

## Increased 1-Year Healthcare Use in Survivors of Severe Sepsis

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## Abstract

**Rationale:** Hospitalizations for severe sepsis are common, and a growing number of patients survive to hospital discharge. Nonetheless, little is known about survivors' post-discharge healthcare use.

**Objectives:** To measure inpatient healthcare use of severe sepsis survivors compared with patients' own presepsis resource use and the resource use of survivors of otherwise similar nonsepsis hospitalizations.

**Methods:** This is an observational cohort study of survivors of severe sepsis and nonsepsis hospitalizations identified from participants in the Health and Retirement Study with linked Medicare claims, 1998–2005. We matched severe sepsis and nonsepsis hospitalizations by demographics, comorbidity burden, premorbid disability, hospitalization length, and intensive care use.

**Measurements and Main Results:** Using Medicare claims, we measured patients' use of inpatient facilities (hospitals, long-term acute care hospitals, and skilled nursing facilities) in the 2 years

surrounding hospitalization. Severe sepsis survivors spent more days (median, 16 [interquartile range, 3–45] vs. 7 [0–29]; P < 0.001) and a higher proportion of days alive (median, 9.6% [interquartile range, 1.4–33.8%] vs. 1.9% [0.0–7.9%]; P < 0.001) admitted to facilities in the year after hospitalization, compared with the year prior. The increase in facility-days was similar for nonsepsis hospitalizations. However, the severe sepsis cohort experienced greater post-discharge mortality (44.2% [95% confidence interval, 41.3–47.2%] vs. 31.4% [95% confidence interval, 28.6–34.2%] at 1 year), a steeper decline in days spent at home (difference-in-differences, -38.6 d [95% confidence interval, -50.9 to 26.3]; P < 0.001), and a greater increase in the proportion of days alive spent in a facility (difference-in-differences, 5.4% [95% confidence interval, 2.8–8.1%]; P < 0.001).

**Conclusions:** Healthcare use is markedly elevated after severe sepsis, and post-discharge management may be an opportunity to reduce resource use.

**Keywords:** patient outcomes assessment; hospitalization; patient readmission; skilled nursing facility; healthcare facilities

Each year, hundreds of thousands of patients in the United States are hospitalized with severe sepsis (1, 2), and a growing number survive to hospital discharge (2–4). Severe sepsis survivors experience increased mortality (5, 6), cognitive and functional decline (7–9), and decreased quality of life (10, 11) that persist for years after their acute illness. The growing population of severe sepsis survivors has been deemed a "hidden public health disaster" (12). Yet, despite rising attention to the long-term consequences of severe sepsis (9, 12), and hospitalization more generally (13–16), newly updated sepsis treatment guidelines have literally no mention of post-discharge care (17). The Agency for Healthcare Research and Utilization recently reported that the United States spends more money on hospitalizations for sepsis than for any other cause (18), yet the full impact of severe sepsis survivorship on the US healthcare system is unknown. A recent study of survivors of a less common acute illness (those requiring prolonged mechanical

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## At a Glance Commentary

## Scientific Knowledge on the

**Subject:** The existing literature on healthcare resource use after severe sepsis has focused on documenting the healthcare spending, and other indirect costs, incurred by patients surviving a severe sepsis hospitalization.

## What This Study Adds to the

Field: Our study adds to the literature in several important ways. First, we provide relevant comparisons with patients' own baseline resource use, and with the resource use of matched survivors of nonsepsis hospitalizations so that the resource use of severe sepsis survivors can be understood within the appropriate context. Second, rather than measuring use as Medicare spending or other abstract measures of cost, we measure the daily use of inpatient healthcare facilities in the 2 years surrounding hospitalization. We believe this outcome is of high interest to both patients and payers.

ventilation) revealed significant postdischarge care needs (19). Such granular information on the clinical course of severe sepsis survivors is not currently available, despite the larger population and budgetary impact of severe sepsis survivors.

With the growing prominence of readmissions as a subject of attention and potential financial penalties (13, 20), and concern that the burden of illness is shifting to the post-acute care setting (9, 21–23), it has become increasingly important to understand the post-discharge needs of severe sepsis survivors as a health system, beyond the compelling patient-centered reasons. Therefore, we analyzed Medicare claims from a nationally representative cohort of older Americans to quantify the inpatient healthcare use of severe sepsis survivors.

We compared the healthcare use of severe sepsis survivors with their own presepsis resource use to examine the extent to which changes are associated with severe sepsis, rather than simply patients' propensity to use care (24). By letting patients serve as their own control subjects, we distinguish use associated with severe sepsis from that of patients' baseline health status, demographic characteristics, and socioeconomic factors that influence healthcare consumption. We also examined the resource use of survivors of matched nonsepsis hospitalizations to understand whether the impact of severe sepsis on subsequent use is different from that of patients with otherwise similar hospitalizations for different reasons. We have previously reported some of these results in an abstract (25).

## Methods

### **Study Population**

The Health and Retirement Study (HRS) is an ongoing, nationally representative, prospective cohort study of older Americans (26). Started in 1992, the HRS has enrolled nearly 30,000 participants, of whom 16,772 have agreed to link their data with Medicare (26). The cohort is reinterviewed every 2 years, with a follow-up rate consistently over 90% (26).

Patients provided informed consent on enrollment in the HRS and again for linkage to Medicare. We screened all respondents with a baseline cognitive and physical function assessment (1998–2004) for whom there were claims-based data on a subsequent hospitalization (1998–2005). A similar cohort was identified previously for inclusion in a study of cognitive and functional disability among severe sepsis survivors (7). The University of Michigan institutional review board approved this research.

# Identification of Severe Sepsis and Nonsepsis Hospitalizations

We used a commonly employed and externally validated claims-based definition of severe sepsis that requires documentation of infection and acute organ dysfunction during a single hospitalization (1, 27). This definition has a specificity of 96.3% and positive predictive value of 70.7% (27). For patients with more than one severe sepsis hospitalization, each hospitalization was included. We also identified all hospitalizations without severe sepsis within HRS-Medicare (1998–2005). We excluded all hospitalizations with inpatient mortality.

## **Data Abstraction**

We abstracted characteristics of the index hospitalizations from Medicare Provider

Analysis and Review (MedPAR) files, including acute organ dysfunctions, critical care use, and mechanical ventilation use (for further details on abstraction of hospitalization characteristics see online supplement). We abstracted baseline deficiencies of activities of daily living (ADLs) and instrumental ADLs from the biennial HRS survey immediately preceding index hospitalization (median, 1.1 yr prior). We determined patients' baseline Charlson Comorbidity Index (28) score using a 1year look-back in Medicare claims. We determined enrollment in hospice from the Hospice Standard Analytical File.

### Matching of Severe Sepsis and Nonsepsis Hospitalizations

We used a coarsened exact matching approach, because this matching method often achieves the best possible covariate balance while retaining sample size (29). We matched severe sepsis and nonsepsis hospitalizations one-to-one, without replacement, on age, sex, Charlson Comorbidity Index score (28), premorbid limitations of the six ADLs and five instrumental ADLs, length of hospitalization, and intensive care use. For the dichotomous variables, sex and intensive care use, we matched exactly. For Charlson Comorbidity Index score, we matched on whether patients had 0, 1, or 2+ comorbidities. For premorbid limitations, we matched on whether patients had 0, 1, or 2+ total deficiencies of ADLs and instrumental ADLs. For length of hospitalization, we matched on lengths of 1, 2, 3, 4, 5, 6, 7, and 8 to 30, 31 to 60, or 61+ days. For age, we used the default automatic binning algorithm in the user-written "cem" command for Stata (StataCorp, College Station, TX) (30).

### Outcomes

Our outcome of interest was use of three types of inpatient healthcare facilities: (1) short-term acute care hospitals (hospitals), (2) long-term acute care hospitals (LTACs) (31), and (3) skilled nursing facilities (SNFs) (32) (definitions provided in Table E1 in online supplement). To measure use, we determined the daily location of each patient for the 365 days preceding index hospitalization and the 365 days after index hospital discharge (or until death) using MedPAR files. Date of death was determined from National Death Index, and confirmed in Medicare denominator files and/or HRS exit interviews.

After determining the daily location of each patient in the 2 years surrounding index hospitalization, we measured the number of days spent at home, in each of the three types of inpatient facilities, and dead. We also determined the number of hospital readmissions and transitions of care locations. We did not include the index hospitalization or transition at index discharge in our use counts.

We considered days at home, days in inpatient facilities, and days dead to be mutually exclusive categories, according to the hierarchy: dead, hospital, LTAC, SNF, home, such that each patient's total days of use could not exceed their days of observation. In contrast, readmission and transition of care counts measured all use. For example, if a patient was admitted from a SNF to a hospital and died the same day, this day would be coded as "dead" (not in a hospital or SNF), but would count as both a readmission and a transition of care.

We assumed that patients were at home for any day they were known to be alive and not admitted to an inpatient facility based on MedPAR records. Because Medicare does not pay for long-term custodial care provided by nursing facilities, patients residing in a nursing home on a permanent basis cannot be identified on a daily basis through MedPAR records (33, 34). Thus, although our results accurately measure healthcare use through Medicare, they likely underestimate total service use.

## **Statistical Analysis**

We present patient characteristics as numbers (percentages), means (SDs), or medians (interquartile ranges [IQR]) as appropriate. We report mortality as means (95% confidence interval [CI]), use outcomes as medians (IQR), and rates per person-year. To compare use between and within cohorts, we used nonparametric tests to account for the skewed distribution of the outcomes (*see* online supplement). To compare changes in use between the severe sepsis and nonsepsis cohorts, we performed difference-in-differences analyses (35).

We conducted all analyses with Stata software version 13 (StataCorp). We used hospitalization as the unit of analysis, unless otherwise specified. We used two-sided significance testing and considered a *P* value less than 0.05 to be significant.

#### **Additional Analyses**

We performed a subgroup analysis of the patients who used intensive care during their hospitalization. Also, because of potential temporal changes in in-hospital mortality, SNF use, and LTAC use, we performed a sensitivity analysis in which we matched severe sepsis and nonsepsis hospitalizations by year of hospital admission, in addition to the six other matching variables.

## Results

### Severe Sepsis and Nonsepsis Cohorts

We identified 1,162 severe sepsis and 23,803 nonsepsis hospitalizations without inpatient mortality (Figure 1). After matching hospitalizations one-to-one on age, sex, premorbid functional limitations, Charlson Comorbidity Index, length of hospitalization, and use of intensive care, we included 1,083 severe sepsis hospitalizations (93.2% of those identified) and 1,083 nonsepsis hospitalizations in the analysis.

In general, the cohorts were older (mean age, 78.6  $\pm$  8.6 yr), predominantly female (53.5%), with a moderate comorbidity burden (mean Charlson Comorbidity Index score, 2.1  $\pm$  1.7), and mild to moderate baseline functional limitations (median total limitations of baseline ADL and instrumental ADLs, 1; IQR, 0–5) (Table 1). Median hospital length of stay was 8 days (IQR, 5–13 d) and 420 patients (38.8%) in each cohort used intensive care during their hospitalization. As expected, the severe sepsis and nonsepsis cohorts did not differ by any of the six matched characteristics.

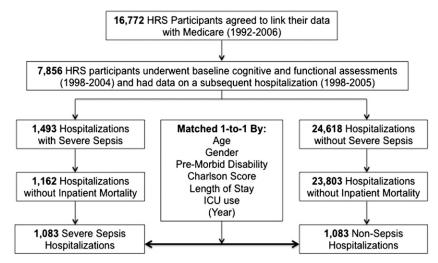
The sepsis cohort experienced a mean of  $1.1 \pm 0.4$  acute organ dysfunctions. Of the 1,083 severe sepsis hospitalizations, 452 (41.7%) had renal dysfunction, 252 (23.3%) had cardiovascular dysfunction, 211 (19.5%) had hematologic dysfunction, 210 (19.4%) had respiratory dysfunction, 104 (9.6%) had neurologic dysfunction, and 7 (0.7%) had hepatic dysfunction.

The 77 (6.6%) unmatched severe sepsis hospitalizations were younger, had longer hospitalizations, and were more likely to use intensive care than the severe sepsis hospitalizations included in the analysis (*see* Table E2). Compared with the matched hospitalization, the unmatched severe sepsis hospitalizations had similar presepsis use, but greater post-sepsis use and greater mortality.

# Mortality, Place of Death, and Hospice Use

The severe sepsis cohort experienced 27.5% 90-day mortality (95% CI, 24.8–30.2%) and 44.2% 1-year mortality (95% CI, 41.3–47.2%), markedly higher than the nonsepsis cohort who experienced 15.5% 90-day (95% CI, 13.4–17.7%) and 31.4% 1-year mortality (95% CI, 28.6–34.2%) (P < 0.01 for each time-point).

Of the 479 severe sepsis patients who died in the year after hospitalization, 231 (48.2%) died at home, 162 (33.8%) died in



**Figure 1.** Identification and matching of severe sepsis and nonsepsis cohorts. We measured premorbid disability as the number of limitations in activities and instrumental activities of daily living. HRS = Health and Retirement Study.

**Table 1.** Demographic Information and Hospitalization Characteristics for Matched

 Survivors of Severe Sepsis and Nonsepsis Hospitalizations

Demographic or Hospitalization Characteristic	Severe Sepsis Cohort (n = 1,083)	Nonsepsis Cohort (n = 1,083)	P Value
Matched characteristics Age, mean (SD) Male, n (%) Charlson score, mean (SD) Premorbid physical disability, median (IQR)	78.5 (8.6) 503 (46.5) 2.1 (1.7)	78.6 (8.6) 503 (46.5) 2.2 (1.8)	0.91 1.00 0.93
Basic ADL limitations Instrumental ADL limitations Total ADL limitations Hospital length of stay (d), median (IQR)	1 (0–3) 0 (0–2) 1 (0–5) 8 (5–14)	1 (0–3) 0 (0–2) 1 (0–5) 8 (5–12)	0.53 0.08 0.27 0.11
Used an intensive care unit, n (%) Unmatched characteristics	420 (38.8)	420 (38.8)	1.00
Self-reported race/ethnicity, n (%) White Black/African American Other Premorbid cognitive function, n (%)	841 (77.7) 219 (20.2) 23 (2.1)	860 (79.4) 198 (18.3) 24 (2.2)	0.51
Normal Mild Impairment Moderate/severe impairment Received mechanical ventilation, n (%)	911 (84.1) 89 (8.2) 83 (7.7) 210 (19.4)	945 (87.3) 54 (5.0) 84 (7.8) 29 (2.7)	<0.01

Definition of abbreviations: ADL = activity of daily living; IQR = interquartile range.

a hospital, and 86 (18.0%) died in a SNF or LTAC. One hundred sixty-two (33.8%) enrolled in hospice before their death. For nonsepsis patients who died in the year after hospitalization, death location and proportion enrolled in hospice were similar to the severe sepsis cohort (P > 0.05 for each).

## Healthcare Use in the Year after Severe Sepsis

The 1,083 severe sepsis survivors required substantial healthcare resources in the year after hospitalization (*see* Table E3). On average, they spent a median 16 days (IQR, 2–45) or 9.6% (IQR, 1.4–33.8%) of their days alive in the year after discharge admitted to an inpatient healthcare facility. Mean use is presented in Table E4.

Two hundred eighty-seven (26.5%) were readmitted within 30 days of hospital discharge, 444 (41.0%) were readmitted within 90 days, and 682 (63.0%) were readmitted at some point in the first year. Three hundred sixty-one (33.3%) were readmitted more than once during the year. Only 222 (20.5%) of the severe sepsis survivors remained alive for 1 year after discharge without being readmitted to a hospital. Visual inspection of patients' daily use in the year after severe sepsis hospitalization suggests that much of the increased healthcare use occurs in the first 90 days (Figure 2B). Indeed, severe sepsis survivors spent a median 9 (IQR, 0–28) days admitted to a healthcare facility in the first 90 days after hospital discharge, compared with a median 0 days (IQR, 0–13) in days 91–365 (P < 0.001). Severe sepsis survivors were hospitalized at a rate of 2.94 hospitalization per patient-year in the first 90 days, compared with 1.54 hospitalization per patient-year in the 91–365 days after hospital discharge.

We present use counts and rates for specific components of care in Tables E3–E5.

# Healthcare Use before and after Severe Sepsis

The healthcare use of the severe sepsis survivors surpassed their own baseline presepsis healthcare use. Patients spent more days (median, 16 [IQR, 3–45] vs. 7 [0–29]; P < 0.001) and a markedly higher proportion of their days alive (median, 9.6% [IQR, 1.4–33.8%] vs. 1.9% [0.0–7.9%]; P < 0.001) admitted to an inpatient healthcare facility in the year after

compared with the year before severe sepsis hospitalization. The rate of hospitalization increased from 1.44 hospitalizations per patient-year before severe sepsis to 1.95 hospitalizations per patient-year after severe sepsis. The rate of inpatient healthcare use increased from 24.2 days per patient-year presevere sepsis to 47.9 days per patient-year post-severe sepsis.

### Changes in Healthcare Use after Severe Sepsis versus Nonsepsis Hospitalizations: A Difference-in-Differences Analysis

The change in facility-days was similar in the severe sepsis and nonsepsis cohorts (Figure 3, Table 2). However, as a result of the increased post-discharge mortality experienced by the severe sepsis cohort, severe sepsis survivors had a steeper decline in the number of days spent at home (difference-in-differences, -38.6 d [95% CI, -50.9 to 26.3]; P < 0.001) and a steeper rise in the proportion of days alive spent admitted to an inpatient healthcare facility (difference-in-differences, 5.4% [95% CI, 2.8-8.1%]; P < 0.001).

# Subgroup Analysis: Hospitalizations with Intensive Care Use

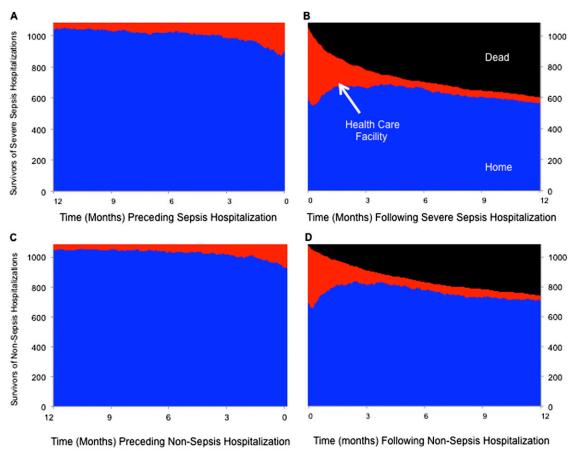
The 420 (38.8%) severe sepsis hospitalizations that used intensive care had increases in use of a similar magnitude to the whole cohort (*see* Table E6). The differencein-differences analysis for the 420 matched severe sepsis and nonsepsis hospitalizations was similar to the difference-in-differences analysis for the full cohorts (Table 2).

### Sensitivity Analysis: Matching on Year

We were able to match 795 (68.4%) of the severe sepsis hospitalizations to nonsepsis hospitalizations by year of admission, in addition to the six other matching variables. The results of pre versus post and difference-in-differences analyses for the year-matched cohorts were not substantially different from the primary analysis (Table 2; *see* Table E7).

## Discussion

Using a nationally representative cohort of older adults, we have shown that the inpatient healthcare facility use of severe sepsis survivors is markedly elevated compared with patients' own presepsis healthcare use. In the year after a severe



**Figure 2.** Use and mortality for 1,083 matched survivors in the 2 years surrounding severe sepsis and nonsepsis hospitalizations. This figure displays the daily location of 1,083 matched survivors of severe sepsis and nonsepsis hospitalizations in the 1 year before (*A* and *C*) and 1 year after hospitalization (*B* and *D*). The index admission is not included. The year prior ends at the day before hospitalization; the year after begins the day after hospital discharge. Patients are depicted as being at home (*blue*), admitted to a healthcare facility (*red*), or dead (*black*).

sepsis hospitalization, patients spent a median of 10% (mean, 25%) of their days alive admitted to an inpatient healthcare facility (a hospital, LTAC, or SNF). Most patients were rehospitalized, many more than once. Only one in five severe sepsis survivors remained alive for a full year after severe sepsis without being rehospitalized.

When compared with survivors of nonsepsis hospitalizations, severe sepsis survivors had similar increases in use, but greater mortality, a steeper decline in days spent at home, and a more dramatic rise in the proportion of days alive spent in an inpatient healthcare facility.

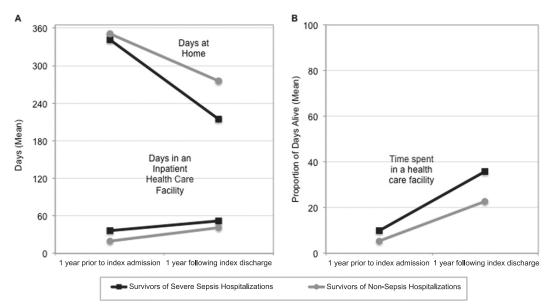
Our analysis provides two complementary perspectives on healthcare use after severe sepsis. The before versus after severe sepsis comparison (also known as within-person analysis) addresses the question: does severe sepsis represent an important change in patients' healthcare use? The answer is a compelling "yes" with,

for example, inpatient days increasing from 24.2 per patient-year before severe sepsis to 47.9 days per patient-year after severe sepsis. The comparison to matched nonsepsis hospitalizations addresses the question: is severe sepsis particularly worse in its disruption to patients' premorbid use than other illnesses with comparably severe inpatient stays? The answer here is more nuanced, with severe sepsis associated with much worse ongoing mortality, a finding that has been seen in many studies (5, 11), but similar changes in absolute healthcare needs relative to other similar hospitalizations. Altogether, we interpret this as meaning that severe sepsis is followed by a significant increased burden of healthcare use, but that increase is not uniquely worse than other severe illnesses requiring significant hospitalization.

Only 39% of the patients in our study were treated in an intensive care unit. Our findings suggest that patients with severe

sepsis, even the majority who do not require intensive care unit admission, often need prolonged and recurrent care in inpatient facilities after hospital discharge. Our findings are consistent a recent study by Liu and coworkers (36) demonstrating that sepsis survivors spend a threefold higher proportion of days alive admitted to a healthcare facility in the year after sepsis hospitalization compared with before. They are also consistent with prior studies that have quantified post-sepsis resource use (37-42), but have not included clear comparisons, control for presepsis healthcare use, or a focus on patientcentered transitions of care.

With the mounting attention on repeat admissions as a quality measure and potentially preventable expense for healthcare systems (13, 20), severe sepsis may be an important opportunity for targeted interventions. The rate of repeat hospitalization observed in our study



**Figure 3.** Difference-in-differences analysis of healthcare use in the severe sepsis and nonsepsis cohorts. The difference-in-differences between the severe sepsis and nonsepsis cohorts was (A) -38.6 days spent at home (95% confidence interval [CI], -50.9 to -26.3 d; P < 0.001), and (B) +5.4% of days alive spent in an inpatient healthcare facility (95% CI, +2.8 to +8,1%; P < 0.001). The difference-in-differences for days spent in a healthcare facility was not significant (-4.2 d; 95% CI, -9.0 to +0.7 d; P = 0.09).

exceeds the average readmission rate for Medicare beneficiaries (13, 43) and the rate observed in our matched nonsepsis hospitalizations, suggesting that severe sepsis survivors may benefit from more intensive post-discharge care management. This might include improved coordination of care, more frequent geriatric medicine involvement to help with new disabilities and functional limitations, support for caretakers (44), or an increased focus on in-hospital maintenance of the ability to live independently.

As the case fatality rate for severe sepsis continues to decline (45), it is important to consider additional metrics to measure the quality of care provided to severe sepsis patients. Post-discharge healthcare use is an attractive metric for several reasons. First, it is meaningful to patients whether, and to what extent, they need to spend time in LTACs and SNFs. Second, to decrease case fatality rates, hospitals may selectively transfer patients to post-acute care facilities, such as LTACs and SNFs (21), shifting the burden of severe sepsis to less studied silos. Third, post-acute care facility use represents the largest source of variation in healthcare spending across the United States (46), so it is worthy of study in and of itself. Thus, we urge researchers to consider post-discharge use as an

 Table 2. Difference-in-Differences Analyses Comparing Changes in Healthcare Use between the Severe Sepsis and Nonsepsis Cohorts\*

	Primary Cohorts (1,083 Matched Pairs)		ICU Subgroup (420 Matched Pairs)		Year-matched Cohorts (795 Matched Pairs)	
	DID Estimate (95% CI)	P Value	DID Estimate (95% CI)	P Value	DID Estimate (95% CI)	P Value
Days at home Days in an inpatient facility Days in a hospital Days in an LTAC Days in a SNF Transitions of care location Proportion of days alive in an inpatient facility	-38.6 (-50.9 to -26.3) -4.2 (-9.0 to 0.7) -1.9 (-4.0 to 0.2) 0.1 (-0.8 to 1.0) -2.4 (-6.0 to 1.3) -0.6 (-1.0 to -0.2) 5.4% (2.8 to 8.1%)	<0.01 0.09 0.08 0.87 0.20 0.01 <0.01	-43.6 (-63.4 to -23.7) -2.5 (-10.4 to 5.5) -2.5 (-6.1 to 1.1) -0.3 (-2.2 to 1.5) 0.4 (-5.2 to 5.9) -0.6 (-1.3 to 0.2) 7.8% (3.3 to 12.3%)	<0.01 0.54 0.18 0.72 0.90 0.13 <0.01	-33.9 (-48.4 to -19.4) -3.2 (-8.8 to 2.4) -1.2 (-3.6 to 1.1) 1.2 (0.3 to 2.2) -3.2 (-7.5 to 1.1) -0.5 (-1.0 to -0.2) 4.1% (0.9 to 7.3%)	<0.01 0.26 0.31 0.01 0.14 0.04 0.01

Definition of abbreviations: CI = confidence interval; DID = difference-in-differences; ICU = intensive care unit; LTAC = long-term acute care hospital; SNF = skilled nursing facility.

\*For each of the three analyses presented here, the severe sepsis and nonsepsis cohorts are matched on age, sex, premorbid disability, Charlson score for comorbidities, length of hospitalization, and whether or not intensive care was used for any portion of the hospitalization. The ICU subgroup is the subset of the primary analysis that used intensive care. The year-matched cohorts were matched on year, in addition to the other six matching variables, and are not a subset of the primary cohort.

important outcome for future observational and interventional studies of severe sepsis patients.

Our study has several potential limitations. We analyzed inpatient healthcare use through Medicare claims. Because of this, we were unable to assess long-term residence in nursing homes that is not covered by Medicare. In addition, we have not quantified informal care needs. Therefore, our results underestimate the magnitude of severe sepsis survivors' total needs (7), concentrating instead on those most relevant to the public cost of severe sepsis. We were not able to assess the extent to which an integrated health system may reduce healthcare use. Rather, our results focus on Medicare, the only data source with national scale and the payer for more than half of severe sepsis hospitalizations (47).

We excluded 7% of the severe sepsis sample because they could not be matched to nonsepsis hospitalizations using our matching algorithm. The excluded severe sepsis hospitalizations had similar baseline use to those included in the analysis. However, they had notably higher use and mortality in the year after severe sepsis. As such, our estimates may underestimate the true impact of severe sepsis on subsequent use. This decision was made so that a consistent group of severe sepsis patients was used for all comparisons.

#### Conclusions

We have shown that healthcare use of severe sepsis survivors is markedly elevated relative to their own premorbid resource use. The change in resource use was similar to that of otherwise similar nonsepsis hospitalizations. This finding suggests that, after controlling for comorbidity burden and functional limitations, much of a patient's subsequent use can be attributed to severity of illness and hospitalization. That is, severe sepsis may be a sufficient cause for a substantial increase in healthcare needs, but it is not a uniquely necessary cause of such an increase relative to other severe illnesses. However, severe sepsis survivors experienced increased mortality relative to

matched nonsepsis survivors. As a result, severe sepsis survivors spent a greater proportion of their days alive admitted to an inpatient healthcare facility and far fewer days at home in the year after hospital discharge. Our findings suggest a need for greater attention to the post-discharge management of severe sepsis survivors and further investigation into the excess mortality experienced by this population. As the case fatality rate for severe sepsis continues to fall, we believe it would be worthwhile to measure post-discharge use in studies of severe sepsis outcomes.

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#### References

- Angus DC, Linde-Zwirble WT, Lidicker J, Clermont G, Carcillo J, Pinsky MR. Epidemiology of severe sepsis in the United States: analysis of incidence, outcome, and associated costs of care. *Crit Care Med* 2001;29:1303–1310.
- Iwashyna TJ, Cooke CR, Wunsch H, Kahn JM. Population burden of long-term survivorship after severe sepsis in older Americans. *J Am Geriatr Soc* 2012;60:1070–1077.
- Kumar G, Kumar N, Taneja A, Kaleekal T, Tarima S, McGinley E, Jimenez E, Mohan A, Khan RA, Whittle J, *et al.*; Milwaukee Initiative in Critical Care Outcomes Research Group of Investigators. Nationwide trends of severe sepsis in the 21st century (2000-2007). *Chest* 2011;140: 1223–1231.
- Martin GS, Mannino DM, Eaton S, Moss M. The epidemiology of sepsis in the United States from 1979 through 2000. N Engl J Med 2003;348: 1546–1554.
- Quartin AA, Schein RM, Kett DH, Peduzzi PN; Department of Veterans Affairs Systemic Sepsis Cooperative Studies Group. Magnitude and duration of the effect of sepsis on survival. *JAMA* 1997;277: 1058–1063.
- Wunsch H, Guerra C, Barnato AE, Angus DC, Li G, Linde-Zwirble WT. Three-year outcomes for Medicare beneficiaries who survive intensive care. JAMA 2010;303:849–856.
- Iwashyna TJ, Ely EW, Smith DM, Langa KM. Long-term cognitive impairment and functional disability among survivors of severe sepsis. *JAMA* 2010;304:1787–1794.
- Guerra C, Linde-Zwirble WT, Wunsch H. Risk factors for dementia after critical illness in elderly Medicare beneficiaries. *Crit Care* 2012;16: R233.
- Yende S, Angus DC. Long-term outcomes from sepsis. Curr Infect Dis Rep 2007;9:382–386.
- Karlsson S, Ruokonen E, Varpula T, Ala-Kokko TI, Pettilä V; Finnsepsis Study Group. Long-term outcome and qualityadjusted life years after severe sepsis. *Crit Care Med* 2009;37: 1268–1274.

- 11. Winters BD, Eberlein M, Leung J, Needham DM, Pronovost PJ, Sevransky JE. Long-term mortality and quality of life in sepsis: a systematic review. *Crit Care Med* 2010;38:1276–1283.
- 12. Angus DC. The lingering consequences of sepsis: a hidden public health disaster? *JAMA* 2010;304:1833–1834.
- Jencks SF, Williams MV, Coleman EA. Rehospitalizations among patients in the Medicare fee-for-service program. N Engl J Med 2009;360:1418–1428.
- Dharmarajan K, Hsieh AF, Lin Z, Bueno H, Ross JS, Horwitz LI, Barreto-Filho JA, Kim N, Bernheim SM, Suter LG, et al. Diagnoses and timing of 30-day readmissions after hospitalization for heart failure, acute myocardial infarction, or pneumonia. JAMA 2013;309:355–363.
- Gill TM, Allore HG, Gahbauer EA, Murphy TE. Change in disability after hospitalization or restricted activity in older persons. *JAMA* 2010; 304:1919–1928.
- Krumholz HM. Post-hospital syndrome: an acquired, transient condition of generalized risk. N Engl J Med 2013;368:100–102.
- Dellinger RP, Levy MM, Rhodes A, Annane D, Gerlach H, Opal SM, Sevransky JE, Sprung CL, Douglas IS, Jaeschke R, *et al.*; Surviving Sepsis Campaign Guidelines Committee including the Pediatric Subgroup. Surviving Sepsis Campaign: international guidelines for management of severe sepsis and septic shock: 2012. *Crit Care Med* 2013;41:580–637.
- Torio CM, Andrews RM. National inpatient hospital costs: the most expensive conditions by payer, 2011. HCUP Statistical Brief #160. Rockville, MD: Agency for Healthcare Research and Quality; 2013 Aug.
- Unroe M, Kahn JM, Carson SS, Govert JA, Martinu T, Sathy SJ, Clay AS, Chia J, Gray A, Tulsky JA, *et al*. One-year trajectories of care and resource utilization for recipients of prolonged mechanical ventilation: a cohort study. *Ann Intern Med* 2010;153:167–175.
- Axon RN, Williams MV. Hospital readmission as an accountability measure. JAMA 2011;305:504–505.
- Hall WB, Willis LE, Medvedev S, Carson SS. The implications of longterm acute care hospital transfer practices for measures of inhospital mortality and length of stay. *Am J Respir Crit Care Med* 2012;185:53–57.

## **ORIGINAL ARTICLE**

- Kahn JM, Benson NM, Appleby D, Carson SS, Iwashyna TJ. Long-term acute care hospital utilization after critical illness. *JAMA* 2010;303: 2253–2259.
- Kahn JM, Kramer AA, Rubenfeld GD. Transferring critically ill patients out of hospital improves the standardized mortality ratio: a simulation study. *Chest* 2007;131:68–75.
- 24. Creditor MC. Hazards of hospitalization of the elderly. *Ann Intern Med* 1993;118:219–223.
- Prescott HC, Langa KL, Iwashyna TJ. Post-discharge health care use is markedly higher in survivors of severe sepsis. *Am J Respir Crit Care Med* 2013;187:A1573.
- Sonnega A, Faul JD, Ofstedal MB, Langa KM, Phillips JWR, Weir DR. Cohort profile: the Health and Retirement Study (HRS). Int J Epidemiol 2014;43:576–585.
- 27. Iwashyna TJ, Odden A, Rohde J, Bonham C, Kuhn L, Malani P, Chen L, Flanders S. Identifying patients with severe sepsis using administrative claims: patient-level validation of the Angus Implementation of the International Consensus Conference Definition of Severe Sepsis. *Med Care* 2014;52:e39–e43.
- Deyo RA, Cherkin DC, Ciol MA. Adapting a clinical comorbidity index for use with ICD-9-CM administrative databases. *J Clin Epidemiol* 1992;45:613–619.
- 29. King G, Neilsen R, Coberley C, Pope JE, Wells A. Comparative effectiveness of matching methods for causal inference. 2011 [accessed 2014 May 1]. Available from: http://gking.harvard.edu/ files/gking/files/psparadox.pdf
- Blackwell M, Iacus S, King G, Porro G. cem: Coarsened Exact Matching in Stata. Stata J 2009;9:524–546.
- 31. Munoz-Price LS. Long-term acute care hospitals. *Clin Infect Dis* 2009; 49:438–443.
- CMS. Medicare coverage of skilled nursing facility care. 2014 [accessed 2014 March 4]. Available from: http://www.medicare.gov/ pubs/pdf/10153.pdf
- 33. Iwashyna TJ. On the detection of nursing home use in Medicare claims. Health Serv Outcomes Res Methodol 2003;4:187–196.
- 34. Yun H, Kilgore ML, Curtis JR, Delzell E, Gary LC, Saag KG, Morrisey MA, Becker D, Matthews R, Smith W, et al. Identifying types of nursing facilities using Medicare claims data: an algorithm and validation. *Health Serv Outcomes Res Methodol* 2010;10:100–110.
- 35. Abadie A. Semiparametric difference-in-difference estimators. *Rev Econ Stud* 2005;72:1–19.

- Liu V, Lei X, Prescott HC, Kipnis P, Iwashyna TJ, Escobar GJ. Hospital readmission and healthcare utilization following sepsis in community settings. J Hosp Med (In press)
- Dick A, Liu H, Zwanziger J, Perencevich E, Furuya EY, Larson E, Pogorzelska-Maziarz M, Stone PW. Long-term survival and healthcare utilization outcomes attributable to sepsis and pneumonia. *BMC Health Serv Res* 2012;12:432.
- Weycker D, Akhras KS, Edelsberg J, Angus DC, Oster G. Long-term mortality and medical care charges in patients with severe sepsis. *Crit Care Med* 2003;31:2316–2323.
- Lee H, Doig CJ, Ghali WA, Donaldson C, Johnson D, Manns B. Detailed cost analysis of care for survivors of severe sepsis. *Crit Care Med* 2004;32:981–985.
- Braun L, Riedel AA, Cooper LM. Severe sepsis in managed care: analysis of incidence, one-year mortality, and associated costs of care. J Manag Care Pharm 2004;10:521–530.
- Letarte J, Longo CJ, Pelletier J, Nabonne B, Fisher HN. Patient characteristics and costs of severe sepsis and septic shock in Quebec. J Crit Care 2002;17:39–49.
- 42. Schmid A, Burchardi H, Clouth J, Schneider H. Burden of illness imposed by severe sepsis in Germany.*Eur J Health Econ* 2002;3:77–82.
- Wier LM, Barrett ML, Steiner C, Jiang HJ. All-cause readmissions by payer and age, 2008. HCUP Statistical Brief #115. Rockville, MD: Agency for Healthcare Research and Quality; 2011 Jun [accessed 2013 Jan 30]. Available from: http://www.hcup-us.ahrq.gov/reports/ statbriefs/sb115/pdf
- Davydow DS, Hough CL, Langa KM, Iwashyna TJ. Depressive symptoms in spouses of older patients with severe sepsis. *Crit Care Med* 2012;40:2335–2341.
- 45. Stevenson EK, Rubenstein AR, Radin GT, Wiener RS, Walkey AJ. Two decades of mortality trends among patients with severe sepsis: a comparative meta-analysis. *Crit Care Med* 2014;42:625–631.
- Institute of Medicine. Variation in health care spending: target decision making, not geography. Washington, DC: The National Academies Press; 2013.
- Elixhauser A, Friedman B, Stranges E. Septicemia in US hospitals, 2009. HCUP Statistical Brief #122. Rockville, MD: Agency for Healthcare Research and Quality; 2011 Oct [accessed 2013 Jan 23]. Available from: http://www.hcup-us.ahrq.gov/reports/statbriefs/ sb122.pdf