain intensity of preterm infants post- inter- vention showed that preterm infants in the lullaby and glucose groups had reduced pain and those in the control group had more pain (p<.05).
Uemat- su and Sabue,	Japan	To examine the effect of music (Brahms lul- laby)	RCT	Sample: 25	The intervention in the experimental groups took	The mean PIPP (3.6 to 2.4) of infants dur- ing the intervention
2018		and nonnutri- tious			place at the beginning of the	was significantly lower than
			No blinding was	Experimental		
		sucking on heel lance in preterm infants.	performed.	group: 15	intervention and lasted until 5 minutes after it began.	points done every 30 seconds after heel
			Control group: 1000		lance (p=.0039 and p<.0001).	
Tang et al., 2018	China	To examine the applica- tion of a mu- sic interven- tion (MI) dur- ing insertion procedures of peripher- ally inserted	RCT	Sample: 60	The MI was performed for 10 minutes before and after the interven- tion from the music player set at a distance of 30 cm. The sound lev-	During PICC place- ment, there was a significant increase in blood oxygen satura- tion (p<.05) and a decrease in heart rate (p<.05) and cortisol accumulation in the intervention group receiving music com- pared to the control
		central cath- eter (PICC) in premature infants.		Experimental group: 30	el was set as 50-60 dB.	group. In addition, MI significantly reduced the pain score and the time required for
				Control group: 30		PICC placement in the MI group compared to the control group.

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Kuçuk Alemdar, 2018	Turkey	To examine the effect of maternal voice re- corded during the opening of the periph- eral vascular access pro- cedure, the breast milk odor and the incubator cover on pain and comfort, in preterm infants.	RCT	Sample: 136 Experimental group (mother's voice): 34 Experimental group (breast milk odor): 33 Experimental group (incubator	In the breast milk group, breast milk taken from the mothers was spilled into a sponge and put 5 cm away from the infants. The mother's voice was played to the babies at a level of 45dB during the intervention and until 15 minutes after the intervention. As an incubator cover, a special white cover was used. In the control group however, routine procedures were ap- plied.	Although there was no significant differ- ence between PIPP scores of the control and intervention groups before periph- eral vascular access procedure (p>0.05), a significant difference was found during and after the intervention (p<0.05).	
				cover): 35			
Kuçuk		To examine the effect of cover- ing the eyes and playing of intrauterine		Control group: 34 Sample: 94	While the intrauterine sound group was given sound at a level of 45dB, the closed eyes group's	It is reported that covering the eyes of	
Alemdar and Kardas Ozdemir, 2017	Alemdar and Kardas Ozdemir,	RCT	Experimental group (intrauter- ine): 32 Experimental group (closing eyes): 32 Control group: 30	eyes were closed 15 minutes before the intervention and kept closed for up to 15 min- utes after the interven- tion. No intervention was made to the control group.	preterm infants dur- ing vascular access positively affects their pain scores after intervention.		
Shah et al., 2017	Australia	To examine the trial of music, su- crose and combination therapy to re- lieve pain dur- ing heel lance in neonates.	RCT	Sample: 35	Every newborn went through all three in- terventions randomly during consecutive heel lance. A video camera in silent mode recorded facial expressions start- ing two minutes before to seven minutes after the heel lance. The vid- eos were then analyzed once per minute by two independent evaluators, who were blind to the	The PIPP revised scores were sig- nificantly lower at all time points after the combination therapy compared to the groups applied only music or sucrose. There was no differ- ence between the PIPP revised scores of the music and su-	
			Blind cross	Experimental	intervention, using the	crose groups.	
Qiu et al., 2017	China	To examine the effect of combined mu- sic and touch intervention (CMT) on pain response in	RCT	group: 35 Sample: 62	PIPP revised scale. Painful interventions in the infants hospitalized for two weeks were recorded. Of the 3707 painful interventions, 1913 were applied to the control group and 1794 were applied to the experimental group. There was no significant difference between these painful interventions.	CMT can signifi- cantly improve the concentration of β-endorphins and reduce the pain re- sponse of preterm	
		premature infants.		Experimental group: 30 Control group: 32	During these entries, the experimental group received combined music and touch intervention, while the control group received standard care.	neonates.	

Kuçuko- glu et al., 2016	Turkey	To examine the effect of white noise in relieving pro- cedural pain arising from vaccination in premature infants.	RCT	Sample: 75 Experimental group: 35 Control group: 40	Premature infants in the study group were ex- posed to white noise us- ing MP3 players placed next to their cradles one minute before vaccina- tion. The white noise continued until one min- ute after vaccination.	control group (PIPP ¼ 14.35±2.59) was significantly higher than that of the study group (PIPP ¼ 8.14±3.14) (p<.05).
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RCT studies. Table 4.1 shows detailed features of the studies and noise and facilitated tucking were found to be more effective at the detailed coding table that shows the study results (Table 5).

Of the studies obtained, 46.9% were conducted in Turkey, 17.7% in China, 11.8% in Iran, 5.9% in Taiwan, 5.9% in Lebanon, 5.9% in Japan and 5.9% in Australia. While 58.8% of the studies used In 14 studies included in our research, voice applications applied a single voice and a single experimental group, 41.2% used more than one voice and experimental group.

In total, there were 19 voice applications in 28 experimental groups. Of the sound interventions used, 26% were white noise, 16% were mother's voice, 21% were lullabies, 10.7% were heartbeats, 21% were music and 5.3% were intrauterine sounds. While the positive effect rate in voice interventions was found to be 88.2%, the studies that were negative or had no effect were determined as 11.8%.

### DISCUSSION

This study was conducted to reach scientific generalizations about the use of voice interventions in pain in premature infants in order to examine their effect on pain in premature infants by synthesizing the findings of RCT between 2002 and 2022. The results of 17 studies were included in the discussion of the study.

Considering publication years of the studies included in this research, most studies have been conducted in 2022, followed by the year 2021 and others (Table 5). The reason that there have been more studies recently can be the desire to support pain management in premature infants using non-pharmacological methods.

Three studies in voice interventions applied to the experimental groups among the included research showed no benefit in pain reduction [25-27]. Sharara-Chami et al. (2022) played music besides analgesic administration given to infants to be circumcised. In this research with 206 samples, there were 103 experimental groups [25]. The infants were subject to pain assessment of two blind and independent observers. An increase was found in heart rate and crying time of the experimental group. However, the played music did not affect pain. In another study by Ren et al. (2022), white noise was given during radial artery blood sampling [26]. From the sample group of 60 people, the experimental group of 29 people listened to white noise at 50 dB and the control group at 0 dB. There was no difference in pain levels between the two groups during and after the intervention. Therefore, white noise did not have any effect on pain. Sener Taplak and Bayat (2021) conducted a study examining the effect of the breast milk odor, white noise and facilitated tucking on pain and physiological parameters during EA in infants under mechanical ventilation [27]. The sample group was divided into 22, 21 and

relaxation before the EA intervention than the other groups. However, no difference was found between the groups in pain reduction.

to the experimental groups were found to be have positive effects on pain [4-8, 28-35]. Sound interventions applied to experimental groups in the studies that our research included simulation of the heartbeat, maternal heartbeat, maternal voice, white noise, lullaby, music and intrauterine sounds.

In the study conducted by Karadag et al. (2022), heartbeat simulation was applied to the experimental group. The PIPP score showed a significant result in the control group (p=0.001) [8]. In a study conducted by Sarhangi et al. (2021), the experimental group listened to their mother's heartbeat during blood collection. In pain assessment measured immediately after the intervention and 10 minutes later, there was a significant result compared to the control group (p<0.05). In their study, Kuçuk Alemdar and Kardas Ozdemir (2017) played intrauterine sounds to the experimental groups [31,34]. There was a significant difference after the NIPS pain assessment (p < 0.05).

Yu et al. (2022), Kahraman et al. (2020) and Kuçuk Alemdar (2018) played mother's voice to the experimental groups [28, 31, 34]. In these studies, there was a significant difference in the results of pain assessment in the experimental groups versus control groups (p<0.001, p<0.001 and p<0.05, respectively) [6].

Dora and Tural (2021), Kahraman et al. (2020) and Kuçukoglu et al. (2016) used white noise as sound intervention in their studies. They found that pain decreased after the intervention applied in the experimental groups (p<0.001, p<0.001 and p<0.05, respectively) [5-7].

Barandouzi et al. (2020), Tang et al. (2018), Uematsu and Sabue (2018), Qiu et al. (2017), and Shah et al. (2017) played music to the experimental groups [29,32,33,35-37]. The results from their studies showed a significant difference in pain scores of the experimental groups compared to the control groups (p=0.009, p<0.05, p<0.0001, p<0.05 and p<0.05, respectively).

Dora and Tural (2021), and Tekgunduz et al. (2019) played lullaby to the experimental groups as sound intervention. There was a significant difference in pain scores of the groups listening to lullaby compared to the groups that did not listen (p<0.001 and p<0.05, respectively) [7,30].

### CONCLUSION

We found 9301 studies related to our topic and published between 2002 and 2022 examining the effect of voice interventions on pain in premature infants. Upon elimination, a total of 17 studies 21, respectively and control group. Infants who underwent white using RCT method were included in our research. In three of

these studies, voice intervention did not give positive results SUGGESTIONS on pain, while in 14 of them gave positive results. In premature infants, voice applications had a positive effect on pain with • a rate of 82.3%. Besides pain assessment, positive results were obtained with the increase in SpO, value, stabilization in heart . rate, decrease in respiratory rate and shortening of crying times. However, meta-analysis studies should be conducted to clearly deem sound applications positive or negative.

Pain management is pointed as one of the independent tasks of a nurse. Therefore, the nurse should evaluate the severity of the The research received no external or grant funding. All authors infant's pain, and plan and implement pharmacological and contributed to the study conception and design. Material non-pharmacological pain management strategy developed with preparation, data collection and analysis were performed by G.M., the health team. Among non-pharmacological pain procedures, O.T. and G.M., O.T., and P.U. participated in conceiving the there are a limited number of evidence-based studies for voice design of the study, collecting the data and the data analysis The interventions (music, lullaby, white noise, mother voice, etc.). The first draft of the manuscript was written by O.T., and supervision results of the current study can be considered an evidence-based was made by G.M., and P.U. All authors commented on previous study in pediatric nursing and can contribute to its use in non- versions of the manuscript. All authors read and approved the pharmacological pain management.

- The frequency of studies with much evidence can be increased. Further double blind studies can be conducted.
- Sample size can be increased.
- Multi-centred studies can be carried out.

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final manuscript. The authors have declared that they have no competing or potential conflicts of interest.

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