

MHz transducer a statistically significantly higher median DT was showed in all skin areas, included thighs ( $p < 0.01$ ). Finally, a positive statistically significant correlation was observed between the two transducers in the evaluation of DT ( $p < 0.0001$ ), as well as between both probes and mRSS ( $p < 0.0001$  for both).

**Conclusions:** This study suggests that subclinical dermal involvement may be detectable by skin high frequency US already in patients with limited cutaneous SSc. This study confirms that DT can be better assessed in SSc patients by using a 22 MHz US probe, and suggests that DT might be underestimated by using US probes of lower frequency (18 MHz). However, the DT values obtained using both probes resulted significantly correlated together and with the mRSS.

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**OP0038 MYOSITIS AUTOANTIBODIES OUTPERFORM CLINICAL SUBGROUP CLASSIFICATION IN PREDICTING MUSCLE WEAKNESS IN MYOSITIS PATIENTS**

I. Pinal-Fernandez<sup>1</sup>, W. Huang<sup>1</sup>, M. Casal-Dominguez<sup>1</sup>, E. Tiniakou<sup>2</sup>, C. Parks<sup>1</sup>, K. Pak<sup>1</sup>, J. Albayda<sup>2</sup>, J.J. Paik<sup>2</sup>, S.K. Danoff<sup>3</sup>, L. Christopher-Stine<sup>2</sup>, A.L. Mammen<sup>1</sup>. <sup>1</sup>Muscle Disease Unit, NIAMS/NIH, Bethesda; <sup>2</sup>Rheumatology; <sup>3</sup>Pulmonology, Johns Hopkins University, Baltimore, United States

**Background:** Myositis patients may be classified as belonging to one of four clinical groups: dermatomyositis (DM), polymyositis (PM), clinically amyopathic dermatomyositis (CADM) or necrotizing myositis (NM). Alternatively, myositis patients may be classified according to myositis autoantibody status.

**Objectives:** The aim of this study was to determine whether clinical groups or myositis autoantibodies provide better prognostic categories with regard to muscle involvement in these patients.

**Methods:** All Johns Hopkins Myositis Center patients from 2002 to 2015 with a myositis specific autoantibody confirmed by two different immunologic techniques were included. Autoantibody groups accounting for less than 2% of the final sample size were excluded. Strength (analyzed as the average of deltoid and hip flexor strength using Kendall's scale) and log transformed CK levels were compared between the different autoantibody groups using multilevel regression models adjusted for age, time from disease onset, sex, race and treatments. Models with different combinations of key variables were compared using the likelihood ratio test to ascertain if autoantibody groups and clinical subgroups provided the same amount of information regarding muscle weakness and CK levels over time.

**Results:** 483 patients with 4181 visits were included and 10 different autoantibody groups were identified. Muscle weakness and CK levels followed a gradient among both antibody and clinical groups. Anti-SRP patients had the greatest weakness, followed by anti-HMGCR, anti-Mi2 and anti-NXP2, and then anti-Jo1. CK levels were highest in anti-HMGCR patients, followed by anti-SRP, anti-PL7, anti-Jo1 and anti-Mi2. Interestingly, strength and CK levels were dissociated in two groups: anti-NXP2 patients had significant weakness with low CK levels and anti-PL7 patients were relatively strong despite high CK levels. Multilevel regression models showed autoantibody groups explained the strength and the CK variability better than the clinical groups (AIC difference > 20). Indeed, adding clinical groups to a model using only autoantibodies did not improve the model's ability to predict strength ( $p = 0.2$ ) and only mildly improved its ability to predict CK ( $p = 0.01$ ). In comparison, adding the autoantibodies to a model using the clinical groups resulted in a marked improvement in predicting both CK and strength (both  $p < 0.001$ ).

**Conclusions:** In patients with myositis, autoantibody status predicts strength and CK levels better than clinical grouping.

**Disclosure of Interest:** None declared

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**SLE, Sjögren's and APS - clinical aspects**

**OP0039 A POPULATION-BASED STUDY ON MORTALITY AND THE INFLUENCE OF MEDICATION USE IN 4356 PATIENTS WITH SYSTEMIC LUPUS ERYTHEMATOSUS AND 21845 MATCHED CONTROLS FROM THE UNITED KINGDOM**

I.E. Bultink<sup>1</sup>, A. Lalmohamed<sup>2,3</sup>, F. de Vries<sup>3,4</sup>. <sup>1</sup>Rheumatology, Amsterdam Rheumatology and Immunology Center, VU University Medical Center, Amsterdam; <sup>2</sup>Clinical Pharmacy, University Medical Center Utrecht; <sup>3</sup>Pharmacoepidemiology and Clinical Pharmacology, Utrecht University, Utrecht; <sup>4</sup>Clinical Pharmacy and Toxicology, Maastricht University Medical Centre, Maastricht, Netherlands

**Background:** Systemic lupus erythematosus (SLE) has been associated with

an increased mortality rate. However, population-based data on all-cause, age-specific and sex-specific mortality risk are limited and data on the influence of medication exposure on mortality risk in SLE are scarce.

**Objectives:** To estimate the magnitude of the risk from all-cause, age-specific, and sex-specific mortality in patients with SLE and relative risks compared with matched controls, and to evaluate the influence of medication exposure on mortality risk in SLE.

**Methods:** We conducted a population-based cohort study using the Clinical Practice Research Datalink (from 1987 to 2012). Each SLE patient ( $n = 4356$ ) was matched with up to 6 controls ( $n = 21845$ ) by age and sex. Multivariate Cox regression analysis estimated adjusted relative rates (RR) of mortality, and time interaction terms to evaluate mortality timing patterns. Time-dependent Cox models were used to evaluate the association of glucocorticoid use and hydroxychloroquine use on mortality and were adjusted for age, sex, lifestyle parameters, comorbidities and comedication.

**Results:** A total of 442 out of 4356 SLE patients died during the study period. Patients with SLE had an increased mortality rate for all-cause mortality compared with age- and sex-matched subjects, after adjustment for confounders (adjusted RR 1.80, 95% CI 1.57–2.08). Glucocorticoid use in the previous six months raised the mortality rate while the adjusted RR was 45% decreased with low dose hydroxychloroquine use. The RR was highest in patients aged 18–39 years (adjusted RR 4.87, 95% CI 1.93–12.3) and slightly higher in females (adjusted RR 1.82, 95% CI 1.56–2.13) compared to male patients (adjusted RR 1.68, 95% CI 1.19–2.39). The mortality rate was significantly increased for patients with a history of dementia, seizures, diabetes, cancer, and renal disease (Table 1).

Table 1. Risk of all-cause mortality within SLE patients ( $n = 4356$ ), stratified according to organ damage (reference = no risk factor)

	Person years (x1000)	Deaths	IR (/1000)	Adjusted RR* (95% CI)
Dementia	0.1	14	140.0	2.99 (1.74–5.14)
Seizures	1.4	37	26.4	2.33 (1.66–3.28)
Cerebrovascular event	1.9	73	38.4	1.28 (0.99–1.65)
Renal disease	2.0	86	43.0	1.40 (1.09–1.78)
Osteoporotic fracture	5.1	110	21.6	1.06 (0.85–1.32)
Diabetes mellitus	0.9	45	50.0	1.90 (1.39–2.59)
Malignancy	2.0	95	47.5	1.90 (1.50–2.40)

\*Adjusted for: recent use of corticosteroids, recent use of antimalarials, recent use of benzodiazepines.

**Conclusions:** Patients with SLE have a 1.8-fold increased mortality rate compared with the general population. Glucocorticoid use, female sex and young age are associated with an increased mortality risk while low dose hydroxychloroquine use significantly reduces the mortality rate. In addition, special attention should be paid to lupus patients with neuropsychiatric complications, diabetes, malignancy or renal disease since these subgroups of patients are at high risk of death.

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**OP0040 INTEGRATION OF SALIVARY-GLAND ULTRASONOGRAPHY IN CLASSIFICATION CRITERIA FOR PRIMARY SJÖGREN'S SYNDROME: AN INTERNATIONAL VIGNETTE-BASED STUDY**

S. Jousse-Joulin<sup>1</sup>, F. Gatineau<sup>2</sup>, C. Baldini<sup>3</sup>, S. Arends<sup>4</sup>, F. Barone<sup>5</sup>, A. Baer<sup>6</sup>, H. Bootsma<sup>7</sup>, S. Bowman<sup>7</sup>, P. Brito-Zeron<sup>8</sup>, D. Corneec<sup>9</sup>, T. Dorner<sup>10</sup>, S. De Vita<sup>11</sup>, B. Fisher<sup>12</sup>, D. Hammenfors<sup>13</sup>, M. Jonsson<sup>14</sup>, X. Mariette<sup>15</sup>, V. Milic<sup>16</sup>, T. Nakamura<sup>17</sup>, W.-F. Ng<sup>18</sup>, E. Nowak<sup>2</sup>, A. Rasmussen<sup>19</sup>, R. Seror<sup>20</sup>, C. Shiboski<sup>21</sup>, T. Nakamura<sup>22</sup>, M. Ramos-Casals<sup>23</sup>, A. Vissink<sup>4</sup>, A. Saraux<sup>24</sup>, V. Devauchelle-Pensec<sup>24</sup>. <sup>1</sup>Rheumatology; <sup>2</sup>CIC, CHU Brest, BREST, France; <sup>3</sup>Rheumatology, University of Pisa, Pisa, Italy; <sup>4</sup>Rheumatology, University Medical Center Groningen, Groningen, Netherlands; <sup>5</sup>Rheumatology, University, Birmingham, United Kingdom; <sup>6</sup>Department of Rheumatology, Johns Hopkins University School, Baltimore, United States; <sup>7</sup>Rheumatology, university of Birmingham, Birmingham, United Kingdom; <sup>8</sup>Department of Rheumatology, Hospital CIMA-Sanitas, Barcelona, Spain; <sup>9</sup>Thoracic Disease Research Unit, Mayo Clinic Rochester, Rochester, United States; <sup>10</sup>Rheumatology, Charité Universitätsmedizin Berlin, Berlin, Germany; <sup>11</sup>Rheumatology, Santa Maria della Misericordia University-Hospital, Udine, Italy; <sup>12</sup>Rheumatology Research Group and Arthritis Research UK Rheumatoid Arthritis RACE, University of Birmingham, Birmingham, United Kingdom; <sup>13</sup>Rheumatology; <sup>14</sup>Clinical Science, university of Bergen, Bergen, Norway; <sup>15</sup>Rheumatology, INSERM U1184, Center Autoimmune Diseases, Paris-Sud, le Kremlin Bicetre, France; <sup>16</sup>Rheumatology, university of Belgrade, Belgrade, Serbia; <sup>17</sup>Department of Immunology and Rheumatology, Nagasaki University Graduate School of Biomedical Sciences, Nagasaki, Japan; <sup>18</sup>Rheumatology, University of New Castle, New Castle, United Kingdom; <sup>19</sup>Arthritis & Clinical Immunology Program, Oklahoma Medical Research Foundation, Oklahoma, United States; <sup>20</sup>Rheumatology, INSERM U1184, Center for Autoimmune Diseases, Paris-Sud, le Kremlin Bicetre, France; <sup>21</sup>Department of Dentistry, University of California, San Francisco, United States; <sup>22</sup>Rheumatology, Nagasaki University Graduate School of Biomedical Sciences, Nagasaki, Japan; <sup>23</sup>Hospital Clinic, university of Barcelona, Barcelona, Spain; <sup>24</sup>Rheumatology, Brest University and INSERMU1227, BREST, France

**Background:** The recent classification criteria sets for primary Sjögren's syn-

drome (AECG 2002 and ACR/EULAR 2017) did not include major salivary glands ultrasonography (SGUS).

**Objectives:** The UTOPIA study was undertaken to determine if and how SGUS may improve the ACR/EULAR criteria.

**Methods:** Twenty four international experts in pSS evaluated on an internet-secure relational database 512 randomly realistic vignettes derived from 150 patients with suspected pSS included in the french DiapSS cohort. Each vignette contained sections on "history" (duration of the symptoms, gender, age), clinical symptoms (dry mouth, dry eyes and systemic manifestations), results of the SGUS evaluation (score > ou < to 2), and results of the major tests to diagnose pSS (Schirmer's test, ocular staining score (OSS), salivary flow, focus score on salivary biopsy, presence of anti-SSA antibodies). Each expert had to score the diagnosis of pSS as absent, unlikely, likely or present for 64 vignettes. Each vignette was evaluated by 3 experts. Diagnosis of pSS was obtained when at least 2 of 3 considered it as likely or present. Univariate and multivariate analysis (Wald test) were performed to evaluate the association between the SGUS criteria, the ACR/EULAR criteria and its different individual items with the diagnosis of pSS as defined by the experts. Data were then replicated on independent cohorts of suspicion of pSS.

**Results:** Univariate and multivariate analyses confirmed that ACR/EULAR criteria and SGUS were independently associated with the diagnosis of pSS. Disease duration, OSS and ocular dryness were not associated with the diagnosis of pSS. Only 6 variables were selected by logistic regression analysis: presence of anti-SSA (weight:4), focus score (weight:3), SGUS (weight:2), Schirmer's test (weight:1), dry mouth (weight:1) and salivary flow rate (weight:1). According to ROC curve analysis, a score of  $\geq 5$  had 96% Se and 84% Sp, compared with 90% Se and 84% Sp for the ACR/EULAR criteria. The corrected C statistic (AUC) for the new weighted score was 0.98.

**Conclusions:** Inclusion of the SGUS item in the ACR/EULAR criteria improves their diagnostic performance.

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#### OP0041 ALL-CAUSE, CARDIOVASCULAR AND MALIGNANCY RELATED MORTALITY IN SYSTEMIC LUPUS ERYTHEMATOSUS (SLE): A POPULATION-BASED STUDY

B. Moghaddam<sup>1</sup>, J.A. Avina-Zubieta<sup>2,3</sup>, E.C. Sayre<sup>2</sup>. <sup>1</sup>Division of Medicine, University of British Columbia, Vancouver; <sup>2</sup>Arthritis Research Canada, Richmond; <sup>3</sup>Division of Rheumatology, University of British Columbia, Vancouver, Canada

**Background:** SLE is associated with increased risk of overall mortality; however mortality trends due to specific causes including cardiovascular disease (CVD), malignancies or other causes are largely unknown.

**Objectives:** Our objective was to assess trends in cause-specific mortality among SLE patients between January 1, 1997 and December 31, 2012 in a general population-based context.

**Methods:** We conducted a population-based matched cohort on SLE patients diagnosed between January 1, 1997 and December 31, 2012 using an administrative health database from the province of British Columbia, Canada. We identified all incident cases of SLE and up to 10 non-SLE controls matched on sex, age, and calendar year of study entry. The cohort was divided into two cohorts based on year of SLE diagnosis (1997–2004 and 2005–2012). All-cause mortality and cause-specific incidence of death rates (IR) were calculated. Cox proportional hazard regression models were used to estimate the mortality hazard ratios (HR), adjusting for possible confounders (i.e. Charlson Comorbidity Index, number of outpatient visits, hospitalization, cardiovascular medications, glucocorticoids and NSAIDs at baseline).

**Results:** 4238 SLE and 42380 matched controls were studied. SLE patients had significantly increased all-cause mortality with HR 1.29 (95% CI, 1.15–1.46) and increased cause-specific mortality from CVD and other causes with HRs of 1.43 (95% CI, 1.15–1.79) and 1.74 (95% CI, 1.46–2.09), respectively. The cohorts did not differ in the rate of death from malignancy. SLE patients had an approximately 2-fold increase in death from other causes in both early (HR 1.86 (95% CI 1.33–2.60)) and recent cohorts (HR 1.90 (95% CI 1.42–2.56)). There was no significant improvement in all-cause and cause-specific mortality trends between the two cohorts.

**Conclusions:** This study demonstrates that despite advances in therapy with novel biologic agents, there are no significant differences in all-cause and CVD mortality from SLE between early and recent cohorts. Death from other causes, which includes a composite of death related to for example renal disease and infections, remains high suggesting areas for future targeted research and therapy.

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#### OP0042 BACTEREMIA IN SYSTEMIC LUPUS ERYTHEMATOSUS PATIENTS FROM RELESSER REGISTRY: RISK FACTORS, CLINICAL AND MICROBIOLOGICAL CHARACTERISTICS AND OUTCOMES

I. Rua-Figueroa<sup>1</sup>, J. López-Longo<sup>2</sup>, M. Galindo-Izquierdo<sup>3</sup>, V. Del Campo<sup>4</sup>, J. De la Torre Cisneros<sup>5</sup>, E. Uriarte<sup>6</sup>, P. Vela<sup>7</sup>, E. Tomero<sup>8</sup>, J. Pego-Reigosa<sup>4</sup>. <sup>1</sup>Rheumatology, Hospital Dr. Negrin, Las Palmas GC; <sup>2</sup>Gregorio Marañón Hospital; <sup>3</sup>Doce de Octubre Hospital, Madrid (Spain), Madrid; <sup>4</sup>Hospital Complex of Vigo and IISGS, Vigo; <sup>5</sup>Reina Sofía Hospital, Córdoba; <sup>6</sup>Donosti Hospital, Guipúzcoa; <sup>7</sup>Alicante G Hospital, Alicante; <sup>8</sup>La Princesa Hospital, Madrid, Spain

**Background:** According to RELESSER (Spanish Society of Rheumatology Lupus Registry) data, bacteremia is the main cause of death by infection in systemic lupus erythematosus (SLE). However, the available information about this severe infection in SLE patients remains scarce.

**Methods:** Retrospective, nested case-control study of SLE patients (ACR-97 criteria) with at least one bacteremic episode and random controls from RELESSER Registry. Descriptive, bivariate and multivariate analysis (logistic regression)

**Results:** 114 bacteremic episodes in 83 patients were recorded. Incidence rate: 2.7/ 1,000 patient-years (n total: 3658). At the time of the bacteremia: median age: 40.5 (8–90) years, 88.6% female, disease duration: 9.7 (IR16.7), median SELENA-SLEDAI: 4 (IR8), 66% with severe flare (SFI criteria), active nephritis: 16.7%, median SLICC/ACR DI: 3 (IR4), any comorbidity: 64% (McCabe-Jackson criteria: 28.1% rapidly or ultimately fatal), more frequently renal failure (15.8%) or diabetes (11.4%). SLE treatment at the time of bacteraemia: 88.6% corticosteroids (68.6% >10mg/day), 57% immunosuppressors (mycophenolate 17.5% and cyclophosphamide 12.3%), 27% antimalarials. 44.7% suffered invasive procedures, more frequently intravascular catheter (24.6%). The bacteremia was nosocomial in 35.1% and the source was more frequently urinary (27.2%). 64% developed systemic inflammatory response syndrome and 35% needed intensive care unit admission, with multiorgan failure in 22.8%. The most frequent microorganism was *E. coli* (29.8%) followed by *Staphylococcus aureus* (16.7%) (22% methicillin-resistant) and *Salmonella spp.* (10.5%). 16% of the gram-negative enteric bacilli were extended-spectrum  $\beta$ -lactamase positive. 17.5% were multidrug resistant. 68.4% started the antibiotherapy before blood culture results, resulting finally active in susceptibility testing in 56 cases (71.8%), indicating an appropriate empirical antibiotic therapy in 49%. The bacteremia-related mortality was 14%. The risk of death was higher in patients with severe sepsis (Pitt index >8) (OR: 13 (IC95%: 3.71–45.17)). The bacteremia was recurrent in 26.3%. Associations with bacteremia in bivariate analysis (114 bacteremias vs 688 controls) are shown in Table 1. Antimalarials were protective. In the multivariate analysis (adjusted for disease duration), only elevated creatinine (OR 1.31 (95% CI 1.01–1.70), p=0.045), diabetes (OR 6.01 (95% CI 2.26–15.95), p=0.000), cancer (OR 5.32 (95% CI 2.23–12.70), p=0.000), immunosuppressors (OR 6.35 (95% CI 3.42–11.77), p=0.000), cyclophosphamide (OR 9.37 (95% CI 5.12–17.14), p=0.000) and SLICC/ACR DI (OR 1.65 (95% CI 1.31–2.09), p=0.000) remained statistically significant.

**Conclusions:** Bacteremia occurred mostly in active SLE, frequently in the context of a severe flare. Gram negative bacilli predominated, with high rate of multidrug resistance. The empiric treatment was inappropriate in a half of the cases. The

**Abstract OP0041** – Table 1. Overall and cause-specific mortality in SLE patients compared to general population; HR, hazard ratio

Cohorts		All-Cause Deaths		CVD Deaths		Malignancy Deaths		Other Causes	
		Deaths	HR (95% CI)	Deaths	HR (95% CI)	Deaths	HR (95% CI)	Deaths	HR (95% CI)
Overall Cohort	SLE (n=4238)	411	1.29 (1.15–1.46)	104	1.43 (1.15–1.79)	95	0.80 (0.63–1.00)	212	1.74 (1.46–2.09)
	Non-SLE (n=42380)	2226	1	622	1	795	1	809	1
Female Cohort	SLE (n=3643)	323	1.34 (1.17–1.54)	75	1.40 (1.08–1.81)	77	0.81 (0.62–1.05)	171	2.05 (1.68–2.50)
	Non-SLE (n=36430)	1697	1	469	1	618	1	610	1
Male Cohort	SLE (n=595)	88	1.06 (0.81–1.39)	29	1.73 (1.10–2.71)	18	0.76 (0.44–1.30)	41	1.06 (0.70–1.62)
	Non-SLE (n=5950)	529	1	153	1	177	1	199	1
Early Cohort (1997–2004)	SLE (n=1678)	98	1.20 (0.96–1.51)	25	1.18 (0.76–1.82)	22	0.68 (0.42–1.09)	51	1.86 (1.33–2.60)
	Non-SLE (n=16780)	508	1	163	1	173	1	172	1
Recent Cohort (2005–2012)	SLE (n=2560)	137	1.13 (0.92–1.39)	25	0.89 (0.57–1.39)	33	0.66 (0.45–0.98)	79	1.90 (1.42–2.56)
	Non-SLE (n=25600)	622	1	179	1	260	1	223	1