An Observational Study on Infective Endocarditis: A Single Center Experience

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1. Background

IE (Infective Endocarditis) is a life-threatening infectious disorder with a relatively high mortality rate if misdiagnosed. There are nonspecific and different symptoms in IE diagnosis and in most cases diagnosis is based on fever, vegetation in echocardiography and positive blood culture in some cases (1). In absence of these symptoms, we cannot exclude IE from differential diagnoses of suspected patients. Due to this fact, some diagnostic criteria such as traditional and modified Duke criteria were proposed for IE diagnosis (2-5). The incidence of IE had been raised in the recent years. Epidemiological studies showed specific features for each country (13). Diagnosis of IE is commonly based on echocardiography, which is the procedure of choice for detection of vegetations (2, 14).

2. Objectives

The present study was designed to evaluate clinical and laboratory presentations of IE patients and comparison between their TTE and TEE findings in Baqiyatallah hospital between 2007 and 2010.

3. Patients and Method

This cross-sectional study was performed between February 2007 and March 2010. We included all patients with endocarditis who were hospitalized in our center. Totally, thirty five patients were included. We confirmed endocarditis diagnosis in these patients according to different including elderly patients with valvular diseases, hemodialysis patients and IV drug abusers. Epidemiological studies showed specific features for each country (13). Diagnosis of IE is commonly based on echocardiography, which is the procedure of choice for detection of vegetations (2, 14).
Duke criteria (2, 15). The definite diagnosis requires positive blood cultures or positive cultures (or histology or PCR of the organism) from vegetation or embolus. TTE and TEE were performed to confirm the diagnosis. Echo features suggestive of IE are moving mass on atrial side of atroventricular valves or LV side of ventriculoarterial, abscess, new paravalvular or valvular leakage and dehiscence of prosthetic valves. These masses show high frequency movements and could be bulky, linear, round or irregular. In data checklist, age, sex, background cardiac disorders and prosthetic heart valves were recorded in the demographic part. Laboratory tests such as Erythrocyte Sedimentation Rate (ESR), C-reactive Protein (CRP), Complete Blood Cell (CBC) and Blood Culture (in three times) were performed. Dyspnea, fever and cardiac murmur, results of laboratory parameters, blood culture, pathologic, antibiotic resistance, vegetation, pericardial effusion, TTE and TEE, treatment type and results of follow-up were recorded in a check list. After follow-up period for about 13 months, outcome of patients were assessed according to improvement in clinical symptoms and laboratory parameters. To eliminate experimenter bias, all echocardiographies were performed by a same echocardiographer.

3.1. Statistical Analysis
Data was entered SPSS 16.0 for Windows (SPSS Inc., Chicago, Illinois, USA). Student’s t-test was used for sub-group analysis. P < 0.05 were assumed to be significant.

4. Results
In the present study, 16 (45%) male and 19 (55%) female patients were included. Their mean age was 56.36 ± 12.44 (35-78) years. Seven patients had a history of mitral valve replacement (MVR), two a history of aortic valve replacement (AVR) and two AVR plus MVR. Seven patients had chronic renal failure, two had diabetes, two hypertension and two kidney transplantation. None of included patients had a history of intravenous drug abuse.

4.1. Clinical Symptoms and Laboratory Findings
Fever was the most common symptoms among study patients. Cardiac murmur, chest pain and pulmonary crackle had highest frequency, respectively. Details of other symptoms were presented in Figure 1. Mean of white blood cell in study patients was 10530 ± 4490 (22 (62.8%) patients had leukocytosis. The mean of hemoglobin and ESR in study patients were 10.74 ± 1.47 and 66.79 ± 31.81. Twenty-two (62.8%) patients had positive findings for CRP. Hematuria and proteinuria were present in 5 (13.8%) and 10 (25.7%) patients, respectively. Four patients had pericardial effusion. Only one patient had a positive result for blood culture (Enterococcus sensitive to the Vancomycin and Amoxicillin). Involved valve in 19 patients was mitral including native and prosthetic. Besides, it was aortic valve in 16 patients (including native and prosthetic) and in one patient was native tricuspid valve.

4.2. Echocardiographic Findings
In echocardiography examination, five (14.2%) patients had mild, seven (20%) moderate and two (5.7%) severe aortic regurgitation. In patients with mitral valve involvement two, five (14.2%) and three patients had mild, moderate and severe mitral regurgitation, respectively. Mild and moderate tricuspid regurgitation were seen in 15 (42.8%) and six (16.7%) patients. Masses suggestive of vegetation were seen in 31 involved valves by TTE, and TEE revealed vegetations in all patients. In prosthetic AV, one patient presented with abscess and the others with vegetation. In those with prosthetic MV, five patients had vegetations and four vegetables with paravalvular leakage. Comparing TTE and TEE in diagnosis of IE patients, there were four false negative cases without false positive cases in TEE findings. Sensitivity and specificity of TEE in diagnosis of IE patients were 100% and 88.6%, respectively. Other findings of the two echocardiographic methods were the same. Findings of TEE and TTE echocardiographic studies were reported in Table 1.

Figure 1. Frequency of Clinical Symptoms in Endocarditis Patients

| Table 1. Echocardiographic Parameters<sup>a</sup> |
|-----------------------------------------------|
| Count Variable       | Mean ± SD  |
|----------------------|------------|
| LVEF                 | 49.72 ± 4.13 |
| LVEDD, mm            | 5.20 ± 0.70 |
| LVESD, mm            | 3.47 ± 0.77 |
| IVSD, mm             | 1.13 ± 0.1  |
| TRPG                 | 32.08 ± 10.77 |
| PAP, mm Hg           | 39.86 ± 11.40 |

<sup>a</sup> Abbreviations: IVSD, inter ventricular septal diameter; LVEDD, left ventricle end diastolic diameter; LVEF, left ventricle ejection fraction; LVESD, left ventricle end systolic diameter; PAP, pulmonary artery pressure; TRPG, tricuspid regurgitation (pressure gradient).
The mean of ejection fraction and pulmonary artery pressure were 49.72 ± 4.13 and 39.86 ± 11.40, respectively. Gender of patients had no significant association with cardiac valve involvement. The mean age of patients with mitral valve involvement was not significantly different from other patients (54.45 ± 12.43 vs. 57.20 ± 12.61; P value > 0.05). Patients with aorta valve involvement were significantly older than those without aortic valve involvement (66.64 ± 8.62 vs. 49.82 ± 9.58; P value = 0.00). Vegetation size in nine (25.8%) patients was higher than 10 mm.

4.3. Antibiotic Therapy and Outcome of Patients

Ceftriaxone was prescribed in all patients with a mean duration of 12.68 ± 12.71 days. After that Vancomycin had the highest frequency with a mean duration of 19.97 ± 12.55 days. After treatment period, six patients (17.1%) died so follow-up (FU) was not possible but other patients had FU. Among them two patients had a history of renal transplantation and received immunosuppressive agents and one patient had a history of arterial emboli and coronary artery bypass grafting operation. Six patients needed valve replacement after IE treatment.

5. Discussion

Endocarditis is a fatal disease and most of untreated patients might lose their life few weeks after the diagnosis (15). IE was developed in patients with valvular abnormalities due to several factors such as rheumatic carditis and some variants of streptococcus infection. These symptoms had higher frequency in developing countries due to higher prevalence of rheumatic heart disease (13). IV drug abusers, elderly patients with nosocomial exposure and hemodialysis patients were added to previous at risk patients (7). In the present study, fever had the highest frequency among clinical symptoms. Leukocytosis and high ESR were found in 62.8% of patients. Only one of our patients had positive findings for blood culture. In the present study, male involvement was 45%, but in similar studies male patients had a higher proportion (between four and even eight times more) than females (16). It could be due to more including IV drug abusers (had higher frequency in male patients) in other studies compared to this investigation. Fever and shaking were the most common clinical symptoms; therefore, we must pay more attention to IE diagnosis in suspected patients with risk factors such as cardiac valve replacement and IV drug abuse. In these cases, more laboratory tests and echocardiographic examination are needed to roll out IE diagnosis.

We had only one patient with positive finding for blood culture, which is the same with other similar studies; only in one study four positive blood cultures were reported among 20 patients with infective endocarditis (17). It could be due to different laboratory techniques for blood culture and previous use of antibiotics. Echocardiography is a noninvasive and available method of choice to evaluate presence of vegetations and its complications. It is recommended to perform TTE in all suspected patients and in both native and prosthetic valves (13, 14) it could help to diagnose IE. Nevertheless, TEE has a high sensitivity for IE, as in our study four patients had vegetations confirmed by TEE (one patient with prosthetic AV, one patient with native valve endocarditis of MV and two with prosthetic MV). However, in other patients with both techniques vegetations, paravalvular leakage or abscess could be evaluated with the same accuracy. Therefore, echocardiography has an important role in IE diagnosis, also repeating the procedure in appropriate intervals is helpful to avoid missing any patient with IE.

5.1. Limitations

The present study had some limitations; first, we performed the study on a small group of IE patients, thus we suggest performing further studies on larger sample size. Second, the present study was performed only in one hospital and multicenter studies with more patients might yield better results. Longer follow-up is also important for better evaluating these patients. Patients with endocarditis need more accurate clinical evaluation and early diagnosis in at risk populations, which can prevent disease progression. It is important to perform echocardiography (TTE and especially TEE) in endocarditis suspected patients after clinical and laboratory evaluation. In dead patients with negative blood culture, PCR was recommended to detect resistant pathogens.

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Authors’ Contributions

Dr. Hosseini was the physician who treated patients and referred them to Echolab; Dr. Bakshian did the analysis; Dr. Moshkani did echocardiographies and filled required questionnaires and did the FU; Dr. Sate filled the questionnaires and did the FU; Dr. Abdar gave important recommendations and revised the manuscript.

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