INCIDENCE AND RISK FACTORS FOR GANGRENE IN PATIENTS WITH SYSTEMIC SCLEROSIS FROM THE EUSTAR Cohort

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Background: Digital ulcers (DUs) affect about half of systemic sclerosis (SSc) patients during disease course. In some patients, peripheral vasculopathy can promote critical ischemia and gangrene, severe complications with potential life threatening consequences. Recently the DUO registry suggested a 18% prevalence of gangrene in DU-SSc patients, with smoking and a high number of DUs being predictive factors. However, little is known about gangrene in unselected SSc patients.

Objectives: To investigate the prevalence, incidence and risk factors for gangrene in the EUSTAR cohort.

Methods: We included patients from the EUSTAR database satisfying the ACR classification criteria for SSc, with at least one visit recording data on gangrene. We extracted from this database data regarding the reporting of DUs, DUs history and digital gangrene. Centres were asked for supplementary data on traditional cardiovascular (CV) risk factors. We analysed by univariable and multivariable logic regression the cross-sectional relationship between gangrene and its potential risk factors such as history of DUs, cutaneous subset, disease duration, autoantibodies, traditional CV risk factors. Furthermore, longitudinal data were analysed by Cox proportional hazards regression.

Results: 1757 patients matched the inclusion criteria (age at inclusion 55.9±14.5 years, disease duration since first non-Raynaud’s phenomenon 7.9±10.3 years and from onset of Raynaud’s phenomenon (RP) 11.1±11.0 years, male sex 16.7%, 24.6% diffuse cutaneous subset (DcSSc)). At inclusion, 8.9% of patients had either current or previous digital gangrene, 15.7% had current DUs and a further 25.8% had previously had DUs. Among the potential risk factors, older age, a history of DUs and the DcSSc subset were statistically significant risk factors in the cross-sectional multivariable model.

For the longitudinal part, during the entire follow-up (median [Q1,Q3] 13.1 [9.6, 19.3] months), 16/771 patients had incident gangrene (2.1%), accounting for an incidence of 1.73/100 patient-years. All 16 patients who developed incident gangrene were the DcSSc subset and longer disease duration: hazard ratio [95% confidence interval]: 8.97 [2.90–27.71] and 1.08 [1.04–1.13] respectively, both p<0.001.

Conclusions: In unselected SSc patients, gangrene still occurs in about 9% of SSc patients. Of the most importance, a history of DUs and, to a lesser extent, the DcSSc subset are strongly and independently associated with gangrene, while traditional CV risk factors were not identified as risk factors. Our results confirm that gangrene is still a concern in SSc. They emphasise the importance of microvascular SSc-associated disease in the pathogenesis of gangrene and suggest that the DcSSc subset should be prioritised for risk-stratification of the patients.

GLOBAL LONGITUDINAL STRAIN AS EARLY PREDICTOR OF SYSTOLIC DYSFUNCTION IN SYSTEMIC SCLEROSIS

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Background: Systemic sclerosis (SSc) is a chronic autoimmune disease of unknown etiology, characterised by microvascular abnormalities, immune abnormalities and progressive cutaneous and internal organs fibrosis. Subclinical heart disease in SSc patients is common but difficult to detect through conventional imaging.

Objectives: We sought to evaluate speckle-tracking derived global longitudinal strain (GLS) as an early marker of subclinical systolic dysfunction in patients with SSc.

Methods: We enrolled 52 patients with SSc and 52 age and gender matched controls. Patients with structural heart disease, heart failure, atrial fibrillation or pulmonary hypertension were excluded. An echocardiographic exam was performed for all patients, and standard and specke-tracking derived variables for the systolic and diastolic function of the left ventricle (LV) and right ventricle (RV) were acquired. SSc variant, antibodies pattern, cardiovascular risk factors and involvement of other organ systems were recorded.

Results: Common parameters of left and right systolic function did not differ between SSc patients and controls and were on average well above the cut-off for normality (all p>0.05). LV and RV GLS were significantly impaired in patients with SSc when compared to healthy controls (−19.2% vs. −21.1%; p=0.009 and −18.2% vs. −22.3%; p=0.012 respectively). In patients with SSc, GLS impairment was greater in basal segments when compared to midventricular and apical ones and homogenous between the endo-, meso-, or epicardial layers of the RV, while LV showed an eccentric pattern with the epicardial layers mostly impaired. Using “20% as a cut-off for GLS, SSc patients had a 2.5-fold increased risk of subclinical LV systolic impairment (OR 2.5; 95% CI 1.4–7.7; p=0.004).

Conclusions: While traditional parameters are ineffective in detecting subclinical systolic impairment, a reduced GLS is common in patients with SSc and is significant for both LV and RV. While GLS impairment recognises a basal-apical gradient, transmural heart involvement seems different between RV and LV, suggesting a different mechanism of disease between the two ventricles.

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