Cardiopulmonary Affection in Patients with Systemic Lupus Erythematosus

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Abstract

Background and Study Aims: Systemic Lupus Erythematosus (SLE) is a systemic autoimmune disease that can affect any part of the body. Early detection and quantification of pathological changes are important for assessing the benefits of cardiopulmonary prevention in SLE management. The aim of the study is to effect of SLE on cardiopulmonary system, and its early detection.

Methods: Fifty patients selected from those attending the outpatient clinics and inpatients, who admitted to Internal Medicine Department of Al-Azhar Assuit University Hospital, from May 2017 to May 2018, fulfilled the American College of Rheumatology (ACR) 1982 revised criteria for classification of SLE. All patients were subjected to complete history taking, clinical examination, routine investigations, transthoracic echo cardiography (Echo) and Computed Tomography (CT) of the chest. The damage was measured using the SLICC (Systemic Lupus International Collaborating Clinics)/ACR damage index (SDI). The disease flare was defined by the increase in the SLE Disease Activity Index (SLEDAI).

Results: The most common Echo finding was pericardial effusion seen in 20 patients (40%), followed by mitral regurgitation in 14 patients (28%), Mitral valve prolapse was seen in 13 patients (26%). The most common CT chest findings was ground glass opacity seen in 15 patients (30%) followed by pleural effusion seen in 14 patients (28%) and pleural thickening in 10 patients (20%). There was non-significant correlation between EF% and SLEDAI (p=0.95), but a negative significant correlation between disease duration and EF% (p=0.02).

Conclusion: All SLE patients even who clinically inactive disease should be screened for the presence of structural cardiac and chest abnormalities. Echocardiography and CT chest can be helpful as a noninvasive diagnostic tool for early detection of such abnormalities.

Keywords: Cardiac changes; Pulmonary changes; Systemic lupus erythematosus

Introduction

SLE is a systemic autoimmune disease that can affect any part of the body. As occurs in other autoimmune diseases, the immune system attacks the body’s cells and tissue, resulting in inflammation and tissue damage James et al., [1]. The course of the disease is unpredictable, with periods of activity alternating with remissions. The disease occurs nine times more often in women that in men, especially in women in child-bearing years ages 15 to 35 Rahmanand Isenberg [2]. Patients with SLE have a high risk of cardiovascular disease. Coronary artery disease, myocarditis, endocarditis, valvular disease and pericarditis are major manifestations of cardiac involvement in SLE patients. Multiple SLE-specific mechanisms, such as autoimmun responses, altered autoantibody and cytokine levels, and lipid dysfunctions can accelerate the progression of overall atherosclerotic burden. Therefore, early detection and quantification of pathological changes are important for assessing the benefits of cardiovascular prevention in SLE management Lin et al., [3]. The respiratory system is more commonly affected in SLE than in any other systemic autoimmune disease and that all its components may be affected. Pleuro-pulmonary involvement was defined as the presence of one or more of the following manifestations: pleuritis, pneumonitis, pulmonary arterial hypertension, shrinking lung syndrome, pulmonary fibrosis, pulmonary hemorrhage, pulmonary thrombosis and lung infarction Salinas et al., [4]. Disease activity varies over time and, at onset, symptoms are nonspecific and may include unexplained fever, extreme fatigue, muscle and joint pain and skin rash. SLE can present by arthritis, nephritis, heart and lung inflammation,
central nervous abnormalities and blood disorders Lapteva et al., [5].
This aims to study the effect of SLE on cardiopulmonary system, and its early detection.

Patients and Methods

This study was carried out on 50 patients (44 females and 6 males); aged 16 to 56 years, and the mean age was 31.12 ± 10.32 years. The mean disease duration was (11.50 ± 7.054). The study was approved by local ethical committee of Assuit Faculty of medicine; Al-Azhar University to evaluate and publish information gained data. All patients fulfilled the American College Of Rheumatology (ACR) 1982 revised criteria for classification of SLE. These patients were selected from those attending outpatient clinics and inpatients who admitted to the Internal Medicine Department of Al- Azhar Assuit University Hospital, from May 2017 to May 2018. Informed and written consent was taken from all patients.

Exclusion criteria

Diabetic patients, patients with associated rheumatic diseases and pregnant women.

All patients will be subjected to the following

History taking and clinical examination: 1) Socio-demographic data including age, sex, occupation, residence, marital status and special habits. History of deep venous thrombosis. 2) If married female: history of recurrent abortion. 3) History of fever, skin rash, purpuric eruption, ecchymotic patches, oral ulceration, photosensitivity, hair loss, arthralgia or arthritis and myalgia. 4) Full physical examination.

Laboratory investigations: Complete blood picture, ANA, Anti double stranded DNA, CRP, ESR.

Transthoracic echocardiographic examination at rest: To assess the following parameters: left ventricular Ejection Fraction (EF), Diastolic function, Valvular state. Pericardial state and pulmonary hypertension.

Computed Tomography (CT) of the chest to assess the following: Pulmonary parenchymal changes, Pulmonary interstitial changes, Pulmonary vascular changes and Mediastinal and pleural changes E-Measurement of activity index; the damage was measured using the SLICC (Systemic Lupus International Collaborating Clinics)/ACR Damage Index (SDI). The disease flare was defined by the increase in the SLE Disease Activity Index Rahman et al., [6].

Statistical analysis

Data were analyzed using Statistical Program for Social Science (SPSS) version 20.0. Quantitative data were expressed as mean ± Standard Deviation (SD). Qualitative data were expressed as frequency and percentage.

The following tests were done: 1) Independent-samples t-test
of significance was used when comparing between two means. 2) Chi-square ($X^2$) test of significance was used in order to compare proportions between two qualitative parameters. 3) Pearson’s correlation coefficient ($r$) test was used for correlating data. 4) Probability (P-value).

**Results**

This study was carried out on 50 patients (44 females and 6 males); aged 16 to 56 years, and the mean age was $31.12 \pm 10.32$ years. The mean disease duration was $(11.50 \pm 7.054)$ (Table 1). All patients fulfilled the American college of rheumatology (ACR) 1982 revised criteria for classification of SLE. These patients were selected from those attending outpatient clinics and inpatients who admitted to the Internal Medicine Department of Al-Azhar Assuit University Hospital, from May 2017 to May 2018. Of the 50 patients, 45 were positive for antinuclear antibody and 30 patients were positive for anti DNA antibody (Figures 1 and 2).

Echocardiography was done in all patients of whom 20 had normal echo findings. The most common finding was pericardial effusion seen in 20 patients, followed by mitral regurgitation in 14 patients; Mitral valve prolapse was seen in 13 patients. The other echocardiography findings were aortic regurgitation in 3 patients, tricuspid regurgitation in 4 patients, pulmonary hypertension in 4 patients, systolic dysfunction and diastolic dysfunction in 5 patients each. Regional hypokinesia was seen in 6 patients (Tables 2 and 3; Figures 3-6). As regard CT chest, there was 13 patients (26%) had normal CT chest. The most common CT chest findings was ground glass opacity seen in 15 patients (30%) followed by pleural effusion seen in 14 patients (28%), pleural thickening in 10 patients (20%), interlobular septal thickening in 9 patients (18%) & mediastinal lymph node enlargement, pulmonary artery trunk broadening, bronchiectasis, consolidation in 5 patients (10%) each, honey comb opacity in 4 patients (8%) and emphysema in one patient (2%) (Table 4 and Figures 6-10).

There was a significant correlation between disease duration and EF% ($p=0.02$) (Figure 11). But There was no significant correlation between disease duration and SLEDAI ($p=0.55$) (Figure 12). Also There was no significant correlation between EF% and SLEDAI ($p=0.95$) (Figure 13).

There was a significant correlation between CRP and SLEDAI ($p=0.01$) (Figure 14). But There was no significant correlation between ESR and SLEDAI ($p=0.92$) (Figure 15).

**Discussion**

Patients with SLE have a high risk of cardiovascular disease. Multiple SLE-specific mechanisms, such as autoimmune responses,
altered autoantibody, cytokine levels, and lipid dysfunctions can accelerate the progression of overall atherosclerotic burden. Therefore, early detection and quantification of pathological changes are important for assessing the benefits of cardiovascular prevention in SLE management Lin et al., [3]. Transthoracic Echocardiography can be helpful as a noninvasive diagnostic tool for early detection such abnormalities, resulting in earlier treatment and reduction in mortality and morbidity rates Abdel Gaffar et al., [7]. Also the respiratory system is more commonly affected in SLE than in any other systemic autoimmune disease and that all its components may be affected Salinas et al., [4]. Currently, CT is the best imaging technology in diagnosing the chest lesions in SLE patients. Chest lesions occur with high frequency, and the CT manifestations are complex and various, pulmonary interstitial changes are the most common Ping et al. [8]. In the present study, the most common Echocardiography finding was pericardial effusion seen in 40% of patients, followed by mitral regurgitation in 28% of patients; Mitral valve prolapse was seen in 26% of patients. The other echocardiography findings were aortic regurgitation in 6% patients, tricuspid regurgitation and pulmonary hypertension in 8% of patients, each. Systolic dysfunction and diastolic dysfunction in 10% of patients, each. Regional hypokinesia was seen in 12% of patients.

This was in agreement with Purushottam et al., [9] who did a study included 50 patients of Systemic Lupus Erythematosus of whom 44 patients were females and 6 patients were males. Majority of patients were between the age group 20 to 40 years. The maximum prevalence of Systemic Lupus Erythematosus was seen in the age group of 31 to 50 years. Echocardiography was done in all patients and found the same results as in our study. Furthermore, they found one patient had Libman-Sacks endocarditis and MVP with mitral regurgitation seen in 10 patients.
In another study by Shazzad et al., [10] fifty lupus patients were enrolled in the study and were evaluated by standard echocardiography with color Doppler. SLEDAI was applied for assessment of disease activity. Out of 50 patients 80% had abnormal echocardiographic findings. Pericardial thickening was found in 19 patients (38%), pericardial effusion in 10 patients (20%), diastolic dysfunction in 36 patients (72%), hypokinesia of ventricular wall in 4 patients (8%), overall valvular abnormalities 20%, commonest being aortic regurgitation in 6 patients (12%), followed by mitral regurgitation in 4 patients (8%), pulmonary hypertension in 3 patients (6%).

In our study, There was a significant correlation between disease duration and EF% (p<0.02). This was in agreement also with Shazzad et al., [10] as they found significant relationship between disease duration and cardiac abnormalities (p<0.01).

As regard CT chest in our study, there was 13 patients (26%) had normal CT chest. The most common CT chest findings was ground glass opacity seen in 15 patients (30%) followed by pleural effusion seen in 14 patients (28%), pleural thickening in 10 patients (20%), interlobular septal thickening in 9 patients (18%) & mediastinal lymph node enlargement, pulmonary artery trunk broadening, bronchiectasis, consolidation in 5 patients (10%) each, honey comb opacity in 4 patients (8%) and emphysema in one patient (2%).

This was in agreement with Ping et al., [8] who did a study included 39 patients (34 females and 5 males), aged from 19 to 74 years old, the mean age was (44.13 ± 12.17) years old. HRCT chest was done for all patients and found the same results as in our study. Furthermore, they found Funicular opacity in 20 patients, Sub-pleural line in 9 patients and Mosaic sign in 3 patients.

In other study by Kakati et al., [11] 38 lupus patients, were enrolled in the study and were evaluated by HRCT chest. The age of the patients ranged from 12 to 60 years (mean age 26.18 yrs) with maximum incidence between 16 to 25 years of age & 3 patients (7.89%) were males. Pleural effusion seen in 4 patients (10.53%), pleural thickening seen in one patient (2.63%), Sub-pleural bands seen in 2 patients (5.26%), Lymphadenopathy seen in 2 patients (5.26%), Bronchiectasis seen in 3 patients (7.89%), Bronchial thickening seen in 15 patients (39.47%), Parenchymal bands seen in 14 patients (36.84%), Air space consolidation seen in 2 patients (5.26%), Ground glass opacification seen in 10 patients (26.32%), Parenchymal micronodules seen in 3 patients (7.89%), Bullae seen in 2 patients (5.26%).

**Conclusion**

It is common to find cardiac and chest involvement in SLE patients. All SLE patients even who clinically inactive disease should be screened for the presence of structural cardiac and chest abnormalities. Trans-thoracic Echocardiography and CT chest can be helpful as a noninvasive diagnostic tool for early detection of such abnormalities, resulting in earlier treatment and reduction in mortality and morbidity rates.

**References**