Title: Age-Stratified Thirty-Day Rehospitalization and Mortality and Predictors of Rehospitalization among Patients with Systemic Lupus Erythematosus: a Medicare cohort study

Complete author names and ORCID iDs:

Maria Schletzbaum, 0000-0002-9009-3620 Amy J Kind, 0000-0002-7183-610X Yi Chen, 0000-0002-1817-5306 Brad C Astor, 0000-0002-0876-0069 Stacy P Ardoin, 0000-0003-4239-3634 Andrea Gilmore-Bykovskyi, 0000-0003-4930-3558 Ann M Sheehy, 0000-0001-6856-4687 Farah A Kaiksow, 0000-0002-1625-2187 W Ryan Powell, 0000-0001-6340-1520 Christie M Bartels, 0000-0001-6523-0374

Funding sources: Primary source of funding was National Institute on Minority Health and Health Disparities Research (NIMHD) award number R01MD010243. Additional support from NIMHD award number F30MD015211, the University of Wisconsin Medical Scientist Training Program (grant number T32 GM140935), and the University of Wisconsin Clinical and Translational Science Award (NIH CTSA grant 1UL1TR002373).

Authors: Maria Schletzbaum^{1,2}, Amy J. Kind MD PhD^{2,3}, Yi Chen MS⁴, Brad C. Astor PhD MPH^{1,2}, Stacy P. Ardoin MD MS⁵, Andrea Gilmore-Bykovskyi PhD RN^{3,6}, Ann M. Sheehy MD MS^{2,3}, Farah A. Kaiksow MD MPP^{2,3}, W. Ryan Powell PhD MA^{2,3}, Christie M. Bartels MD MS^{2,3}

Affiliations:

¹University of Wisconsin School of Medicine and Public Health, Department of Population Health Sciences, Madison, WI, US

²University of Wisconsin School of Medicine and Public Health, Department of Medicine, Madison, WI, US

³Center for Health Disparities Research, University of Wisconsin School of Medicine and Public Health, Madison, WI, US

⁴University of Wisconsin School of Medicine and Public Health, Department of Biostatistics and Medical Informatics, Madison, WI, US

⁵Division of Pediatric Rheumatology, Nationwide Children's Hospital, Columbus, OH, US

⁶University of Wisconsin – Madison, School of Nursing, Madison, WI, US

Conflicts of Interests:

CMB receives peer-reviewed institutional grant funding from Independent Grants for Learning and Change (Pfizer) for research unrelated to this study. All other authors declare no conflicts.

Corresponding author:

Christie M Bartels, MD MS 1685 Highland Ave #4132 Madison, WI 53705 Phone: (608) 262-8305 cb4@medicine.wisc.edu

Key Indexing Terms: Systemic Lupus Erythematosus, Hospital Readmission,

Medicare, Age Group, Health Services Research, Cohort Study

Word Count: Abstract: 250 Body text: 3,495

ABSTRACT

Objective

Recent studies suggest young adults with lupus have high 30-day readmission rates, which may necessitate tailored readmission reduction strategies. To aid in risk-stratification for future strategies, we measured 30-day rehospitalization and mortality among Medicare beneficiaries with lupus and determined rehospitalization predictors by age.

Methods

In a 2014 20% national Medicare sample of hospitalizations, rehospitalization risk and mortality within 30 days of discharge were calculated for young (18-35), middle-aged (36-64), and older (65+) beneficiaries with and without lupus. Multivariable GEE models were used to predict rehospitalization among lupus patients by age group using patient, hospital, and geographic factors.

Results

Among 1.39 million Medicare hospitalizations, 10,868 involved lupus beneficiaries. Hospitalized young adult lupus beneficiaries were more racially diverse, living in more disadvantaged areas, and had more comorbidities than older lupus and non-lupus beneficiaries. Thirty-day rehospitalization was 36% among young adult lupus beneficiaries – 40% higher than peers without lupus and 85% higher than older (65+) lupus beneficiaries. Longer length of stay and higher comorbidity risk score increased odds of rehospitalization in all age groups, while specific comorbid condition predictors

and their impact varied. Our models, which incorporated neighborhood-level socioeconomic disadvantage, had moderate-to-good predictive value (c-statistics 0.67-0.77), outperforming administrative data models lacking comprehensive social determinants in other conditions.

Conclusions

Young adults with lupus on Medicare had very high 30-day rehospitalization at 36%. Considering socioeconomic disadvantage and comorbidities provided good prediction of rehospitalization risk, particularly in young adults. Young lupus beneficiaries with comorbidities should be a focus of programs aimed at reducing rehospitalizations.

Patients with systemic lupus erythematosus (SLE or lupus) experience high rates of hospital readmission (24-27%) ^{1,2}. Medicare covers over a third of US lupus hospitalizations ^{2,3}, and more than half of Medicare lupus hospitalizations occur in patients under 65, who qualified for Medicare due to disability or end stage renal disease (ESRD) ². In 2012, the Centers for Medicare & Medicaid Services (CMS) implemented a program that financially penalizes hospitals for unplanned readmissions for six conditions as an indicator of poor inpatient care quality ⁴. While readmissions related to SLE are not included in the program, rehospitalization within 30 days represents an important care quality marker. We previously reported higher 30-day readmissions in young adults compared to older age groups (27% vs 17%), yet drivers of readmission in young adults remain unclear ².

Many young adults with SLE were diagnosed before age 18. Making up 20% of lupus cases in the US ⁵⁻⁷, early-onset versus adult-onset patients are more likely to identify as Black, Hispanic, Asian, or American Indian ^{5,8}, have lower socioeconomic status ⁷, and have greater organ damage ⁸. Young adults with lupus also use emergency care more frequently than older SLE patients ⁹. Lower socioeconomic status and greater disease activity further increase odds of emergency visits ⁹. Disproportionately high acute care use and rehospitalizations might indicate sub-optimal outpatient management, leading to increased healthcare costs and poor outcomes ^{10,11}. In other conditions, risk-stratification to target high risk patients has helped reduce rehospitalizations and healthcare costs ¹²⁻¹⁶.

Our first objective was to assess rates of 30-day rehospitalization and mortality in young

adults with SLE compared to non-SLE and older SLE Medicare beneficiaries. Then, to risk stratify, our second objective was to identify predictors of 30-day rehospitalization among young adults with SLE compared to older SLE beneficiaries.

METHODS

Study population

We performed a cohort study utilizing a geo-linked 20% random sample of Medicare beneficiaries. Hospitalizations of adult beneficiaries between January 1st and November 30th, 2014 were eligible for inclusion; a beneficiary could contribute multiple hospitalizations. Inclusion required hospitalization with a lupus diagnosis code (ICD-9-CM 710.0) and discharge before November 30, 2014 (Figure 1). At least 12 months of continuous Medicare A and B coverage with any claims before index hospitalization were required for baseline comorbidity assessment. Beneficiaries lacking at least 30 days of A/B coverage after discharge were excluded unless they died in the 30 days. Due to alternative claims processing, beneficiaries with Health Maintenance Organization plans or Railroad worker benefits were excluded as their Medicare data may be incomplete. Consistent with standard rehospitalization metrics, long-term acute care facility, psychiatric, rehabilitation, cancer, children's hospitals, and drug treatment hospitalizations were excluded 10. Beneficiaries were excluded from regression analysis if they were missing geographical residence.

This study was approved by the Health Sciences Minimal Risk Institutional Review

Board at the University of Wisconsin School of Medicine and Public Health with a waiver

of individual informed consent as a secondary analysis of administrative claims data

(study ID: 2020-0438).

SLE definition

Patients were defined as having SLE if the hospitalization was associated with an International Classification of Diseases, Ninth Revision, diagnosis code for SLE (ICD-9 710.0) [specificity of 99.4%] at any diagnosis level ^{2,17,18}.

Outcomes

The primary outcome for this study was all-cause readmission to a hospital setting within thirty days of discharge ¹⁹. All-cause thirty-day mortality (defined using the National Death Index) was evaluated as a secondary outcome.

Predictors

Potential 30-day readmission predictors included both individual and contextual factors to capture broad influences on rehospitalization ^{11,15,20}: demographic and socioeconomic status (SES) variables, geographic area, comorbidities, and characteristics of the index hospital. Age was measured at index hospitalization. Age groups were young (18-35), middle-aged (36-64), and older adults (65+). Medicare-reported race and ethnicity were included as a proxy for lived experiences of these groups, including structural and institutional racism. For analysis, race and ethnicity responses were consolidated to Asian, Black, American Indian, Hispanic, White, and Other/Unknown using the Research Triangle Institute (RTI) variable ²¹⁻²⁴. Medicaid status ²⁵ was also included. The Area Deprivation Index (ADI) rank score was used to capture neighborhood-level socioeconomic disadvantage based on nine-digit ZIP code ^{26,27}. The ADI includes 17 variables, reflecting census block-level income, education,

employment, and housing quality. To account for geographical context, patients were classified into isolated, small rural, large rural, and urban areas based on Rural-Urban Commuting Area (RUCA) codes by ZIP code ^{28,29}.

The following health status variables were included: disability status, CMS's hierarchical condition category community risk score (HCC) ³⁰, Elixhauser Comorbidity Index conditions ³¹, and length of the index hospital stay. Disability was determined by original qualification for Medicare due to disability, regardless of subsequent eligibility (i.e., ESRD, age 65) ²⁵. The length of stay for the index hospitalization was included as a proxy for medical complexity.

Characteristics of the index treating hospital were evaluated, including affiliation with a medical school, tertile of discharge volume, and critical access hospital status.

Analysis

Descriptive summaries of patient and hospital characteristics for hospitalizations of beneficiaries with SLE and without SLE by age group are provided. The non-SLE group was composed of hospitalizations that were not associated with a SLE diagnosis code from the 20% random Medicare sample in the same period. This non-SLE group was used for comparison of observed 30-day rehospitalization and mortality risk but was not used in rehospitalization prediction models.

Observed risk of 30-day rehospitalization and 30-day mortality

Rehospitalization and mortality risk within 30 days of index hospitalization discharge with Clopper-Pearson 95% confidence intervals were calculated by age group among

SLE and non-SLE hospitalizations.

Predictors of 30-day rehospitalization among SLE hospitalizations by beneficiary age group

We used generalized estimating equation (GEE) models with logistic link functions to predict 30-day odds of rehospitalization, clustering by beneficiary to account for multiple hospitalizations. *A priori* variables were selected based on the literature for their association with acute care use among SLE patients; these variables were age, sex, race and ethnicity, disability, length of stay, ADI national rank, HCC community risk score, and renal failure status ^{2,9,18,32-35}. Other potential predictors, based on theoretical models of rehospitalization, were eligible for selection by LASSO, which maximizes a model's predictive power while minimizing factors in the model for higher efficiency ³⁶.

Two modeling approaches were used. In the first, LASSO variable selection was performed across all SLE associated hospitalizations, regardless of beneficiary age. These selected predictors, along with the *a priori* variables, were combined into one model to predict 30-day rehospitalization and applied separately to young, middle-aged, and older SLE patient groups. This approach allows for quantitative comparisons of odds ratios across age groups. In the second approach, 30-day rehospitalization predictors were selected separately within each beneficiary age group and then combined with the *a priori* variables, resulting in a unique set of predictors (and model) for each age group. This approach allows for a qualitative comparison of relevant predictors between SLE patient age groups. Model performance was assessed by c-statistic.

Sensitivity analysis for obstetrical hospitalizations

Given the age and female predominance of the cohort, sensitivity analyses were performed to determine whether obstetrics-related hospitalizations influenced the findings. First, 30-day rehospitalization rates were calculated and compared for obstetrics-related index hospitalizations, based on having a primary Clinical Classification Software diagnostic code of 11.* (excluding 11.1.* - contraceptive care), and non-obstetrical hospitalizations among SLE and non-SLE beneficiary hospitalizations. Second, LASSO variable selection and GEE models on the SLE cohort were re-performed excluding obstetrical hospitalizations. LASSO-selected variables and significant predictors of rehospitalization were compared for models including and excluding obstetrics-related hospitalizations.

Analysis was conducted using SAS version 9.4 (SAS Institute, Cary, NC).

RESULTS

There were 10,868 SLE associated Medicare hospitalizations in the sample (Figure 1). Beneficiaries with SLE were significantly younger and more likely to be Black or Hispanic than beneficiaries without SLE (Table 1). Disability, receipt of Medicaid, and multiple comorbid conditions were more frequent among young adults with SLE than among older adults with SLE or among the age matched non-SLE Medicare population. Young adults with SLE tended to have a higher illness burden, as measured by HCC, and to live in more disadvantaged neighborhoods relative to both older SLE beneficiaries and those without SLE.

Accepted Articl

Observed 30-day rehospitalization and mortality risk

Unadjusted 30-day rehospitalization risk was highest in young adult Medicare beneficiaries. In young adults, rehospitalization risk was 10% higher among SLE beneficiaries compared to non-SLE beneficiaries (36% vs. 26%; p<.001). Adults 65+ with SLE had a rehospitalization risk of 20% (Figure 2A). Thus, the rehospitalization risk for young adults with SLE was 40% higher than their peers and 85% higher than the risk for older adults with SLE.

The unadjusted 30-day mortality risk for young adults with and without SLE was similar at 0.5% and 0.7% (Figure 2B). Mortality risk increased with age in SLE and non-SLE beneficiaries, but was statistically lower for those with SLE compared to beneficiaries without SLE in the middle-aged and older age groups.

Rehospitalization predictors across all ages

When examining predictors across all ages of SLE beneficiaries, the variables selected by the LASSO procedure, after accounting for *a priori* variables, were all comorbid conditions: coagulopathy, congestive heart failure, drug use disorder, fluid and electrolyte disorders, and paralysis (Table 2).

When this model was applied separately to each SLE age group, length of index hospitalization and HCC risk score were significant predictors of rehospitalization in all ages. Longer index hospitalization showed a similar increase in odds in each age group (aOR ~1.03). Higher HCC risk scores were associated with higher odds of rehospitalization, with greater impact on middle-aged and older adults (young adult aOR 1.10, 95% CI 1.03, 1.18; middle-aged 1.18, 1.14-1.23; older adults 1.21, 1.17-1.26).

Within the young and middle-aged adult strata, but not among older adults, younger beneficiaries were at higher risk of rehospitalization. Each additional year of age was associated with a 6.4% decrease in odds of rehospitalization among young adults and a 2.1% decrease among middle-aged adults.

In young adults, fluid and electrolyte disorders (aOR 2.35; 95% CI 1.53, 3.61), congestive heart failure (2.05; 1.44, 2.91), drug use disorder (1.58; 1.10, 2.27), and coagulopathy (1.46; 1.03, 2.06) substantially increased the odds of rehospitalization. Paralysis was associated with lower rehospitalization odds (0.23; 0.07, 0.78). The impact of these comorbid conditions generally decreased in the older age groups. In middle-aged adults, only drug use disorder (1.73; 1.31, 2.27) and fluid and electrolyte disorders (1.26; 1.05, 1.52) remained significant. Among older adults, congestive heart failure (1.32; 1.08, 1.61) was the only significant predictor beyond length of stay and HCC risk score.

The c-statistic for the model was 0.766 for young adults, 0.703 in middle-aged adults, and 0.673 in older adults.

Age-specific rehospitalization predictors

With the sample restricted to young adults with SLE, more variables were added by the LASSO procedure as predictors and included index hospital characteristics (index hospital medical school affiliation and discharge volume) and additional comorbid condition indicators (deficiency anemia, diabetes, hypertension with complications, and valvular disease) (Table 3). With the addition of more comorbid condition variables, coagulopathy became non-significant (1.36; 0.97, 1.90), while valvular disease was

associated with increased odds (1.74; 1.14, 2.65). Index hospitalization at a medical school affiliated hospital was significantly protective for rehospitalization (0.68; 0.50, 0.92).

When LASSO variable selection was performed among middle-aged beneficiaries with SLE, additional variables selected as predictors included critical access status of the index hospital along with alcohol use disorder, liver disease, pulmonary circulation disorders, and weight loss (Table 3). Coagulopathy, congestive heart failure, and paralysis were no longer selected for the model. Drug use disorder (1.67; 1.23, 2.18) and liver disease (1.32; 1.03, 1.68) conferred additional risk for rehospitalization, while hospitalization in a critical access hospital (0.30; 0.10, 0.95) and weight loss (0.71; 0.55, 0.92) were associated with decreased odds.

Among older adults with SLE, only three variables were selected as predictors by the LASSO procedure: congestive heart failure, other neurological conditions, and paralysis. Congestive heart failure (1.35; 1.11, 1.64) and other neurological conditions (1.26; 1.01, 1.58) were significantly associated with increased odds of rehospitalization, while paralysis trended towards decreased odds (0.58; 0.33, 1.00).

The c-statistic for the young adult model was 0.775, 0.706 for the middle-aged adult model, and 0.672 for the older adults model.

Sensitivity analysis for obstetrical hospitalizations

Obstetrics-related hospitalizations made up 0.52% of SLE-related hospitalizations compared to 0.19% of non-SLE hospitalizations. While rehospitalization rates were much lower for obstetric hospitalizations among young adults without SLE (9.8% vs 27.6% in non-obstetric; p<0.0001), rehospitalization rates for obstetric and non-obstetric hospitalizations among young adults with SLE were similarly high (31.7% vs 36.5%; p=0.53). When obstetric hospitalizations were excluded, rehospitalization rates showed very minimal change and there were no changes in the significant predictors of rehospitalization.

DISCUSSION

In this study, we found the 30-day rehospitalization risk among young adult Medicare beneficiaries with SLE to be 36%; 40% higher than young adult beneficiaries without SLE, and 85% higher than SLE beneficiaries 65 or older. Young adult beneficiaries with SLE were more racially diverse, from more disadvantaged neighborhoods, and had a higher burden of comorbidities than middle-aged and older adult SLE beneficiaries. Our models had moderate-to-good predictive ability for 30-day rehospitalization among SLE beneficiaries using geo-linked administrative data, demonstrating value for riskstratification. Medical school affiliation of the index hospital was protective for readmissions among young adults with lupus in this Medicare sample. This may indicate positive effects of programs or policies at medical school affiliated hospitals and potential for wider dissemination efforts. To our knowledge, this is among the first studies to evaluate readmissions for young adults with lupus. It adds to findings from multi-payor SLE cohorts that have reported associations of younger age with greater Emergency Department use ⁹ and direct healthcare costs ³⁷. Our findings underscore the high rehospitalization risk among young adult Medicare beneficiaries with SLE and

suggest the need to include young adults with SLE in targeted efforts to reduce rehospitalization.

In Kangovi and Grande's framework for readmissions, rehospitalization is not only a result of inpatient care quality and patient health status, but also outpatient care quality, access to care, and patient socioeconomic resources ¹¹. Uniquely, our models incorporated rich information on the social determinants of health: patient's neighborhood disadvantage (comprised of 17 indicators of socioeconomic resources), Medicaid status, and rurality. We found that the highest risk group, young adult beneficiaries with SLE, lived in more disadvantaged socioeconomic contexts, with a median neighborhood disadvantage index 16 percentile points higher than the U.S. median, and 91% qualified for Medicaid. Young adult Medicare beneficiaries with SLE were also highly diverse at 52% Black and 15% Hispanic. Lower access to care for Black and Hispanic Americans compared to White Americans is well documented, and prior research has shown these groups may also receive lower quality lupus care ^{38,39}.

An inverse relationship between age and rehospitalization rates has been observed in other early-onset chronic conditions, such as Type 1 Diabetes ⁴⁰. Young adults with SLE may disproportionately experience additional barriers to outpatient services (e.g., childcare, transportation), potentially contributing to their higher rates of rehospitalization. In the general population, those aged 18-39 were 10% more likely than adults 55 or older to report non-financial barriers leading to delayed or unmet care ⁴¹.

To obtain Medicare coverage individuals under age 65 must have end stage renal

disease or have qualified for Social Security disability benefits ⁴² for ≥24 months.. The five-step qualification process begins by confirming income <\$1,070/month. Young adults with SLE additionally had higher prevalence of many significant comorbid conditions than any other SLE age group (Table 1). In addition to HCC risk score and renal failure status, several other comorbidities potentially related to SLE pathophysiology (coagulopathy, congestive heart failure, fluid and electrolyte disorders, and paralysis) were selected for model inclusion. Notably, renal failure was not associated with odds of rehospitalization. Fluid and electrolyte disorders may have captured the sequalae of renal failure that predict rehospitalization. While obstetrical hospitalizations had significantly lower rehospitalization rates among beneficiaries without SLE, obstetrical hospitalizations of lupus beneficiaries still had high levels of rehospitalization. This suggests readmission reduction efforts should extend to obstetrical care for patients with lupus.

Among young and middle-aged adults, drug use disorder was associated with increased rehospitalization odds. Patients with lupus often need chronic pain management as part of their SLE treatment and are 2 to 3 times more likely to be taking prescribed opioids than non-SLE patients, with greater prevalence of previous opioid-related encounters among young adults with lupus ^{43,44}. Thus, these findings raise concerns about the contribution of substance use disorders, particularly in relation to prescribed opioids, to rehospitalization.

Among young adults with SLE, index hospitalization at a medical school-affiliated hospital was associated with lower odds of rehospitalization. This aligns with previous

literature showing higher quality of care and lower 30-day post-discharge mortality for major teaching hospitals compared to non-academic hospitals for many common inpatient conditions ^{45,46}. Our finding prompts interest in identifying practices at academic centers that may reduce rehospitalization for potential dissemination.

Our models outperformed prior U.S. population-based studies on health condition-specific readmission risk prediction using retrospective administrative data which reported moderately predictive models (c-statistic range of 0.55 to 0.65) ¹⁵. Our method, combining *a priori* and LASSO-selected variables including a neighborhood-level social determinants of health index, performed better (c-statistic range of 0.67 to 0.77) than those in the literature. These results suggest the value of incorporating location-based metrics of SES, such as the ADI, in readmission risk prediction models. Such metrics are now freely accessible and can be integrated into both retrospective administrative and real-time electronic health record data ^{27,47}. Notably, while the fitting of age-specific models revealed some unique significant predictors in each age group, it had little effect on overall predictive value compared to the model developed with patients of all ages. The presented lupus-specific prediction models can aid healthcare and insurance systems in deciding where to focus resources to reduce rehospitalizations by risk-stratifying among the high-risk SLE patient population.

While there are many strengths of this study, including a large national cohort of lupus patients, we acknowledge some limitations. Since this study utilized claims data, we do not have some patient health measures such as functional status or current SLE disease activity. Individuals must meet additional requirements to receive Medicare

benefits under age 65. Thus, results under age 65 may not generalize to the US lupus population, particularly patients with better health status, lower prevalence of ESRD, or those unable to navigate disability qualification. However, one-third of individuals with SLE in the U.S. have public insurance and one-half of SLE-related hospitalizations and Emergency Department visits are covered by Medicare or Medicaid ^{10,48,49}, making this cohort relevant to practice and policy. Higher rates of ED use among young adult lupus patients in a multi-payor lupus cohort 9 suggest that the higher risk of acute healthcare use in young adult lupus patients is not unique to Medicare beneficiaries. Poor health status of Medicare beneficiaries also underscores the importance of developing interventions for this group. Reported race and ethnicity in Medicare data have known issues with misclassification ^{22,23}. While we used the more valid RTI measure, approximately 4% of patients may be misclassified. Native American and Asian patients are more likely to be misclassified ²², which could affect comparisons involving these groups. As administrative data were used in this study, conditions and billing codes may not have been uniformly coded for all beneficiaries. However, previously validated, published algorithms for administrative data were used to define SLE and other comorbidities ^{18,31}. Coding bias may also have been reduced since all individuals in the cohort had the same insurance coverage.

While Medicare covers many lupus patients with SLE, multi-payor or Medicaid cohorts should be evaluated to compare rates and validate the rehospitalization predictors among young adults, including more patients without ESRD. Further research comparing rehospitalization reasons should be performed among young adults with SLE. Additional evaluations are needed of socioeconomic resource and access barriers

for young adults with SLE and their relation to rehospitalization, ambulatory versus acute care use, and healthcare costs. For Medicare lupus-related hospitalizations, the administrative data-based predictive models had moderate-to-good predictive value and could be applied to prioritize lupus patients for readmission interventions. These areas of inquiry can be leveraged to inform program and policy development to reduce rehospitalization and improve care for young adults with SLE.

Young adult Medicare beneficiaries with SLE were 40% more likely to be rehospitalized within 30 days compared to age matched non-SLE beneficiaries and 85% more likely than older SLE beneficiaries. This elevated rehospitalization rate among young adults with SLE in Medicare may be partially explained by higher neighborhood disadvantage and greater prevalence among Black and Hispanic patients, who generally have lower access to healthcare and receive lower quality lupus care than White patients ^{38,39}. Young adult beneficiaries with SLE also had more comorbidities which increased their odds of rehospitalization. The presented lupus-specific prediction models had moderate-to-good prediction for 30-day rehospitalization, informing risk-stratification. Together, these findings suggest a critical need to develop targeted interventions, alongside young adults with SLE, to provide greater outpatient support, address disease management barriers, and reduce costly rehospitalizations.

ACKNOWLEDGMENTS

Authors would like to thank Monica Messina for manuscript formatting.

REFERENCES

- Elixhauser A, Steiner C. Readmissions to U.S. Hospitals by Diagnosis, 2010: Statistical Brief #153. Healthcare Cost and Utilization Project (HCUP) Statistical Briefs. Rockville (MD)2006.
- 2. Bartels CM, Chodara A, Chen Y, et al. One Quarter of Medicare Hospitalizations in Patients with Systemic Lupus Erythematosus Readmitted within Thirty Days. Semin Arthritis Rheum 2021;51:477-85.
- 3. Ward MM. Hospital experience and mortality in patients with systemic lupus erythematosus: which patients benefit most from treatment at highly experienced hospitals? J Rheumatol 2002;29:1198-206.
- 4. Centers for Medicare & Medicaid Services. Hospital Readmissions Reduction Program (HRRP) Archives. [Internet. Accessed October 1, 2019.] Available from:

 https://www.cms.gov/Medicare/Medicare-Fee-for-Service-Payment/AcuteInpatientPPS/HRRP-Archives.
- 5. Pons-Estel GJ, Alarcon GS, Scofield L, Reinlib L, Cooper GS. Understanding the epidemiology and progression of systemic lupus erythematosus. Semin Arthritis Rheum 2010;39:257-68.
- 6. Ardoin SP, Daly RP, Merzoug L, et al. Research priorities in childhood-onset lupus: results of a multidisciplinary prioritization exercise. Pediatr Rheumatol Online J 2019;17:32.
- 7. Levy DM, Kamphuis S. Systemic lupus erythematosus in children and adolescents. Pediatr Clin North Am 2012;59:345-64.
- 8. Tucker LB, Uribe AG, Fernandez M, et al. Adolescent onset of lupus results in more aggressive disease and worse outcomes: results of a nested matched case-control study within LUMINA, a multiethnic US cohort (LUMINA LVII). Lupus 2008;17:314-22.
- 9. Panopalis P, Gillis JZ, Yazdany J, et al. Frequent use of the emergency department among persons with systemic lupus erythematosus. Arthritis Care Res (Hoboken) 2010;62:401-8.
- 10. Jencks SF, Williams MV, Coleman EA. Rehospitalizations among patients in the Medicare fee-for-service program. N Engl J Med 2009;360:1418-28.
- 11. Kangovi S, Grande D. Hospital readmissions--not just a measure of quality. JAMA 2011;306:1796-7.
- 12. Halfon P, Eggli Y, Pretre-Rohrbach I, Meylan D, Marazzi A, Burnand B. Validation of the potentially avoidable hospital readmission rate as a routine indicator of the quality of hospital care. Medical Care 2006;44:972-81.
- 13. Amarasingham R, Moore BJ, Tabak YP, et al. An Automated Model to Identify Heart Failure Patients at Risk for 30-Day Readmission or Death Using Electronic Medical Record Data. Medical Care 2010;48:981-8.
- 14. Hansen LO, Young RS, Hinami K, Leung A, Williams MV. Interventions to reduce 30-day rehospitalization: a systematic review. Ann Intern Med 2011;155:520-8.
- 15. Kansagara D, Englander H, Salanitro A, et al. Risk prediction models for hospital readmission: a systematic review. JAMA 2011;306:1688-98.
- 16. Kripalani S, Theobald CN, Anctil B, Vasilevskis EE. Reducing hospital readmission rates: current strategies and future directions. Annu Rev Med 2014;65:471-85.
- 17. Hanly JG, Thompson K, Skedgel C. Identification of patients with systemic lupus erythematosus in administrative healthcare databases. Lupus 2014;23:1377-82.
- 18. Yazdany J, Marafino BJ, Dean ML, et al. Thirty-day hospital readmissions in systemic lupus erythematosus: predictors and hospital- and state-level variation. Arthritis Rheumatol 2014;66:2828-36.

This accepted article is protected by copyright. All rights reserved.

Accepted Articl

- 19. Centers for Medicare & Medicaid Services. 2015 Measure Information about the 30-day All-Cause Hospital Readmission Measure. [Internet. Accessed February 5, 2021.]

 Available from: https://www.cms.gov/Medicare/Medicare-Fee-for-Service-Payment/PhysicianFeedbackProgram/Downloads/2015-ACR-MIF.pdf.
- 20. Joynt KE, Jha AK. A path forward on Medicare readmissions. N Engl J Med 2013;368:1175-7.
- 21. Eicheldinger C, Bonito A. More accurate racial and ethnic codes for Medicare administrative data. Health Care Financ Rev 2008;29:27-42.
- 22. Grafova IB, Jarrin OF. Beyond Black and White: Mapping Misclassification of Medicare Beneficiaries Race and Ethnicity. Med Care Res Rev 2020:1077558720935733.
- 23. Jarrin OF, Nyandege AN, Grafova IB, Dong X, Lin H. Validity of Race and Ethnicity Codes in Medicare Administrative Data Compared With Gold-standard Self-reported Race Collected During Routine Home Health Care Visits. Med Care 2020;58:e1-e8.
- 24. Bonito AJBC, Eicheldinger C, L C. Creation of new race-ethnicity codes and socioeconomic status (SES) indicators for Medicare beneficiaries. Final report, sub-task 2. Report No.: AHRQ Publication No. 08-0029-EF. Rockville, MD: 2008.
- 25. Centers for Medicare & Medicaid Services, Research Data Assistance Center. Original Reason for Entitlement Code. [Internet. Accessed February 25, 2021.] Available from: https://resdac.org/cms-data/variables/original-reason-entitlement-code.
- 26. Singh GK. Area deprivation and widening inequalities in US mortality, 1969-1998. Am J Public Health 2003;93:1137-43.
- 27. Kind AJ, Jencks S, Brock J, et al. Neighborhood socioeconomic disadvantage and 30-day rehospitalization: a retrospective cohort study. Ann Intern Med 2014;161:765-74.
- 28. Washington State Department of Health. Guidelines for Using Rural-Urban Classification Systems for Community Health Assessment. 2016. [Internet. Accessed May 14, 2021.] Available from: https://www.doh.wa.gov/Portals/1/Documents/1500/RUCAGuide.pdf.
- 29. U.S. Department of Agriculture Economic Research Service. Rural Classifications: Overview. 2021. [Internet. Accessed October 5, 2021.] Available from: https://www.ers.usda.gov/topics/rural-economy-population/rural-classifications/.
- 30. Pope GC, Kautter J, Ellis RP, et al. Risk adjustment of Medicare capitation payments using the CMS-HCC model. Health Care Financ Rev 2004;25:119-41.
- 31. Elixhauser A, Steiner C, Harris DR, Coffey RM. Comorbidity measures for use with administrative data. Med Care 1998;36:8-27.
- 32. Feldman CH, Xu C, Williams J, Collins JE, Costenbader KH. Patterns and predictors of recurrent acute care use among Medicaid beneficiaries with systemic lupus erythematosus. Seminars in Arthritis and Rheumatism 2020;50:1428-36.
- 33. Chan K, Dekis A, Clarke AE, et al. Hospitalizations in patients with systemic lupus erythematosus: updated analyses from 2006 to 2011. Arthritis Research & Therapy 2012;14:A59.
- 34. Ward MM. Avoidable hospitalizations in patients with systemic lupus erythematosus. Arthritis Rheum 2008;59:162-8.
- 35. Sule S, Fivush B, Neu A, Furth S. Increased hospitalizations and death in patients with ESRD secondary to lupus. Lupus 2012;21:1208-13.
- 36. Tibshirani R. Regression shrinkage and selection via the Lasso. J Roy Stat Soc B Met 1996;58:267-88.
- 37. Panopalis P, Yazdany J, Gillis JZ, et al. Health care costs and costs associated with changes in work productivity among persons with systemic lupus erythematosus. Arthritis Rheum 2008;59:1788-95.
- 38. Pryor KP, Barbhaiya M, Costenbader KH, Feldman CH. Disparities in Lupus and Lupus Nephritis Care and Outcomes Among US Medicaid Beneficiaries. Rheum Dis Clin North Am 2021;47:41-53.

- 39. Yazdany J, Trupin L, Tonner C, et al. Quality of care in systemic lupus erythematosus: application of quality measures to understand gaps in care. J Gen Intern Med 2012;27:1326-33.
- 40. Everett E, Mathioudakis NN. Association of socioeconomic status and DKA readmission in adults with type 1 diabetes: analysis of the US National Readmission Database. BMJ Open Diabetes Res Care 2019;7:e000621.
- 41. Kullgren JT, McLaughlin CG, Mitra N, Armstrong K. Nonfinancial Barriers and Access to Care for U.S. Adults. Health Services Research 2012;47:462-85.
- 42. US Congressional Research Service Report. Primer on Disability Benefits: Social Security Disability Insurance (SSDI) and Supplemental Security Income (SSI) [Internet. Accessed March 25, 2022.] Available from: https://sgp.fas.org/crs/misc/RL32279.pdf.
- 43. Somers EC, Lee J, Hassett AL, et al. Prescription Opioid Use in Patients With and Without Systemic Lupus Erythematosus Michigan Lupus Epidemiology and Surveillance Program, 2014-2015. MMWR Morb Mortal Wkly Rep 2019;68:819-24.
- 44. Bartels C, Schletzbaum M, Chen Y, Kind A. Opioid-Related Encounters as a Predictor of 30-Day Readmissions in Lupus [abstract]. Arthritis Rheumatol 2020;72 (suppl 10).
- 45. Ayanian JZ, Weissman JS. Teaching hospitals and quality of care: a review of the literature. Milbank Q 2002;80:569-93, v.
- 46. Burke LG, Frakt AB, Khullar D, Orav EJ, Jha AK. Association Between Teaching Status and Mortality in US Hospitals. JAMA 2017;317:2105-13.
- 47. Kind AJH, Buckingham WR. Making Neighborhood-Disadvantage Metrics Accessible The Neighborhood Atlas. N Engl J Med 2018;378:2456-8.
- 48. Lim SS, Bayakly AR, Helmick CG, Gordon C, Easley KA, Drenkard C. The incidence and prevalence of systemic lupus erythematosus, 2002-2004: The Georgia Lupus Registry. Arthritis Rheumatol 2014;66:357-68.
- 49. GfK Roper Public Affairs. Lupus Awareness Survey for the Lupus Foundation of America [Executive Summary October 2012]. [Internet. Accessed October 5, 2020.] Available from: https://b.3cdn.net/lupus/2489f6ca2bcbde1818_ggm6i6gzi.pdf.

This accepted article is protected by copyright. All rights reserved.

FIGURE LEGENDS

Figure 1. Flow diagram describing creation of study cohorts from a 20% national Medicare sample of index hospitalizations between January 1, 2014 and November 30, 2014.

Figure 2. Observed 30-day rehospitalization rates (panel A) and mortality rates (panel B) among SLE (black) and non-SLE (grey) Medicare beneficiaries by age category. Error bars represent 95% confidence intervals.

TABLE 1

Table 1. Characteristics of Hospitalizations of Medicare Beneficiaries

| Patient Characteristic | Non-SLE | SLE Age 18-35 | SLE Age 36-64 | SLE Age 65+ |
|------------------------------------|-----------------|-----------------|-----------------|-----------------|
| n (%) | n= 1,378,654 | n= 1,133 | n= 4,855 | n= 4,880 |
| Female Sex | 776,803 (56) | 1,037 (92) | 4,331 (89) | 4,355 (89) |
| Age (median, IQR) | 75.0, 67.2-83.2 | 29.8, 26.8-32.5 | 52.5, 45.4-59.0 | 73.4, 69.3-79.7 |
| Race/Ethnicity Asian | 14,920 (1) | 24 (2) | 52 (1) | 37 (1) |
| Black | 168,877 (12) | 586 (52) | 1,880 (39) | 687 (14) |
| Hispanic | 27,180 (2) | 170 (15) | 297 (6) | 71 (1) |
| American Indian | 10,068 (1) | 26 (2) | 64 (1) | 39 (1) |
| White | 1,378,654 (83) | 253 (22) | 2,493 (51) | 3,992 (82) |
| Other/Unknown | 19,022 (1) | 74 (7) | 69 (1) | 54 (1) |
| ADI National Rank (median, IQR) | 51, 28-74 | 66, 42-86 | 63, 39-83 | 50, 26-72 |
| Medical School Affiliated Hospital | 709,740 (52) | 734 (65) | 2,809 (58) | 2,424 (50) |
| Critical Access Hospital | 45,383 (3) | * | 51 (1) | 104 (2) |
| Discharge Volume: Highest Tertile | 467,995 (34) | 450 (40) | 2,023 (42) | 1,744 (36) |
| Middle Tertile | 454,667 (33) | 427 (38) | 1,614 (33) | 1,701 (35) |
| Lowest Tertile | 455,992 (33) | 256 (23) | 1,218 (25) | 1,435 (29) |
| Ever Received Medicaid | 411,694 (30) | 1,036 (91) | 2,912 (60) | 1,059 (22) |
| Disability on Enrollment | 440,957 (32) | 898 (79) | 4,590 (95) | 1,559 (32) |
| Length of Stay (median days, IQR) | 4, 2-6 | 4, 2-6 | 4, 2-6 | 4, 2-6 |
| HCC Risk Score (median, IQR) | 2.27, 1.26-3.89 | 3.92, 2.24-6.07 | 3.25, 1.90-5.32 | 2.8, 1.70-4.42 |
| Renal Failure | 308,476 (22) | 745 (66) | 1,839 (38) | 1,244 (25) |
| Coagulopathy | 94,073 (7) | 363 (32) | 830 (17) | 488 (10) |
| Congestive Heart Failure | 291,295 (21) | 345 (30) | 1,139 (23) | 1,134 (23) |
| Deficiency Anemia | 429,475 (31) | 900 (79) | 2,547 (52) | 1,917 (39) |
| Fluid and Electrolyte Disorders | 406,930 (30) | 794 (70) | 2,234 (46) | 1,674 (34) |
| Hypertension with Complications | 983,804 (71) | 895 (79) | 3,566 (73) | 3,601 (74) |
| Other Neurological Disorder | 223,999 (16) | 338 (30) | 1,104 (23) | 823 (17) |
| Paralysis | 50,851 (4) | 23 (2) | 196 (4) | 124 (3) |
| Pulmonary Circulation Disorder | 79,098 (6) | 209 (18) | 605 (12) | 446 (9) |
| Valvular Disease | 159,653 (12) | 145 (13) | 629 (13) | 678 (14) |
| Alcohol Use Disorder | 49,741 (4) | 36 (3) | 164 (3) | 22 (0) |
| Diabetes | 495,115 (36) | 209 (18) | 1,651 (34) | 1,403 (29) |
| Drug Use Disorder | 55,168 (4) | 308 (27) | 630 (13) | 78 (2) |
| Liver Disease | 64,096 (5) | 75 (7) | 441 (9) | 227 (5) |
| Weight Loss | 106,444 (8) | 210 (19) | 572 (12) | 436 (9) |

^{*}Suppressed due to small cell size. ADI = Area Deprivation Index. HCC Risk Score = Hierarchical condition category community risk score.

Accepted Articl

TABLE 2

Table 2. Odds Ratios for 30-Day Rehospitalization across SLE Age Strata with LASSO Selected Predictors from all SLE Beneficiaries

| | | SLE Age 18-35 | SLE Age 36-64 | SLE Age 65+ |
|---|-----------------|-------------------|-------------------|--------------------|
| | | n= 1,103 | n= 4,748 | n= 4,789 |
| Age at Index Admission (per year) | | 0.94 (0.90, 0.98) | 0.98 (0.97, 0.99) | 0.99 (0.98, 1.01) |
| Sex | Female | 1.16 (0.70, 1.93) | 0.90 (0.69, 1.17) | 0.887 (0.68, 1.16) |
| Race/Ethnicity | White | ref | ref | ref |
| | Asian | 0.42 (0.16, 1.12) | 0.75 (0.35, 1.57) | 0.81 (0.37, 1.78) |
| | Black | 0.79 (0.52, 1.20) | 1.13 (0.94, 1.36) | 1.07 (0.85, 1.35) |
| | Hispanic | 0.61 (0.36, 1.01) | 0.98 (0.71, 1.36) | 0.83 (0.41, 1.70) |
| | American Indian | 1.07 (0.48, 2.36) | 1.31 (0.76, 2.26) | 1.02 (0.55, 1.92) |
| | Other/Unknown | 0.89 (0.47, 1.67) | 0.67 (0.35, 1.25) | 1.17 (0.56, 2.46) |
| Ever received Medicaid | | 1.00 (0.58, 1.73) | 0.93 (0.78, 1.11) | 0.98 (0.79, 1.22) |
| ADI National Rank (per decile increase) | | 1.00 (0.95, 1.06) | 1.02 (0.99, 1.05) | 0.98 (0.95, 1.01) |
| Disability on Enrollment | | 0.93 (0.61, 1.42) | 0.94 (0.67, 1.31) | 0.98 (0.82, 1.18) |
| Length of Stay (per day) | | 1.04 (1.02, 1.06) | 1.04 (1.02, 1.05) | 1.03 (1.01, 1.04) |
| HCC Community Risk Score (per unit) | | 1.10 (1.03, 1.18) | 1.18 (1.14, 1.23) | 1.21 (1.17, 1.26) |
| Renal Failure | | 1.08 (0.74, 1.58) | 1.09 (0.89, 1.34) | 1.01 (0.84, 1.23) |
| Coagulopathy | | 1.46 (1.03, 2.06) | 1.09 (0.87, 1.37) | 1.15 (0.91, 1.45) |
| Congestive Heart Failure | | 2.05 (1.44, 2.91) | 1.15 (0.93, 1.42) | 1.32 (1.08, 1.61) |
| Fluid and Electrolyte Disorders | | 2.35 (1.53, 3.61) | 1.26 (1.05, 1.52) | 1.17 (0.97, 1.41) |
| Paralysis | | 0.23 (0.07, 0.78) | 0.75 (0.49, 1.15) | 0.61 (0.37, 1.02) |
| Drug Use Disorder | | 1.58 (1.10, 2.27) | 1.73 (1.31, 2.27) | 1.25 (0.75, 2.06) |
| C-statistic | | 0.766 | 0.703 | 0.673 |
| | | | | |

Variables below the horizontal line were selected by LASSO variable selection performed on all participants with SLE. ADI = Area Deprivation Index with higher values indicating greater disadvantage. HCC = Hierarchical condition category community risk score is scaled to 1 for average risk with higher values indicating greater comorbidities and healthcare utilization.

TABLE 3

Table 3. Odds Ratios for 30-Day Rehospitalization across SLE Age Groups using Age Strata-Specific LASSO Selected Predictors

| | SLE Age 18-35 | SLE Age 36-64 | SLE Age 65+ |
|---|-------------------|-------------------|-------------------|
| | n= 1,103 | n= 4,748 | n= 4,789 |
| Age at Index Admission (per year) | 0.93 (0.90, 0.96) | 0.98 (0.97, 0.99) | 1.00 (0.98, 1.01) |
| Sex Female | 1.13 (0.68, 1.89) | 0.93 (0.72, 1.19) | 0.89 (0.68, 1.17) |
| Race/Ethnicity White | ref | ref | ref |
| Asian | 0.29 (0.10, 0.81) | 0.76 (0.36, 1.59) | 0.86 (0.40, 1.86) |
| Black | 0.72 (0.48, 1.08) | 1.13 (0.94, 1.35) | 1.08 (0.85, 1.36) |
| Hispanic | 0.61 (0.36, 1.03) | 1.01 (0.72, 1.40) | 0.87 (0.43, 1.75) |
| American Indian | 1.08 (0.52, 2.22) | 1.19 (0.71, 2.02) | 1.02 (0.53, 1.94) |
| Other/Unknown | 0.87 (0.46, 1.65) | 0.67 (0.26, 1.23) | 1.19 (0.57, 2.51) |
| Ever received Medicaid | 0.88 (0.50, 1.54) | 0.93 (0.78, 1.11) | 0.98 (0.79, 1.21) |
| ADI National Rank (per decile increase) | 1.00 (0.95, 1.05) | 1.02 (0.99, 1.05) | 0.98 (0.95, 1.01) |
| Disability on Enrollment | 1.00 (0.69, 1.46) | 0.92 (0.66, 1.29) | 0.98 (0.81, 1.18) |
| Length of Stay (per day) | 1.03 (1.01, 1.05) | 1.04 (1.02, 1.05) | 1.03 (1.01, 1.04) |
| HCC Community Risk Score (per unit) | 1.09 (1.02, 1.17) | 1.19 (1.15, 1.23) | 1.22 (1.18, 1.27) |
| Renal Failure | 1.00 (0.69, 1.44) | 1.15 (0.95, 1.39) | 1.03 (0.85, 1.24) |
| Medical School Affiliated Hospital | 0.68 (0.50, 0.92) | _ | - |
| Discharge Volume: Highest Tertile | ref | _ | _ |
| Middle Tertile | 0.73 (0.47, 1.13) | - | - |
| Lowest Tertile | 0.95 (0.67, 1.36) | _ | _ |
| Critical Access Hospital | - | 0.30 (0.10, 0.95) | _ |
| Deficiency Anemia | 1.26 (0.76, 2.10) | - | - |
| Congestive Heart Failure | 2.02 (1.46, 2.81) | - | 1.35 (1.11, 1.64) |
| Coagulopathy | 1.36 (0.97, 1.90) | - | - |
| Hypertension with Complications | 1.05 (0.65, 1.68) | - | - |
| Fluid and Electrolyte Disorders | 2.30 (1.49, 3.56) | 1.29 (1.07, 1.55) | - |
| Other Neurological Conditions | - | - | 1.26 (1.01, 1.58) |
| Paralysis | 0.20 (0.06, 0.63) | - | 0.58 (0.33, 1.00) |
| Pulmonary Circulation Disorders | - | 1.27 (0.98, 1.63) | <u>-</u> |
| Valvular Disease | 1.74 (1.14, 2.65) | - | - |
| Alcohol Use Disorder | - | 1.51 (0.97, 2.36) | - |
| Diabetes | 1.37 (0.94, 1.99) | - | - |
| Drug Use Disorder | 1.57 (1.12, 2.21) | 1.67 (1.23, 2.18) | - |
| Liver Disease | - | 1.32 (1.03, 1.68) | - |
| Weight Loss | - | 0.71 (0.55, 0.92) | - |
| C-statistic | 0.775 | 0.706 | 0.672 |

Variables below the horizontal line were selected by LASSO variable selection procedure performed on the respective age stratum of beneficiaries with SLE. ADI = Area Deprivation Index with higher values indicating greater disadvantage. HCC = Hierarchical condition category community risk score is scaled to 1 for average risk with higher values indicating greater comorbidities and healthcare utilization.

Accepted Article

FIGURE 1

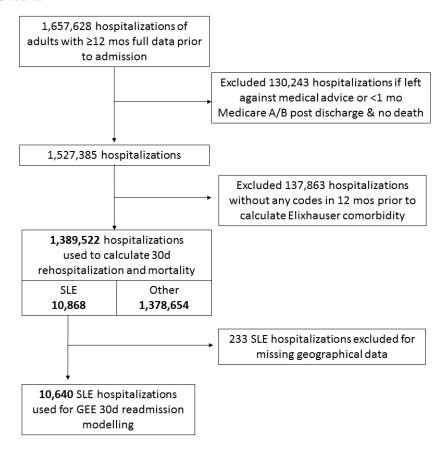


Figure 1. Flow diagram describing creation of study cohorts from a 20% national Medicare sample of index hospitalizations between January 1, 2014 and November 30, 2014.

254x254mm (96 x 96 DPI)

FIGURE 2

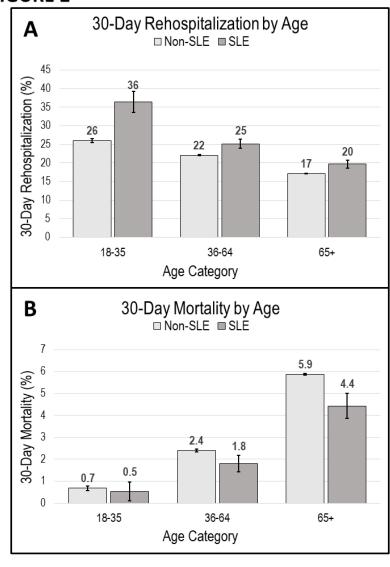


Figure 2. Observed 30-day rehospitalization rates (panel A) and mortality rates (panel B) among SLE (black) and non-SLE (grey) Medicare beneficiaries by age category. Error bars represent 95% confidence intervals.

203x279mm (96 x 96 DPI)