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Targeted newborn hearing screening in the neonatal intensive care unit of Assiut University Hospital



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Abstract

Background: Early detection of congenital hearing loss (HL) and appropriate intervention is essential to minimize its dramatic impact. Target-based newborn hearing screening (TNHS) was initially targeted high-risk register (HRR) newborn. The advance techniques in neonatology in neonatal intensive care unit (NICU) lead to decrease mortality among low birth weight and preterm infants, but significant morbidities including hearing loss persist. In this study, we aimed to assess the prevalence of hearing loss and the percentage of different risk factors among the HRR neonates admitted to NICU at Assiut University Hospital.

Results: The prevalence of hearing loss among the study group is 1%. The most common risk factor present between the study group was prematurity associated with the low birth weight (57%). The combination of preterm and low birth weight has statistically significant effect on hearing loss (p < 0.006). There was statistically significant difference in the pass rate of the transient evoked otoacoustic emission (TEOAE) frequencies included in the screening program. The highest pass rate 91% and 85% was at 4 KH and 3 KH, respectively, while the least pass rate was at 1 KH (11.5%).

Conclusion: TEOAE screener is safe, noninvasive, and doesn't require sedation and cost-effective method in the newborn hearing screening at NICU. For HRR, it is better to do screening before hospital discharge so allowing enough time for comorbidities of the neonate to improve. Neonatal hearing screening in high-risk neonates in NICU allows early identification of children with hearing loss with early intervention.

Keywords: Newborn hearing screening, High-risk register, Neonatal intensive care unit

Background

In many world population, the prevalence of moderate and severe bilateral hearing loss (HL) (> 40 dB) is 2–3 per 1000 live births in well baby nursery population [1, 2] and 2–4 in 100 infants in an intensive care population [3–5] as reported by Wroblewska-Seniuk et. al. [6]. The most obvious effect of childhood hearing loss is on

language development, but it also has an impact on literacy, self-esteem, and social skills [7–9]. Early detection of congenital HL and appropriate intervention is essential to minimize its dramatic impact [10, 11]. The Joint Committee on Infant Hearing (JCIH) in 2007 recommended that screening should be completed by 1 month of age, and all infants with hearing loss should be identified by 3 months of age and receive intervention by 6 months of age [11].

The screening for hearing loss must fulfill all the criteria for universal screening. First of all, its prevalence is very high. Secondly, it has severe consequences for the

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affected child and its family if it is not diagnosed early and managed appropriately. Finally, the techniques used to diagnose this problem are relatively inexpensive, easily available, accurate, objective, and reliable [12]. There are two types of newborn hearing screening: (1) target-based newborn hearing screening (TNHS) was initially targeted toward those newborn "at risk" for hearing loss called high-risk register (HRR). (2) Universal newborn hearing screening (UNHS) is based on the principle that all neonates and infants should have access to objective screening to allow for early intervention [13].

In Egypt, UNHS has been implemented in 2019 by Egyptian Ministry of Health. The UNHS protocol in Egypt composed of two stages: first stage is conducted in primary healthcare centers during the days of the national program of thyroid screening using transient evoked otoacoustic emission (TEOAE). Newborns who failed the test rescreened 1 week later by TEOAE in the same centers. The second stage is conducted to diagnose all newborns who failed the second screening of first stage and conducted on audiology clinic in tertiary hospitals of ministry of health. In Assiut Governorate, the diagnostic stage is done on Assiut General Hospital. Before the implantation of the national newborn hearing screening program in 2019, there was no universal or targeted newborn hearing screening programs in Assiut. To our knowledge, there were no studies done to screen newborn hearing loss in Assiut University Hospital or even in Assiut Governorate.

There are many advances in neonatology that have improved outcomes for infants born premature and/or at low birth weight in neonatal intensive care unit (NICU) such as assisted ventilation and extracorporeal membrane oxygenation therapy. These advance techniques lead to decrease mortality among low-birth-weight infants, but significant morbidities including hearing loss persist.

The present study aimed to assess the prevalence of hearing loss and the percentage of different risk factors among the high-risk register neonates admitted to NICU at Assiut University Hospital.

Subjects and methods

Subjects

The study is a prospective cross-sectional study that was conducted on 200 neonates of both sexes who were admitted to NICU at Assiut University Hospital. The 200 neonates considered HRR according to the JCIH 2007 position statement [11]. The research was carried out between March 2020 and January 2021, and informed written consent was obtained from all parents/ guardians of all participants under 16 years old in the study. Ethical

approval was obtained by the ethical committee, Faculty of Medicine, Assiut University, Egypt.

Methods

Each subject was presented to the following:

- 1 Complete history taking include personal history (name, age, sex); prenatal history (infection, drugs, trauma, miscarriage, irradiation); natal history (gestational age and mode of delivery); postnatal history (birth weight, respiratory distress, cyanosis, jaundice, convulsions, cerebral hemorrhage, ototoxic medications intake and its duration, mechanical ventilation and its duration, exchange transfusion, and duration of NICU admission); family history (consanguinity, congenital anomalies, hearing impairment).
- 2 *Clinical examination* (for any congenital anomalies, external examination of auricles, otoscopic examination for patency and structure of external auditory canal).
- 3 Hearing screening by TEOAE: all neonates were screened by TEOAE, and the results were interpreted according to Rhode Island criteria in which pass result means signal-to-noise ratio ≥ 3dB at three distinct frequencies between 1 and 4 kHz) [14]. The first stage was conducted on 200 neonates, while the second stage was done for only nine neonates who failed the first stage.
- 4 Diagnostic ABR was performed for the neonates who failed the second stage screening, and for the neonates with hyperbilirubinemia, to not miss the auditory neuropathy, after hospital discharge and before age of 3 months at Audiovestibular Medicine Unit, Assiut University Hospital.

Results

The study included 200 neonates (n = 400 ears) of both sexes. They were incubated at NICU of Assiut University Hospital in the period between March 2020 and January 2021. Their demographic data is shown in Table 1.

Risk factors among the study group

Most of neonates of the study group had multiple risk factors at the same time like prematurity, low birth weight (LBW), and hyperbilirubinemia required blood transfusion (Table 2). The most common combined risk factors of hearing loss were prematurity plus LBW; they were presented together in 57% of the total neonates, while the most common single risk factor found alone in the study group was the prematurity (25%) followed by LBW (10.5%).

Results of first-stage screening

One-hundred ninety-one out of 200 neonates (95.5%) had passed the first-stage screening, and only nine

Table 1 Demographic data of the study group

	No. (<i>n</i> = 200)	%
Gender		
Male	118	59
Female	82	41
Age range group		
< 1 week	52	26
-1 < 2 weeks	75	37.5
-2 < 3 weeks	33	16.5
-3 < 4 weeks	13	6.5
-4 < 5 weeks	27	13.5
Range (min-max)	2–40	
Mean \pm SD	14.5 ± 9.39	
Gestational age		
< 37 weeks	123	61.5
≥ 37weeks	77	38.5
Range (min-max)	26-40	
Mean ± SD	34.8 ± 3.5	
Body weight level		
500 gm-1.7 kg	124	62
> 1.7 kg	76	38
Delivery		
Cesarean	120	60
Vaginal delivery	80	40

Table 2 Distribution of risk factors among the study group

Risk factor	No.	%
Preterm	50	25.0
Low BW	21	10.5
Hyperbilirubinemia with blood transfusion	5	2.5
Craniofacial anomalies	1	0.5
Brain anoxia	1	0.5
F.H of HL + preterm + low BW	1	0.5
Preterm + low BW	114	57.0
Preterm + hyperbilirubinemia with blood transfusion	1	0.5
Preterm + craniofacial anomalies	2	1.0
Preterm + low BW + craniofacial anomalies	1	0.5
Preterm + low BW + hyperbilirubinemia with blood transfusion	1	0.5
Preterm + low BW + craniofacial anomalies +brain anoxia	1	0.5
Brain anoxia + MV	1	0.5
Total	200	100.0

BW birth weight, FH family history, HL hearing loss, MV mechanical ventilation

neonates (4.5%) had failed the test. Distribution of the risk factors of the study group according to the result of the first-stage screening is shown in Fig. 1.

Tracking of the neonates who failed the first-stage screening

Four out of nine neonates who failed the first-stage screening succeeded to the second-stage screening, and the other five neonates dropped out after the first stage (4 of them died and one was missed). One of the four neonates passed the second-stage screening and advised to follow up. The other three neonates failed TEOAE and referred for the diagnosis by ABR (Fig. 2). One neonate had a normal hearing sensitivity, and the other two neonates had severe hearing loss. Based on the diagnostic ABR results, the prevalence of hearing loss in the present study is 1% (2 out of 200 cases).

Neonates with hyperbilirubinemia

The first-stage screening was carried on seven neonates that have hyperbilirubinemia and blood transfused to them. Five out of the seven passed the 1st stage screening and referred for the diagnosis by ABR, while two neonates did not pass the 1st stage and did not survive to the 2nd stage screening. Four neonates out of the five who passed the 1st stage screening had normal hearing sensitivity and one dropped out (her mother refused to conduct ABR) (Table 3).

Relationship between studied risk factors and HL

Logistic regression analysis for ABR according to risk factor was done (Table 4) to show the relationship between common risk factors in the study group and the hearing loss. There was statically significant effect of the combination of the risk factors of (prematurity + low birth weight) on hearing loss (p < 0.006); this means that the probability of the hearing loss is significant, when the two risk factors combined together in the neonate. Also, there was a statically significant effect of the risk factor of prematurity alone on the hearing loss (p < 0.037). Despite the statically significant effect of prematurity alone and (prematurity + low birth weight) on hearing loss, we could not generalize these results as two cases cannot be the cut point used to say highly significant relation between one of high-risk factors and HL.

Pass rate of TEOAE frequencies in the study group

Five frequencies were counted in during the 1st stage screening by TEOAE (1, 1.5, 2, 3, and 4 KH). Chi-square test showed no significant difference in pass rate of those frequencies between both ears, while there was a significant difference between the pass rate of frequencies in the same ear (p < 0.001); the highest pass rate was on 4 KH (91%) followed by 3 KH (85%), and least pass rate was on 1 KH (11.5%) (Fig. 3).

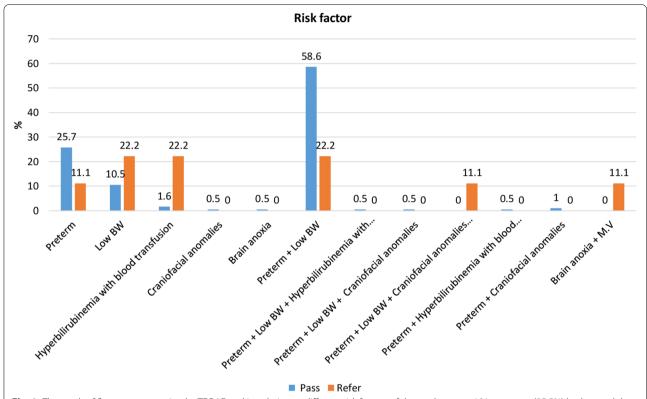


Fig. 1 The result of first-stage screening by TEOAE and its relation to different risk factors of the study group: 191 neonates (95.5%) had passed the 1st stage screening; most of the 112 neonates (58.6%) were preterm + low birth weight. Nine neonates (4.5%) failed the 1st stage screening, and their distributions were as the follows: one neonate was preterm, two neonates were low BW, two neonates had hyperbilirubinemia with blood transfusion, two neonates were preterm + low BW, one neonate was preterm + low BW + craniofacial anomalies + brain anoxia, and one neonate was with brain anoxia on mechanical ventilation (MV)

Discussion

The study was carried out in the form of targeted hearing screening program restricted to a study group of 200 neonates incubated in NICU in the time period between March 2020 and January 2021 and were subjected to risk factors for hearing loss. The most common risk factor in the study was prematurity; total premature neonates were 164 (82%). Prematurity alone as only risk factor was 25% while combined with low birth weight was 57% in 114 neonates. Our finding is in accordance with other literature in which prematurity was the most common risk factors among neonates in NICU [15]. On the other hand, our result did not match with other studies in which other risk factors were the major one like severe birth asphyxia and hyperbilirubinemia [16] and ototoxic medications [17]. The difference between the studies in the common risk factors may be due to the variability in environmental conditions surrounding the pregnancy or birth and the difference of medical care between the hospitals.

The probability of hearing loss increases with an increase in the number of risk factors [17]. In our study, there is

statically significant effect of the combination of preterm and low birth weight on hearing loss (p < 0.006). HL is a severe consequence of prematurity; its prevalence is inversely related to the maturity of the baby. Premature infants have many concomitant risk factors which influence the occurrence of hearing deficit [18]. On another hand, the risk factors for hearing loss in Ohl et al. study were neurological disorders, asphyxia, family history of hearing loss, and TORCH (toxoplasmosis, other agents, rubella, cytomegalovirus, herpes simplex) infection, but hearing loss was not associated with low birth weight or birth before 34 weeks of gestation which is different from our results [19].

In the present study, based on the diagnostic ABR results, the percentage of hearing loss is 1% (2 cases). The prevalence of hearing loss among neonates in NICU is variable in different studies. Some studies reported prevalence close to our results 1.6 % [20], 1.7 % [21], and 0.8 to 2.0 % [22], while others reported higher prevalence 5.09 % [23], 4,3 % [24], 4.55 % [19], and 4.91% [25]. The difference in the prevalence of hearing loss might be related to several factors like differences in screening and diagnostic methods of hearing loss [23]; HI may be affected by

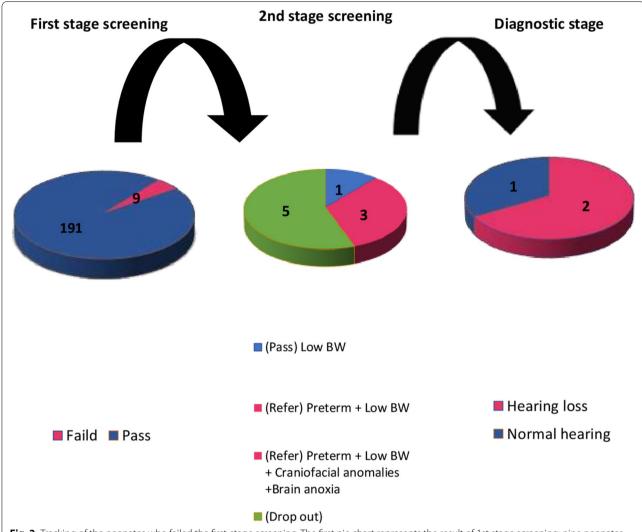


Fig. 2 Tracking of the neonates who failed the first-stage screening. The first pie chart represents the result of 1st stage screening; nine neonates (4.5%) failed the TEOAE test. The second pie chart represents the results of 2nd stage screening; five out of nine neonates who failed the 1st stage screening dropped out, one neonate passed TEOAE, and the other three neonates failed TEOAE and referred for the diagnosis by ABR. The third pie chart represents the results of the diagnostic stage; one neonate (preterm + low birth weight) had normal hearing, and the other two neonates (preterm + low BW; preterm + low BW + craniofacial anomalies + brain anoxia) had hearing loss

indirect medical factors as the availability and quality of healthcare provisions including birth clinics, NICU, and vaccination programs.

The risk factors for the development of ANSD have been studied in different literatures, and associations have been gathered. Some of the risk factors that have been identified are intracranial hemorrhage, asphyxia, hyperbilirubinemia, prematurity, low birth weight, neonatal ventilation, ototoxic drug exposure, dysmorphic features, Apgar scores, respiratory distress, cytomegalovirus infection, sepsis, meningitis, asphyxia, and family history of hearing loss [26–30]. In the current study, seven neonates (3.5%) had hyperbilirubinemia with levels

exceeded 20 mg/dl, and all had blood exchange within 24 h. The four out of seven who completed the diagnostic test had normal hearing.

Possible explanation that neonates with hyperbilirubinemia in the current study had no auditory neuropathy is the early blood transfusion. These neonates were subjected to blood transfusion early which helped to rapidly decrease the level of the bilirubin in the blood before the affection of the auditory nerve and decrease the possibility of auditory neuropathy spectrum disorder (ANSD) to occur. This agrees with a screening study conducted by Xu et al. (2019) on a group of neonates who met the standard of exchange transfusion

Table 3 Results of neonates with hyperbilirubinemia required blood transfusion

Distribution of risk factors	First TEOAE		Drop out (<i>n</i> = 3)	ABR normal hearing (n = 4)	
	Pass	Refer	No.	No.	
Hyperbilirubinemia with blood transfusion ($N = 5$)	3	2	2	3	
Preterm + hyperbilirubinemia with blood transfusion ($N = 1$)	1	0	0	1	
$\label{eq:preterm} \mbox{Preterm} + \mbox{low BW} + \mbox{hyperbilirubinemia with blood transfusion } (N = 1)$	1	0	1	0	

BW, birth weight

Table 4 logistic regression analysis for ABR results according to risk factors

Risk factor	EXP (B)	95 % CI f	95 % CI for EXP (B)		
		Lower	Upper		
Preterm	18.197	1.199	276.128	0.037*	
Low BW	2.708	0.258	28.449	0.407	
Hyper bilirubinemia with blood transfusion	0.129	0.006	2.681	0.186	
Preterm + low BW	16.097	2.215	116.959	0.006**	

EXP (B): this is the exponentiation of the B coefficient, which is an odds ratio. This value is given by default because odd ratios can be easier to interpret than the coefficient, which is in log-odds unit. Cl: confidence interval of the difference of a numerical field that is expected to contain the true value of a statistical parameter to be known for a population. BW, birth weight. *Statistically significant difference (p < 0.05). **Statistically significant difference (p < 0.01)

and another group of neonates in the same period who did not require exchange transfusion. They found lower incidence of auditory neuropathy in the exchange transfusion group than in the phototherapy group. They also noted that plasma exchange can rapidly reduce serum

UB level to 40–60% of the original level, and when it is combined with blue light phototherapy, the UB level can be decreased to 60–80% of original level 24 h later [31]. So, more attention should be paid on the effective role of blood transfusion in the prevention of auditory neuropathy in the future.

Another point to be mentioned in the current study is the pass rate of the TEOAE frequencies included in the screening program. There was a statistically significant difference between the pass rate of frequencies in the same ear (p < 0.001); the highest pass rate was on 4 KH (91%) followed by 3 KH (85%), and the least pass rate was on 1 KH (11.5%). This difference of the pass rate between the frequencies is reported in other screening programs in which higher frequencies (e.g., 2-4 or 2-5 kHz) had lower referral rates than lower frequencies (1-4 kHz) [32]. The stiffness of middle ear increases due to reducing the middle-ear air space by the presence of amniotic fluid and mesenchyme in the middle ear which occurs in early newborn life and therefore affects the transmission

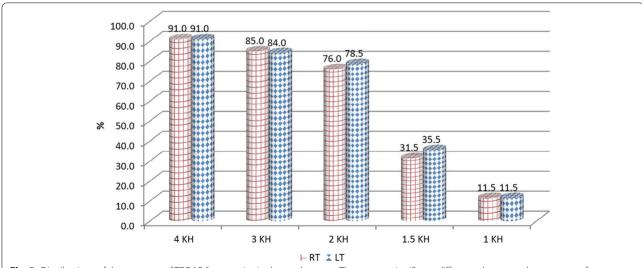


Fig. 3 Distributions of the pass rate of TEOAE frequencies in the study group. There was a significant difference between the pass rate of frequencies in the same ear (*p* < 0.001); the highest pass rate was on 4 KH (91%) followed by 3 KH (85%), and the least pass rate was on 1 KH (11.5%)

of lower-frequency sounds which can be associated with greater referral rates [33]. So, the use of higher frequencies TEOAE instead of lower frequencies will be valuable to ensure better result of newborn hearing screening.

Conclusions

In conclusion, we have screened 200 neonates incubated in NICU at Assiut University Hospital in the time period between March 2020 and January 2021. The prevalence of hearing loss is 1% among the study group. The most common risk factor present between the study group was prematurity associated with the low birth weight (57%). The combination of preterm and low birth weight has statistically significant effect on hearing loss (p < 0.006). There was statistically significant difference in the pass rate of the TEOAE frequencies included in the screening program. The highest pass rate 91% and 85% was at 4 KH and 3 KH, respectively, while the least pass rate was at 1 KH (11.5%).

Abbreviations

ABR: Auditory brainstem response; ANSD: Auditory neuropathy spectrum disorder; HL: Hearing loss; HRR: High-risk register; JCIH: The Joint Committee on Infant Hearing; LBW: Low birth weight; NICU: Neonatal intensive care unit; TEOAE: Transient evoked otoacoustic emission; TNHS: Targetbased newborn hearing screening; UNHS: Universal newborn hearing screening.

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Authors' contributions

KMO: patients' interview, gathering and analysis of the results, and writing the main manuscript; ESM, study design and analysis of the results and review the main manuscript; EAFS, study concept and design and analysis of the results and review of the manuscript. NHRAA, study design and patients' interview and review of the manuscript. MAAA: study concept and design, patients' interview, gathering and analysis of the data, and drafting and critical review of manuscript. The author(s) read and approved the final manuscript.

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Availability of data and materials

Data generated or analyzed during this study are included in this published article and available from the corresponding author on reasonable request

Declarations

Ethics approval and consent to participate

Informed written consent was obtained from all parents/ guardians of all participants under 16 years old in the study. Ethical approval was obtained by the ethical committee, Faculty of Medicine, Assiut University, Egypt. Approval number (17100662).

Consent for publication

Not applicable

Competing interests

The authors declare that they have no competing interests.

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