Significance of cardiac murmurs in term neonates admitted to a neonatal intensive care unit of a tertiary level hospital in India

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Abstract

Introduction: There is a paucity in the literature of prospective studies on neonatal heart murmurs and their correlation with the diagnostic yield on 2D echocardiography.

Objectives: To assess the prevalence of cardiac murmurs in neonates and their significance using 2D Echocardiography/Colour Doppler.

Method: A prospective descriptive analytical study was carried out on 100 term neonates admitted to the neonatal intensive care unit (NICU) of a tertiary level hospital from April - November 2017 with audible systolic (grade 2/6 or above), diastolic (2/4 or above) or continuous murmurs. These patients were evaluated with a detailed 2D Echocardiogram/Colour Doppler on portable Sonosite / Epiq using an 8-12 Hz probe. Patient with isolated small patent foramen ovale (<5mm) and those admitted to the NICU with diagnosed cardiac disease were excluded from study. A 100 term neonates without murmur were used as controls. The results obtained were statistically analysed using SPSS software. p < 0.05 was taken as significant Institutional Ethics Committee approval was obtained prior to study.

Results: Of the 100 neonates with murmur, 65 (65%) had significant structural defects and 35 (35%) had no structural heart defects. The mean hospital stay was 13.03 days. Seventy five (75%) neonates had good outcome and 25 (25%) had poor outcome in the form of prolonged NICU stay and mortality. Of the 100 neonates without murmur, 97 (97%) had good outcome and 3 (3%) had poor outcome due to non-cardiac issues.

Conclusions: The prevalence of murmurs among neonates was 6.2 per 1,000 live births in inborn deliveries (excluding previously diagnosed cases). Complex structural abnormalities were found in 28% of neonates with murmurs. The rest had isolated lesions or single structural lesions such as VSD (19%), PDA (18%) or ASD (2%).

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(Key words: Congenital heart disease, cardiac murmur, neonate)

Introduction

Congenital heart disease (CHD) has an incidence of around 0.8-1 per 1000 live births globally1,2. Early detection will improve management and outcome3. Ultrasound scanning during pregnancy and postnatal clinical examination have a low rate of CHD detection4,5,6. Routine clinical examination diagnoses less than 50% of the children with CHD5. Electrocardiography (ECG) and chest x-ray contribute little to diagnosis of heart murmurs7,8,9. Murmur is the commonest reason for cardiac consultation10. Echocardiography should complement clinical diagnosis if easily available, to avoid missing life threatening cardiac lesions11,12. Paucity in the literature of prospective studies on neonatal heart murmurs and their correlation with diagnostic yield on echocardiography, prompted us to undertake this study.

Objectives

To assess the prevalence of cardiac murmurs and their significance in babies admitted to a neonatal intensive care unit (NICU) compared to a control group of term neonates without cardiac murmurs using 2D Echocardiography/ Colour Doppler.

Method

A prospective descriptive analytical study was carried out on 100 neonates with varied diagnoses admitted to the 150 bedded neonatal intensive care unit (NICU) of a tertiary level hospital. Institutional Ethics Committee approval was obtained prior to the study. NICU setting has both intramural and extramural services. All full term neonates in whom cardiac murmurs were detected on routine examination were included in the study. Babies with audible murmurs were assessed with a detailed 2D Echocardiogram/Colour Doppler on portable Sonosite / Epiq using an 8-12 Hz probe. Neonates included in study were those with audible, systolic (grade 2/6 or above) and diastolic
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(grade 2/4 or above) murmurs heard by fellow or consultant and confirmed by cardiologist or continuous murmurs detected by NICU. Study period was 8 months from April - November 2017. Full term (>37 weeks) neonates with murmurs were included and infants with previously documented or suspected CHD on fetal echocardiogram, Patient with isolated small patent foramen ovale (<5mm) and those admitted in NICU with diagnosed cardiac disease were excluded from the study. One hundred full term neonates without murmurs were taken as controls. Murmur, for study purposes was defined as audible systolic murmur (≥grade 2/6), diastolic (≥grade 2/4) and continuous murmur.

Case record form for the study was designed and included age at presentation, gender, address, family history, antenatal history, gestational age, birth weight, heart rate (HR), respiratory rate (RR), saturation (SPO2), clinical examination details including findings on general examination, cardiac examination, comorbid findings, cyanosis, chest x-ray (if available) and 2 D echocardiogram findings. Based on echocardiographic data neonates were divided into those with innocent and pathological murmurs. Number of days of hospitalisation and outcome were noted. The course of the neonate in hospital and the diagnosis on discharge were also noted. If the child needed intervention or surgery for cardiac condition it was recorded. The outcomes were classified as favourable/good in the form of early recovery and brief NICU stay from the cardiac point of view and not favourable/poor in the form of prolonged NICU stay or mortality.

Statistical analysis: The results obtained were statistically analysed by Chi square test and graphically displayed. *p*<0.05 was taken as significant. All statistical analyses were performed using SPSS software version 23.

The guideline for cardiac murmur identified at newborn examination is shown in Figure 1.

![Figure 1: Guideline for cardiac murmur identified at newborn examination](image-url)
Results

There were 3903 live births recorded during the eight month period. Of them, 100 neonates had audible murmurs. Of the 100 neonates, 66% presented between 0 to 10 days of life, 22% between 11 to 20 days of life, and 12% between 21 to 28 days of life. The mean age of presentation of murmur in our study was 8.4 days. Among the inborn babies, the prevalence of murmurs, in babies previously not known to have congenital heart disease, was 6.2 per 1,000 live births. Incidence of murmurs was 39.2 per 1,000 live births in the outborn, identified through precordial auscultation. The outborn babies were excluded from prevalence calculation as birth prevalence could not be estimated. The male: female ratio of the 100 neonates with murmur was 1.22:1. The mean maternal age was 24.5 years, ranging from 18 to 42 years. The mean heart rate was 133 beats per minute.

On clinical examination, 68% neonates had ejection systolic murmur, 3% had continuous murmur and 29% had pansystolic murmur. The intensity of the murmur was not related to the presence or absence of CHD. The mean percentage oxygen saturation was 96.7%, 83% having saturations between 95-100%, 6% between 90-94%, 8% between 85-89% and 3% having saturations less than 85%. Neonates with cyanosis were 13% and neonates without cyanosis were 87%. Among the 100 neonates with murmur, the common lesions found on echocardiography were ventricular septal defect (19%), patent ductus arteriosus (18%), atrial septal defect (2%) and cyanotic heart with multiple structural lesions (28%). Anatomic lesions diagnosed on 2D echocardiography/Colour Doppler in neonates with murmur are shown in Table 1.

<table>
<thead>
<tr>
<th>Type of lesion</th>
<th>No. of neonates</th>
</tr>
</thead>
<tbody>
<tr>
<td>Peripheral branch pulmonary artery stenosis</td>
<td>35</td>
</tr>
<tr>
<td>Ventricular septal defect (VSD)</td>
<td>19</td>
</tr>
<tr>
<td>Patent ductus arteriosus (PDA)</td>
<td>18</td>
</tr>
<tr>
<td>Transposition of the great arteries (TGA) + VSD + PDA</td>
<td>4</td>
</tr>
<tr>
<td>VSD + atrial septal defect (ASD)</td>
<td>2</td>
</tr>
<tr>
<td>Double outlet right ventricle (DORV) + single ventricle (SV) + large ASD</td>
<td>2</td>
</tr>
<tr>
<td>DORV + VSD + pulmonary stenosis (PS)</td>
<td>2</td>
</tr>
<tr>
<td>Hypoplastic left heart syndrome (HLHS)</td>
<td>2</td>
</tr>
<tr>
<td>Coarctation of aorta (COA) + ASD</td>
<td>2</td>
</tr>
<tr>
<td>Tetralogy of Fallot (TOF) + pulmonary atresia + large PDA</td>
<td>2</td>
</tr>
<tr>
<td>Ostium secundum ASD + PDA</td>
<td>1</td>
</tr>
<tr>
<td>VSD + COA</td>
<td>1</td>
</tr>
<tr>
<td>TGA + intact ventricular system (IVS) + PDA</td>
<td>1</td>
</tr>
<tr>
<td>Ebstein anomaly</td>
<td>1</td>
</tr>
<tr>
<td>Dysplastic tricuspid valve (TV) + Severe tricuspid regurgitation (TR)</td>
<td>1</td>
</tr>
<tr>
<td>Dysplastic TV + tiny PDA</td>
<td>1</td>
</tr>
<tr>
<td>Coronary sinus (CS) Total anomalous pulmonary venous connection (TAPVC) + small VSD</td>
<td>1</td>
</tr>
<tr>
<td>CS TAPVC + small ASD</td>
<td>1</td>
</tr>
<tr>
<td>Tricuspid atresia (TA) + large PDA</td>
<td>1</td>
</tr>
<tr>
<td>Pulmonary atresia + IVS + PDA</td>
<td>1</td>
</tr>
<tr>
<td>Partial anomalous pulmonary venous connection (PAPVC) + VSD + PDA</td>
<td>1</td>
</tr>
<tr>
<td>Complex = Complete atroventricular canal defect (CAVCD) + DORV + TGA + TAPVC.</td>
<td>1</td>
</tr>
<tr>
<td>Total</td>
<td>100</td>
</tr>
</tbody>
</table>

Table 1: Anatomic lesions diagnosed on 2D echocardiography/Colour Doppler in neonates with murmur

Anatomic lesions diagnosed on 2D echocardiography/Colour Doppler in neonates without murmur are shown in Table 2.

Figure 2 shows the classification of murmurs after 2D echocardiography.

<table>
<thead>
<tr>
<th>Type of lesion</th>
<th>Number</th>
</tr>
</thead>
<tbody>
<tr>
<td>Normal</td>
<td>71</td>
</tr>
<tr>
<td>Patent ductus arteriosus (PDA)</td>
<td>10</td>
</tr>
<tr>
<td>Patent foramen ovale (PFO)</td>
<td>13</td>
</tr>
<tr>
<td>PFO + PDA</td>
<td>6</td>
</tr>
</tbody>
</table>

Table 2: Anatomic lesions diagnosed on 2D echocardiography/Colour Doppler in neonates without murmur (n=100)
Of the 100 neonates, 65% had significant structural defects (Figure 3). Of the 65%, 11% were duct dependent lesions needing change of routine care. Ventricular septal defect (VSD) was diagnosed in 19%. Most VSDs were small and in the perimembranous and muscular portion of the interventricular septum.

Of the 100 neonates with murmur, 48% were hospitalised between day 1 to day 10, 37% between day 11 to day 20, 14% between day 21 to day 30 and 01% was hospitalised more than 30 days (Figure 4). The mean hospital stay was 13.03 days.

Of the 100 neonates without murmur, 81% were hospitalised between day 1 to day 10, 15% between day 11 to day 20, 04% between day 21 to day 30 and none were hospitalised more than 30 days. Of the 100 neonates with murmur, 75% had good outcome and 25% had poor outcome in the form prolonged NICU stay and mortality (Table 3).

Table 3: Outcome

<table>
<thead>
<tr>
<th>Outcome</th>
<th>Neonates with murmur Number (%)</th>
<th>Neonates without murmur Number (%)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Good</td>
<td>75 (75)</td>
<td>97 (97)</td>
<td>0.000007</td>
</tr>
<tr>
<td>Bad</td>
<td>25 (25)</td>
<td>03 (03)</td>
<td></td>
</tr>
<tr>
<td>Total</td>
<td>100 (100)</td>
<td>100 (100)</td>
<td></td>
</tr>
</tbody>
</table>

Comorbid conditions noted in neonates with murmur were anorectal malformation, sepsis, arterio-venous (AV) malformation, congenital diaphragmatic hernia and tracheoesophageal fistula. Of the 100 neonates without murmur, 97 (97%) had good outcome and 3 (3%) had bad outcome due to non-cardiac issues. P value is significant (p =0.000007). Simple congenital heart diseases like VSD, atrial septal defect (ASD) and patent ductus arteriosus (PDA) were managed on medical follow up. Complex congenital heart disease required interventions in the form of palliative or corrective surgery.

Discussion

CHD is still an important cause of morbidity and mortality\textsuperscript{13}. Neonatal heart murmur is the commonest cause of cardiologist consultation in neonatal intensive care units and nurseries\textsuperscript{14}. Prevalence of heart murmur in neonates varies from 6-770 per 1000 live births\textsuperscript{11,15}. In present study the prevalence of murmurs was 6.2 per 1,000 live births in inborn. This prevalence is low as antenatally diagnosed cases were not included. The outborn babies were excluded from prevalence calculation as birth prevalence could not be estimated.
On clinical examination of 100 neonates with murmur, 68% had ejection systolic murmurs, 3% had continuous murmurs and 29% had pansystolic murmurs. The intensity of the murmur was not related to the presence or absence of CHD. The mean percentage oxygen saturation was 96.7%. Whilst 13% neonates had cyanosis, 87% were without cyanosis. Thus, in this study, acyanotic heart disease was more prevalent than cyanotic heart disease. A study by Tanner et al found benign murmurs in 48.4% and structural cardiac defects in 51.6% neonates. If a heart murmur is audible, the probability of an underlying cardiac anomaly is 65%. In our study 65% of neonates with heart murmur had underlying structural cardiac malformations, 19% being VSDs alone and 11% being duct dependent lesions needing change of routine care. Most VSDs were small and in the perimembranous and muscular portion of the interventricular septum. Studies using echo-Doppler cardiography have demonstrated a high prevalence of muscular VSDs in neonates.

Clinical examination of the neonate can detect only 44-45% of cardiac anomalies presenting in infants. Our study was limited to live born neonates and focused on the clinical significance of precordial murmurs heard during the routine neonatal examination. Among the 100 neonates with murmur, the common lesions found on echocardiography were VSD (19%), PDA (18%), ASD (2%) and cyanotic heart with multiple structural lesions (28%).

With respect to outcome, 75% of neonates had good outcome and 25% had poor outcome in the form prolonged NICU stay more than 30 days and mortality. Comorbid conditions noted in neonates with murmur were anorectal malformation, sepsis, AV malformation, congenital diaphragmatic hernia and tracheoesophageal fistula which added to the mortality and prolonged morbidity. Neonates with duct dependent circulation needed urgent attention and were referred for intervention or surgery depending upon the clinical scenario, indicating that early detection and diagnosis helps to offer better care and timely interventions. Hence innocent murmurs are only innocent after echocardiographic diagnosis. Most of these patients with neonatal murmurs were asymptomatic and majority of these defects were small and closed spontaneously.

Conclusions
The prevalence of murmurs among neonates was 6.2 per 1,000 live births in inborn deliveries (excluding previously diagnosed cases). Complex structural abnormalities were found in 28% of neonates with murmurs. The rest had isolated lesions or single structural lesion and the most common single lesions were VSD (19%), PDA (18%), and ASD (2%).

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