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AB0425

CLINICAL AND PATHOLOGICAL FEATURES OF BREAST CANCER IN PATIENTS WITH SYSTEMIC SCLEROSIS: PRELIMINARY DATA FROM THE SCLERO-BREAST STUDY

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Background: Systemic Sclerosis (SSc) is a rare and life-threatening connective tissue disease characterized by vascular dysfunction, specific autoimmune abnormalities and fibrosis of the skin and internal organs. Previous studies have shown a 1.5-fold increase in cancer risk in SSc patients compared with the general population, including breast cancer (BC). The relationship between BC and SSc has long been discussed but past research has been contradictory and inconclusive on this topic.

Objectives: The aim of our project was to analyze clinical and pathological characteristics of BC developed by SSc subjects and possible correlations with scleroderma features. Here we present the preliminary data from the Sclero-Breast study.

Methods: Our observational retrospective multicenter study enrolled 33 SSc women with a personal history of BC identified at two Rheumatology/SSc Units in the north of Italy between January 2017 and December 2019 (lc/dcSSc 23/9, 1 unknown; mean age at SSc onset 57 years, range 32-73). All patients underwent general and instrumental assessment: smoking habits; presence of skin ulcers, calcinosis, teleangiectasia; presence of gastro-intestinal and kidney involvement; interstitial lung disease (at HR-CT); pulmonary function tests; ECG abnormalities; echocardiographic assessment of pulmonary arterial hypertension (PAH); videocapillaroscopic pattern; autoantibody profile; exposure to immunosuppressive and vasoactive therapies; status at last follow-up evaluation and cause of death. Clinical and pathological characteristics of BC were also evaluated: age at diagnosis; menopausal status; histotype; hormone receptor status; MIB1, HER2 expression; clinical and pathological stage at diagnosis; metastatic sites; type of loco-regional treatment (surgery and radiotherapy); type of systemic treatment (neoadjuvant/adjuvant chemotherapy and endocrine treatment); other cancers and time from diagnosis of the first disorder to the second one.

Results: A total of 54.5% of subjects developed BC before SSc (median interval of 5 years), whereas 45.5% of patients developed BC after SSc (median delay of 8 years). 54.5% of patients showed interstitial lung disease and the cause of death of the 6 deceased subjects was PAH. A significant association ($p < 0.05$) was observed between the use of immunosuppressive therapy and diffuse skin extension, negative ACA, positive Anti-Scl-70 and interstitial lung disease, but not with BC status. 93.1% of patients were diagnosed with an early-stage tumor, 70.8% of invasive carcinomas with a low MIB-1, 8.3% with a tubular histotype, while 42.8% presented with a Luminal A-like tumor. 66.6% underwent breast conserving surgery and 55.5% RT after surgery. 40% of patients developed interstitial lung disease after RT and 20% dcSSc.

Conclusion: According to our preliminary data, SSc patients developed BC at good prognosis, suggesting a de-escalation strategy of cancer therapies. On these grounds, a proper screening is mandatory in order to allow for early cancer detection in SSc patients. Further investigations on larger numbers of patients are needed. First of all, they would further clarify the intriguing relationship between BC and SSc. Secondly, they would help to explore the common biological and molecular pathways at the basis of these two disorders, with the aim to improve BC diagnosis and prognosis and to personalize oncological targeted treatments in this subset of fragile patients.

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AB0426

CANCER SCREENING IN IDIOPATHIC INFLAMMATORY MYOPATHIES: TEN YEARS EXPERIENCE FROM A SINGLE CENTER

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Background: There is a well-recognized association between cancer and myositis, so cancer screening at diagnosis is recommended.

Objectives: We aim to report the results of our cancer screening strategy and to ascertain the reliability of using PET/CT to identify cancer-associated myositis (CAM) in a large cohort of patients with myositis from a single center over 10 years.

Methods: This retrospective observational study included all patients diagnosed with any type of myositis except for inclusion body myositis. Cancer screening strategy was individualized according to clinical and serological data, including PET/CT as the main test to detect occult cancer (OC). Procedures derived from a positive PET/CT were registered. Qualitative data expressed as percentages, and quantitative data as the median with the interquartile range were analyzed. A ROC curve was used to estimate the reliability of PET/CT for CAM diagnosis.

Results: Seventy-seven out of 131 patients underwent a PET/CT for OC screening. The performance of the PET/CT in patients with myositis at disease onset yielded an area under the curve ROC of 0.87 (0.73-0.97) for CAM diagnosis. Invasive procedures in 7 (9%) patients without a final diagnosis of cancer did not cause derived complications. Patients not evaluated for OC did not develop cancer after a median follow-up of 3.3 years (1.7-6.7).

Conclusion: Cancer screening strategy should be individualized. PET/CT at myositis onset seems to be an efficient approach to rule out CAM. This practice does not seem to significantly increase harm to patients related to the additional tests needed to clarify inconclusive results.

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AB0427

ANTI-NOR90 AUTOANTIBODIES: FAVORABLE OR UNFAVORABLE PROGNOSIS?

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Background: Anti-NOR 90 autoantibodies (anti-NOR90 Ab) are autoantibodies that target nucleolar transcription factor 1 or hUBF, involved in transcription of RNA polymerase I. These autoantibodies have been detected in 6.1% of patients with Systemic Sclerosis (SSc), but their clinical or prognostic significance has not been clearly defined. Anti-NOR90 Ab have been mostly associated with limited scleroderma with mild organ involvement and can also be found in other rheumatic diseases such as rheumatoid arthritis, systemic lupus erythematosus or Sjogren's syndrome.

Objectives: The aim of this study was to identify the main clinical characteristics of patients with positive anti-NOR90 in our Centre.

Methods: This is a retrospective, descriptive, cross-sectional study of all patients with positive anti-NOR90 Ab between January 2013 and December 2020 in a single center. Autoantibodies testing was performed using Euroimmun EUROLINE SSc profile IgG autoAb assay kit. Patient demographics, clinical characteristics, associated diagnoses, laboratory and immunological findings were collected.

Results: We identified a total of 26 patients with at least a positive value for anti-NOR90 Ab (Table 1). In most cases anti-NOR90 patients were ANA positive, predominantly with nucleolar pattern and coexisted with other SSc autoantibodies. 12 patients had rheumatic diseases and two had SSc, both with limited cutaneous SSc and absence of organ involvement. 14 patients had no definite diagnosis. Clinical features of anti-NOR90 patients are represented in Figure 1. Five patients presented Raynaud's phenomenon, two cases with pathological nailfold capillaroscopy and one patient had SSc. There was no patient with skin ulcers, calcinosis, interstitial lung disease or pulmonary hypertension. Four patients had gastroesophageal reflux disease and one patient presented antral vascular ectasia. Six patients developed some neoplasm.

Figure 1. Clinical characteristics of anti-NOR90 Ab patients.

Conclusion: In our case series anti-NOR90 Ab were associated with multiple rheumatic diseases with heterogeneity of clinical manifestations. We did not observe a further progression to SSc or presence of organ involvement or severe scleroderma, so these autoantibodies could be related with a favorable prognosis. In contrast with previous reports, a striking association with cancer has been detected in our population.

Table 1. Demographic characteristics and main diagnoses of anti-NOR90 positive patients.

Characteristics	Total Anti-NOR90: 26 patients
Sex, n	19 women/ 7 men
Age, mean (years)	58.9 IQR [46.3-72.2]
Race, n	Asian: 1; Hispanic: 7; Caucasian: 18
Smoker, n	3
Positive ANA (>1/160), n	24
Pattern, n	7 Homogeneous, 4 Nucleolar, 4 Speckled, 1 Centromere, 5 Speckled -nucleolar, 3 Homogeneous-nucleolar.
Positive ENA, n	3
	2 Anti-SSA-Ro52 and Ro60, 1 anti-RNP and anti-Sm
Systemic sclerosis autoantibodies, n	<ul style="list-style-type: none"> • Anti-Ku: 7 • Anti-U3RNP (Fibrilarin): 6 • Anti-RNA polymerase III: 5 • Anti Th/To: 4 • Anti-centromere: 4 (CENP B +/- CENPA) • Anti-topoisomerase I: 2 • Anti-Ro52: 3 • Anti-PM-Scl: 3
Main diagnosis, n	12 Rheumatic diseases: 2 systemic Sclerosis (2 limited/0 diffuse) 1 rheumatoid arthritis 1 LES 1 Sjögren's syndrome, 3 undifferentiated connective tissue disease 2 overlap (1 Sjögren + LES, 1 Sjögren + MCTD 1) s: 1 morphea, 1 cutaneous graft versus host disease 6: 2 solid organ cancer (bladder, kidney), 1 lung adenocarcinoma, 1 multiple myeloma, 1 acute myeloid leukemia: 1 basal cell carcinoma.
Neoplasm, n	

Abbreviations: LES: systemic lupus erythematosus; MCTD: mixed connective tissue disease

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AB0428

ULTRASOUND ASSESSED NAIL BED THICKNESS AS AN EARLY SCREENING MARKER FOR INTERSTITIAL LUNG DISEASE IN SCLERODERMA

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Background: Interstitial lung disease (ILD) is the most frequent pulmonary manifestation of scleroderma and is seen in 80% of the patients with diffuse and in 20% of the limited forms of scleroderma.¹ Even seen in early stages of the disease with varying degrees of severity, it does not correlate with the clinical manifestations.² Hence, it is difficult to select patients for ILD screening based on symptoms. Digital clubbing is a common feature of ILD and has been associated with poor prognosis.³

Objectives: To assess the ultrasound measured nail bed thickness (NBT) in cases of scleroderma as a screening marker to detect ILD and its inter-digital variation.

Methods: A prospective observational study involving 15 patients each of scleroderma with ILD, without ILD and controls visiting our clinic from August 2020 to December 2020 were included. The presence of ILD was confirmed by HRCT thorax. Patients with a history of smoking, alcohol intake or pre-existing cardio-respiratory illnesses were excluded. Ultrasound assessed NBT was calculated using Philips affiniti 70 eL18-4 probe. A total of 436 nail beds were recorded (142 of scleroderma with ILD, 144 of scleroderma without ILD and 150 of control group). 14 nails were excluded due to trauma. Statistical analysis was done using ANOVA test. Results were considered significant at p value < 0.05.

Results: On comparing nail bed thickness among scleroderma with ILD, scleroderma without ILD and control groups, the difference in NBT was statistically significant in all the three groups, except for left 1st finger (Table 1). NBT's among scleroderma patients without ILD and control group did not show significant difference between the two groups. Whereas, when scleroderma patients with ILD were compared with control group, statistical significance were found (p<0.001) in all the fingers of both the hands, except 1st and 2nd finger of left and right hand respectively. On comparing NBT's among individual fingers of right hand in patients of scleroderma with ILD and without ILD, a statistically significant difference among the two groups in 2nd (p=0.002), 4th (p=0.009) and 5th finger (p<0.001) was noted. In left hand, same fingers had statistically significant difference (p<0.001).

Conclusion: Patients of scleroderma with ILD are predisposed for an increase in the NBT, which can be detected using ultrasound. Thus, an increased NBT may have a role in early screening for the detection of ILD in scleroderma patients. Further studies with large patient data should be done to look at ultrasound assessed NBT as it may be utilized as a marker for ILD in scleroderma, as a widely available and an inexpensive tool.

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Table 1. Nail bed thickness in patients of scleroderma with ILD, without ILD and control groups.

		Scleroderma with ILD		Scleroderma without ILD		Control		
		n=15		n=15		n=15		
	Finger	Mean	SD	Mean	SD	Mean	SD	p-value
Right	1 st	1.86	0.239	1.71	0.081	1.69	0.107	0.021
	2 nd	1.73	0.303	1.50	0.079	1.61	0.108	0.003
	3 rd	1.89	0.200	1.75	0.091	1.50	0.131	<0.001
	4 th	1.80	0.263	1.39	0.147	1.35	0.078	<0.001
Left	5 th	1.80	0.119	1.24	0.095	1.29	0.069	<0.001
	1 st	1.98	0.442	1.72	0.084	1.68	0.050	0.078
	2 nd	1.82	0.123	1.57	0.125	1.57	0.092	<0.001
	3 rd	2.05	0.309	1.76	0.078	1.42	0.109	<0.001
	4 th	1.92	0.246	1.36	0.130	1.35	0.107	<0.001
	5 th	1.81	0.158	1.15	0.368	1.23	0.103	<0.001

Figure 1. NBT measurement of right 2nd finger in scleroderma patient with ILD on ultrasound