INFECTIVE ENDOCARDITIS IN CHILDREN

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ABSTRACT

A total of 14 cases of infective endocarditis (IE) in children aged 6 months to 10 years were seen from December 1987 to December 1992 at the pediatric unit of Ayatollah Taleghani Medical Center. The majority of patients (12 of 14) were between 5 and 10 years of age. Acyanotic congenital heart disease was known to preexist in 78.6% and rheumatic valvular heart disease in 21.4% of cases. Organisms were identified by blood culture in 50%, the majority being Streptococcus viridans (36% of cases). Vegetations were detected by echocardiography in 64%. The overall death rate was 21.4%. Two patients with aortic stenosis developed IE after cardiac surgery for repair of stenotic valves and both of them died. One patient with rheumatic aortic and mitral valve insufficiency underwent valve replacement due to intractable heart failure.

In conclusion, the frequent presence of underlying heart disease, the rarity of IE in infancy, and the ominous prognosis of postoperative endocarditis and aortic valve involvement are emphasized.

Key Words: Children, Infective endocarditis (IE), Congenital heart disease (CHD), Valvular heart disease.

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INTRODUCTION

Infective endocarditis (IE) is a severe infection which is becoming more frequently recognized in children. The reported incidence of IE in infancy and childhood is one in 4500 hospital admissions.14 Most studies of IE in children have been reported from industrialized countries,6,10,11,12,15 whereas only a few have been reported from nonindustrialized countries.1,7 In this study, I will present my experience with IE in a group of Iranian children who were admitted to a pediatric department over a 5-year period.

MATERIALS AND METHODS

Charts were reviewed on 14 patients discharged from the pediatric ward of the Ayatollah Taleghani Medical Center from 1987 to 1992 with a diagnosis of infective endocarditis. The entire hospital course of each patient was reviewed. The following data was acquired from the patients' medical record: basic identifying information, dates of hospitalization, follow-up status, antecedent diseases, predisposing factors, symptoms, signs, laboratory data, treatment, course, and complications. Patients with previously known congenital or rheumatic heart disease (RHD) were accepted into the study if they fulfilled at least two of the following criteria:

1- Vegetations verified by echocardiography
2- Fever without any extracardiac foci
3- Bacteremia without an extracardiac focus
4- Symptoms compatible with embolism

Standard methods were employed for obtaining and processing blood cultures, as well as for the identification and isolation of microorganisms.
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Table I. Significant clinical and laboratory features of 14 children with IE and their treatment, complications and outcomes

<table>
<thead>
<tr>
<th>Case</th>
<th>Age</th>
<th>Sex</th>
<th>Clinical and laboratory findings at time of diagnosis</th>
<th>Underlying heart disease</th>
<th>Possible source</th>
<th>Blood culture</th>
<th>Echocardiography</th>
<th>Treatment</th>
<th>Complications</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>9 years</td>
<td>F</td>
<td>Fever, heart murmur, T ESR, leukocytosis</td>
<td>PDA unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>2</td>
<td>9 years</td>
<td>F</td>
<td>Fever, heart murmur, CHF, spina bifida, anemia, T ESR, leukocytosis</td>
<td>PDA unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>3</td>
<td>9 years</td>
<td>M</td>
<td>Fever, heart murmur, anemia, spina bifida, petechiae, T ESR, leukocytosis</td>
<td>VSD unknown</td>
<td>Klebsiella</td>
<td>RV vegetation</td>
<td>Coagulase negative staphylococci</td>
<td>Chloramphenicol</td>
<td>None</td>
<td>Survived</td>
</tr>
<tr>
<td>4</td>
<td>2 years</td>
<td>M</td>
<td>Fever, heart murmur, CHF, anemia, T ESR</td>
<td>PDA unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>5</td>
<td>7 years</td>
<td>F</td>
<td>Fever, heart murmur, CHF, anemia, T ESR</td>
<td>RHD MR unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>6</td>
<td>9 years</td>
<td>F</td>
<td>Fever, heart murmur, T ESR, leukocytosis</td>
<td>VSD unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>7</td>
<td>6 years</td>
<td>F</td>
<td>Fever, heart murmur, CHF, T ESR</td>
<td>VSD+VSD unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>8</td>
<td>5 years</td>
<td>M</td>
<td>Fever, heart murmur, spina bifida, T ESR, leukocytosis</td>
<td>AS unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Unknown</td>
</tr>
<tr>
<td>9</td>
<td>9 months</td>
<td>M</td>
<td>Fever, heart murmur, CHF, spina bifida, anemia, T ESR, leukocytosis</td>
<td>VSD unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Unknown</td>
</tr>
<tr>
<td>10</td>
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<td>F</td>
<td>Fever, heart murmur, T ESR</td>
<td>PDA unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Penicillin</td>
<td>Severe</td>
<td>Survived</td>
</tr>
<tr>
<td>11</td>
<td>9 years</td>
<td>F</td>
<td>Fever, heart murmur, CHF, anemia, hemoptysis, T ESR</td>
<td>AS unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Died</td>
</tr>
<tr>
<td>12</td>
<td>7 years</td>
<td>M</td>
<td>Fever, heart murmur, anemia, T ESR, leukocytosis</td>
<td>AS unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Died</td>
</tr>
<tr>
<td>13</td>
<td>8 years</td>
<td>M</td>
<td>Fever, heart murmur, CHF, RHD MR+RI unknown</td>
<td>VSD unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Died</td>
</tr>
<tr>
<td>14</td>
<td>9 years</td>
<td>M</td>
<td>Fever, heart murmur, CHF, anemia, petechiae, T ESR, leukocytosis</td>
<td>RHD MR unknown</td>
<td>Streptococci</td>
<td>Coagulase negative staphylococci</td>
<td>no vegetation</td>
<td>Vancomycin</td>
<td>Severe</td>
<td>Died</td>
</tr>
</tbody>
</table>

M = male; F = female; CHF = congestive heart failure; T = increased; PDA = patent ductus arteriosus; VSD = ventricular septal defect; RHD = rheumatic heart disease; MR = mitral regurgitation; COA = coartation of aorta; AS = aortic stenosis; AR = aortic regurgitation; PA = pulmonary artery; RV = right ventricle; MV = mitral valve; AV = aortic valve; P = penicillin; S = streptomycin; C = ceftriaxone; G = gentamicin; V = vancomycin; DRC = disseminated intravascular coagulation.

Predisposing events, including dental procedures, were accepted as portal of entry if they occurred within 4 weeks prior to the first symptoms of IE. Recent cardiac surgery (less than 2 months before onset of symptoms) was accepted as a predisposing event. Recent antibiotic treatment was defined as antibiotic therapy within two weeks prior to admission.

Parenteral antibiotics were started in all patients after taking multiple blood cultures. In patients with a good general condition, antibiotic therapy was usually initiated with a combination of aqueous penicillin G and an aminoglycoside. In patients with critical general conditions, vancomycin plus gentamicin was administered. All patients were treated intravenously for a minimum period of 4 weeks. Autopsy was not possible in any of the patients who died, mainly because parents refused permission and wanted to bury the dead immediately according to traditional and religious rites.

RESULTS

In this study, which was performed in a university hospital, all pediatric cases with a discharge diagnosis of IE were reviewed. Their significant clinical and laboratory features are summarized in Table I. The mean age of the patients was 7 years with a range of 6 months to 10 years. 86% (12 of 14) were 5 to 10 years old. Of 14 children, 8 (57%) were boys and 6 (43%) were girls (M/F = 1.3:1). Fever of variable duration and grade, loss of appetite and weight loss were seen in all patients. Cardiac murmurs were auscultated in all patients and congestive heart failure (CHF) was present in 9 (64%) cases.

Congenital heart disease (CHD) was the predominant lesion, occurring in 11 (78.6%) children. 4 (28.6%) patients had ventricular septal defect (VSD) and in one of them it was accompanied by coarctation of the aorta. 4 had patent ductus arteriosus (PDA) and the other three (21.4%) suffered from aortic stenosis. In 3 patients (21.4%) rheumatic valvular heart disease was diagnosed. In all of them mitral valve incompetence was present and in a 6 year old boy it was accompanied by aortic valve insufficiency. All of the above anomalies were acyanotic.

Splenomegaly was detected in 4 (28.6%) patients. Peripheral mucocutaneous manifestations such as Osler's nodes, Janeway lesions and splinter hemorrhage were not seen in any patient, but two (14%) had subconjunctival petechiae. Anemia (defined as a hematocrit concentration of less than 10g/dL) was present in 9 (64%) cases. An elevated erythrocyte sedimentation-
tion rate (defined as an ESR greater than 35 mm in the first hour by the Westergren method) was present in all patients and in the majority of them it was greater than 50 mm in the first hour. A mild leukocytosis (defined as a leukocyte count greater than 13000/mm$^3$) was demonstrated in 7 patients (50%). Microscopic hematuria was seen in one patient. Blood cultures were positive in 7 patients (50%) and negative in the rest. Of 7 culture-positive cases, *Streptococcus viridans* was found in 5, *Staphylococcus aureus* in 1, and *Klebsiella* in 1 case.

Echocardiography (M-mode and two-dimensional) was performed in all patients and vegetations were detected in 9 (64%) of them. In one patient (no.13) with RHD, surgery was required because of intractable heart failure. His aortic valve was repaired and mitral valve replaced. One patient (no.9) had been discharged by his parents before completing his course of therapy and in spite of physician’s advice. Three patients (21.4%) died, two of which were boys with aortic stenosis who developed IE after cardiac surgery and repair of their stenotic valves. One of these two children (case no. 11) died because of a sudden embolic cerebrovascular accident and the other one died due to renal impairment manifested by oliguria and an elevated serum creatinine level. The third patient died in septic shock.

### DISCUSSION

The present study is an analysis of patients with IE diagnosed over a period of 5 years at our center. In this study, 12 of 14 patients were between 5 and 10 years of age. This is in accordance with the observation of IE in older children.$^5,10$

In Western reports, positive blood cultures were high and varied from 80% to 92% in different series.$^2,4,14$ In two reviews from developing countries,$^1,7$ positive blood cultures were reported in 47.8% and 58% respectively. In our study bacterial isolation was achieved in 7 (50%) patients. We ascribe the high proportion of negative cultures to previous antibiotic therapy, fastidious organisms with difficult culture requirements, very low numbers of circulating organisms, and less than meticulous culture techniques. Suppression of blood culture positivity by outpatient antibiotic therapy in patients with bacterial endocarditis has been reported by Pazin et al.$^{13}$ Patients who have received antibiotics should have more cultures taken initially, and all patients suspected of having IE should have additional blood cultures drawn if the first cultures fail to grow an organism and symptoms persist.$^2$

The introduction of echocardiography has made it possible to demonstrate vegetations in 75-80% of cases of IE.$^9$ Hence one would expect that the diagnosis of IE could be established earlier. There is, however, no doubt that in the future the transesophageal echocardiographic approach will yield a better access in clinically suspected cases with normal transthoracic echocardiographic studies.$^{16}$ In the present study, two-dimensional echocardiography revealed vegetations in 9 of 14 (64%) cases.

Similar to several previous reports,$^{11,14}$ in the present review no dental procedures were performed prior to IE. Poor dental hygiene is possibly of greater importance than the dental procedures themselves as a factor of susceptibility to IE.

A special note concerning "high-risk" individuals is appropriate. In the pediatric age group, these so-called "high-risk" individuals include those with prosthetic heart valves, those with aortic valve stenosis or insufficiency, and those patients with surgically constructed systemic to pulmonary arterial shunts.$^8$ In a study by Bahl et al.$^1$ including 23 patients with right heart endocarditis, five of six patients with postoperative endocarditis died, signifying an ominous prognosis for
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this subgroup when treated by medical means only. In this report, aortic valve involvement was associated with a high rate of complications and mortality. This group of patients may benefit from early surgical intervention.

REFERENCES

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