### Supplement to:

# Use of SLICC criteria in a large, diverse lupus registry enables SLE classification of a subset of ACR-designated incomplete lupus subjects

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#### Supplementary Methods. SLE classification of study subjects.

Extensive effort was made to obtain complete medical records for all patients enrolled in the LFRR, particularly for clinical encounters that would be relevant to SLE classification, such as rheumatology, dermatology, and hematology records. Rheumatology records were available for a large majority of subjects in this study. Records were reviewed for SLE classification criteria by a rheumatologist, rheumatology-trained physician assistant, or rheumatology-trained nurse. Medical record review was standardized using an extensive and detailed form that weighs and scores individual criteria documented by the care provider as well as additional clinical observations, laboratory results, and patient complaints. Recording the progression of these various forms of evidence over time (in most cases, multiple years of medical records) provides a clinical picture for each patient.

To supplement the medical records, blood samples were collected at the time of enrollment and tested for autoantibodies in the College of American Pathologists-certified Clinical Immunology Laboratory at OMRF (see "Methods: Autoantibody detection"). For classification purposes, subjects were considered autoantibody positive if a positive result was obtained by in-house testing or recorded in the medical record.

Organ System	SLE <sup>SLICC-only(1)</sup>	SLE <sup>ACR-only(2)</sup>	SLE <sup>both(3)</sup>	
SLICC criteria/ sub-criteria	(n=178)	(n=85)	(n=3,312)	(n=291)
Mucocutaneous, n (%)				
Acute/Subacute Cutaneous Ras Chronic Cutaneous Rashes	hes 105 (59.0)	74 (87.1)	2868 (86.6)	158 (54.3)
Oral/ Nasal Ulcers Alopecia		P<0.0001	P<0.0001	P=0.320
Musculoskeletal, n (%)	67 (37.6)	46 (54.1)	2344 (70.8)	131 (45.0)
Arthritis		P=0.012	P<0.0001	P=0.117
Pulmonary, n (%)	5 (2.8)	5 (5.9)	966 (29.2)	11 (3.8)
Pleuritis		P=0.233	P<0.0001	P=0.575
Cardiovascular, n (%)	6 (3.4)	5 (5.9)	576 (17.4)	11 (3.8)
Pericarditis		P=0.347	P<0.0001	P=0.818
Renal, n (%)	23 (12.9)	9 (10.6)	1262 (38.1)	13 (4.5)
Renal		P=0.589	P<0.0001	P=0.001
Neurologic, n (%)	22 (12 A)	5 (5.9)	585 (17.7)	4 (1.4)
Neurologic	22 (12.4)	P=0.113	P=0.071	P<0.0001
Hematologic, n (%)				
Anemia	06 (40 2)	28 (32.9)	2458 (74.2)	72 (24.7)
Leukopenia/Lymphopenia Thrombocytopenia	86 (48.3)	P=0.019	P<0.0001	P<0.0001

Supplementary Table 1. Organ system involvement in SLE and ILE patients.

<sup>1</sup>SLE<sup>SLICC-only</sup> were classified with SLE by SLICC criteria, but not ACR criteria. <sup>2</sup>SLE<sup>ACR-only</sup> were classified with SLE by ACR criteria, but not SLICC criteria. <sup>3</sup>SLE<sup>both</sup> were classified with SLE by both SLICC and ACR criteria. <sup>4</sup>ILE patients met 3 ACR criteria and were not classified with SLE by SLICC criteria. Bold P-values are significant (P<0.05) compared to SLE<sup>SLICC-only</sup>.

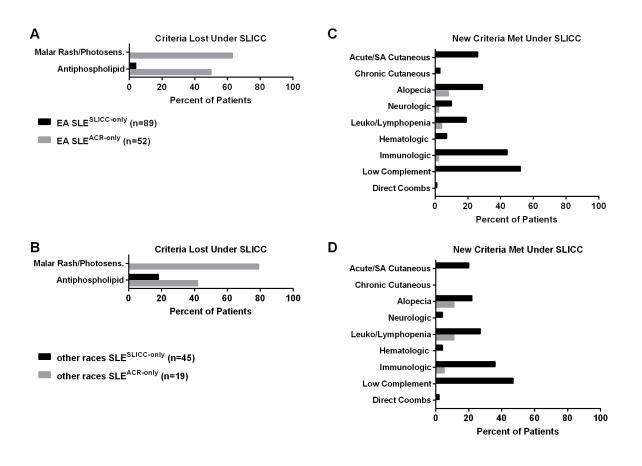
SLICC Criteria	ACR 1997 Criteria		
<u>Clinical</u> Acute/subacute cutaneous lupus malar rash, photosensitive lupus	Malar rash		
rash, and other lupus rashes Chronic cutaneous lupus	Discoid rash		
discoid rash and other lupus rashes	Photosensitivity		
Oral or nasal ulcers Non-scarring alopecia	Oral or nasal ulcers		
Arthritis	Arthritis		
Serositis	Serositis		
Renal disorder	Renal disorder		
Neurologic	Neurologic		
seizures, psychosis, mononeuritis multiplex, myelitis, peripheral or cranial neuropathy, or acute confusional state	seizures or psychosis		
Hemolytic anemia	Hematologic		
Leukopenia or lymphopenia	hemolytic anemia, leukopenia,		
Thrombcytopenia	lymphopenia, or thrombcytopenia		
mmunologic	Immunologic		
Anti-dsDNA	anti-dsDNA, anti-Sm, or		
Anti-Sm	antiphospholipid (abnormal		
Antiphospholipid medium- or high-titer anticardiolipin IgA, IgG, or IgM Low complement	anti-cardiolipin IgG or IgM)		

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**Direct Coombs' test** 

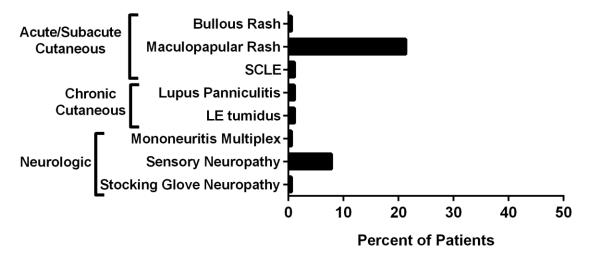
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Supplementary Figure 1. Comparison of SLE classification criteria under 2012 SLICC and **1997 ACR criteria.** SLICC criteria (left) classify subjects with SLE if they meet ≥4 criteria, including at least one clinical and one immunologic criterion, or if they exhibit biopsy-proven lupus nephritis with positive ANA or anti-dsDNA.<sup>1</sup> The 1997 ACR criteria (right) classify subjects with SLE if they meet  $\geq$ 4 criteria.<sup>2</sup> Criteria are listed in bold type. Sub-criteria that fulfill the given criterion are in italics, when SLICC and ACR sub-criteria differ. Comparable SLICC and ACR criteria are aligned in the table. Patients may gain criteria under SLICC criteria compared to ACR criteria by the addition of new criteria (alopecia, low complement, positive Direct Coombs) or new sub-criteria (acute cutaneous, chronic cutaneous, neurologic), less stringent thresholds for leukopenia/lymphopenia, and/or the separation of hematologic and immunologic sub-criteria into separate main criteria. Patients may lose criteria under SLICC criteria compared to ACR criteria by the combination of malar rash and photosensitivity into a single SLICC criterion, and/or the more stringent threshold for anti-phospholipid positivity. <sup>1</sup>Petri M, Orbai A-M, Alarcón GS, et al. Derivation and validation of the Systemic Lupus International Collaborating Clinics classification criteria for systemic lupus erythematosus. Arthritis Rheum 2012;64(8):2677-86. <sup> $^{2}$ </sup>Hochberg MC. Updating the American College of Rheumatology revised criteria for the classification of systemic lupus erythematosus. Arthritis Rheum 1997;40(9):1725.



**Supplementary Figure 2. Criteria contributing to SLE classification under SLICC criteria vary by race.** (A-B) Reasons for loss of criteria under SLICC criteria compared to ACR criteria were evaluated in SLE<sup>SLICC-only</sup> (black) and SLE<sup>ACR-only</sup> (gray) patients of European American (EA; A) and other races (B). (C-D) New criteria met under SLICC criteria compared to ACR criteria were evaluated in SLE<sup>SLICC-only</sup> (black) and SLE<sup>ACR-only</sup> (gray) patients of EA (C) and other races (D). SA=subacute

## SLICC Sub-Criteria in SLE<sup>SLICC-only</sup> Subjects



Supplementary Figure 3. Inclusion of maculopapular rash and sensory neuropathy as SLICC sub-criteria allows SLE classification of patients who do not meet ACR classification standards. Graph shows the percent of SLE<sup>SLICC-only</sup> patients positive for new SLICC sub-criteria. One additional SLE<sup>SLICC-only</sup> patient (not shown) failed to meet the more stringent anti-cardiolipin threshold under SLICC, but met the SLICC anti-phospholipid criterion by anti-β2-glycoprotein positivity.