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SYSTEMIC LUPUS ERYTHEMATOSUS AND STATINS IN GLADEL 2.0: ARE CARDIOVASCULAR RISK PREVENTION GUIDELINES BEING FOLLOWED?

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BACKGROUND

- Patients with systemic lupus erythematosus (SLE) have a significantly increased risk of atherosclerotic cardiovascular disease (ASCVD).
- European Alliance of Associations for Rheumatology (EULAR) recommends a comprehensive assessment and management of cardiovascular risk (CVR), following general population guidelines¹.
- Statin therapy plays a key role in reducing CVR and preventing ASCVD events².

METHODS

- This was an observational and cross-sectional study of GLADEL 2.0, a multi-ethnic Latin-American SLE cohort.
 - » Demographics, comorbidities, medications, disease activity, and laboratory data were analyzed.
- Statin eligibility was determined using the 2019 American College of Cardiology/American Heart Association (ACC/AHA) and the 2021 European Society of Cardiology (ESC) guidelines.
- CVR was assessed using the ASCVD risk calculator from the ACC/AHA, Systemic Coronary Risk Evaluation 2 (SCORE2), and Pan American Health Organization (PAHO) risk scores.
- Cohen's Kappa coefficient was used to determine inter-guideline agreement.
- Both guidelines only consider patients ≥40 years of age as candidates for CVR prevention with statins, due to eligibility dependence on CVR scores.
 - » Therefore, a comparative analysis was conducted between patients above and below this age threshold.

TABLE 1.
Baseline clinical characteristics, disease activity, damage index and treatments.

| | TOTAL (N=1083) |
|--|-------------------|
| Age, years, median (Q1, Q3) | 35.4 (15.6, 77.5) |
| ≥40 years, n (%) | 394 (36.4) |
| Female, n (%) | 970 (89.6) |
| Mestizo, n (%) | 701 (64.7) |
| Obesity, n (%) | 187 (18.5) |
| Disease duration, years, median (Q1, Q3) | 5.6 (0.6, 61.6) |
| SLEDAI, median (Q1, Q3) | 5 (0, 57) |
| SDI, median (Q1, Q3) | 0 (0, 9) |
| HDL-c (mg/dL), median (Q1, Q3), n=504 | 49 (11, 102) |
| LDL-c (mg/dL), median (Q1, Q3), n=480 | 103 (28, 328) |
| Triglycerides (mg/dL), median (Q1, Q3), n= 557 | 124 (32, 619) |
| Diabetes, n (%) | 45 (4.2) |
| Hypertension, n (%) | 293 (27.2) |
| Dyslipidemia, n (%) | 129 (12) |
| Active smoking, n (%) | 54 (5.1) |
| Chronic kidney disease, n (%) | 61 (5.6) |
| Acute myocardial infarction, n (%) | 8 (0.7) |
| Statins, n (%) | 155 (20.4) |
| Prednisone, n (%) | 758 (72.5) |
| Hydroxychloroquine, n (%) | 876 (84) |

HDL-c=high-density lipoprotein cholesterol, LDL-c=low-density lipoprotein cholesterol, SLEDAI=Systemic Lupus Erythematosus Disease Activity Index, SDI=Systemic Lupus International Collaborating Clinics/American College of Rheumatology Damage Index.

TABLE 2.
Comparison of clinical characteristics based on their age group.

| | < 40 YEARS | ≥ 40 YEARS | p VALUE |
|--|-----------------|-----------------|----------------------|
| Obesity, n (%) | 102 (15.8) | 85 (23.3) | 0.0031 ^a |
| Disease duration, years, median (Q1, Q3) | 4.1 (0.2, 28.9) | 9.9 (0.6, 61.6) | <0.0001 ^b |
| SLEDAI, median (Q1, Q3) | 6 (0, 57) | 3 (0, 38) | <0.0001 ^b |
| HDL-c, mg/dL, median (Q1, Q3), n=504 | 47 (11, 102) | 52 (15, 100) | 0.0003 ^b |
| LDL-c, mg/dL, median (Q1, Q3), n=480 | 105.5 (37, 241) | 101 (28, 328) | 0.1404 ^b |
| Triglycerides, mg/dL, median (Q1, Q3), n=557 | 123 (39, 619) | 127 (32, 525) | 0.4044 ^b |
| Diabetes, n (%) | 12 (1.7) | 33 (8.4) | <0.0001 ^a |
| Active smoking, n (%) | 30 (4.5) | 24 (6.2) | 0.0503 ^a |
| Statins, n (%) | 94 (19.8) | 61 (21.3) | 0.0510 ^a |
| Prednisone, n (%) | 525 (78.5) | 233 (61.8) | <0.0001 ² |

^aChi-Square p-value. ^bKruskal-Wallis p-value. HDL-c=high-density lipoprotein cholesterol, LDL-c=low-density lipoprotein cholesterol, SLEDAI=Systemic Lupus Erythematosus Disease Activity Index.

OBJECTIVE

- This study aimed to determine the proportion of patients with SLE eligible for statin use for primary ASCVD prevention, based on American and European CVR guidelines.

RESULTS

- Among 1083 patients in the GLADEL 2.0 SLE cohort, only 394 (36.4%) were older than 40 years of age (Table 1).
- ACC/AHA could only be calculated for 164 patients, PAHO for 351 patients, and SCORE2 for 181 patients.
- Most of these patients were categorized as having a low CVR, regardless of the calculator used.
- A total of 15 patients were indicated to receive statin therapy based on European guidelines; among these, only 5 (33%) had a previous prescription for a statin.
- Among 50 patients eligible for statin therapy based on American guidelines, only 13 (26%) had been prescribed them.
- Inter-guideline agreement on statin eligibility was fair (Cohen's Kappa = 0.35; 95% CI: 0.15–0.55).
- Patients under 40 years of age were more obese and had higher disease activity compared with those older than 40 years of age (Table 2).

CONCLUSIONS

- This study reveals a gap in the management of CVR among patients with SLE in the GLADEL 2.0 cohort, as only a small percentage are candidates for statin therapy, primarily due to the predominance of younger patients under age 40.
- » Traditional CVR assessment tools fail to encompass this group, which is at an elevated risk for accelerated atherosclerosis.
- Among eligible patients, statin prescribing rates remain low, suggesting a missed opportunity in proactive cardiovascular prevention.
- The moderate agreement between differing guidelines highlights the inconsistency in risk assessment and management approaches for this population.
- There is a need for tailored strategies to comprehensively evaluate and address CVR in younger patients with chronic inflammatory conditions like SLE.
- Implementing more inclusive guidelines could enhance preventative measures and improve outcomes for patients at increased risk for ASCVD.

REFERENCES

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- ²Yousef Yengej FA, et al. *Neth J Med.* 75(3):99–105.

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