Insurance status and SCORTEN are associated with mortality in Stevens-Johnson Syndrome/Toxic Epidermal Necrolysis: a retrospective review comparing Black and non-Black patients Sach Thakker, BS,¹ Micah Belzberg, MD,² Jun Kevin Kang, MD²



BACKGROUND

- Stevens-Johnson syndrome/toxic epidermal necrolysis (SJS/TEN) is a life-threatening mucocutaneous disease classically triggered by a drug. Most common culprits are allopurinol, aromatic anticonvulsants, antibiotics, and nonsteroidal anti-inflammatory drugs.¹
- SJS is defined as <10% body surface area (BSA) detached, SJS/TEN overlap is defined 10% to 30% BSA detached, and TEN is defined as > 30% BSA detached.¹
- Prompt identification and discontinuation of a culprit drug is critical to improving patient outcomes and preventing recurrence.^{1,2}
- Hsu et al. studied the National Inpatient Sample and identified that SJS/TEN disproportionately affects Asian and Black patients.³
- This has been linked to use of certain medications and HLA haplotypes in these populations.⁴ However, there is limited data on other factors which may mediate the outcomes of SJS/TEN in minority populations. Recent research has emerged linking factors such as comorbidities and kidney function to SJS/TEN severity.²

Prognostic factors	Points
Age > 40 years	1
Heart Rate >120 bpm	1
Malignancy	1
Initiate detachment >10%	1
BUN > 10 mmol/L	1
Bicarbonate <20 mmol/L	1
Glucose >14 mmol/L	1
SCORETEN	Mortality (%)
SCORETEN 0-1	Mortality (%) 3
SCORETEN 0-1 2	Mortality (%) 3 12
SCORETEN 0-1 2 3	Mortality (%) 3 12 35
SCORETEN 0-1 2 3 4	Mortality (%) 3 12 35 58
SCORETEN 0-1 2 3 4 ≥5	Mortality (%) 3 12 35 58 90



Figure 1: Score of TEN (SCORTEN) criteria and mortality percentage Figure 2: SJS, SJS/TEN, TEN based off BSA distribution

METHODS

- All adult (age ≥18 years) patients diagnosed with SJS/TEN between 2010 to 2021 at the Johns Hopkins Hospital and Johns Hopkins Bayview Medical Center were included in this retrospective cohort review. The diagnosis of SJS/TEN was established by a dermatology consultant based on compatible clinical and histological features.
- All patients were screened to ensure their diagnosis met the predefined clinical criteria (eg, presence of epidermal detachment, involvement of ≥ 2 mucosae, atypical target lesions, histologic evidence supportive of SJS/TEN, and exclusion of differential diagnoses).
- Variables listed in results were manually collected from patient charts and analyzed through univariate and multivariable regression models.
- Statistical analyses were performed with Microsoft Excel (2023). Nominal variables were compared with chi squared tests while metric variables were assessed with t tests. Logistic regression analyses were also performed.

OBJECTIVE

- The primary objective of this study is to identify differences in SJS/TEN presentation and disparities in management and outcome between Black and non-Black patients at a large, tertiary care medical center with a diverse patient population.
- The secondary objective is to identify factors associated with increased risk of mortality in SJS/TEN at a large, tertiary care medical center with a diverse patient population.

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				RES	SULIS						
	All Patients N(%) or Avg	Non-Black N(%) or Avg	Black N(%) or Avg	p-value		Dead N(%) or Avg	Alive N(%) or Avg	OR(95% CI)	p-value		
N	89	38	51		Ν	18	71				
Female	53(60%)	21(55%)	32(63%)	0.47685	Female	10(56%)	43(61%)	0.81(0.29-2.31)	0.69901		
Age	54.0	53.6	54.3	0.88518	Age	61.2	52.2		0.11228		
House Value	\$314,918	\$521,703	\$155,130	0.00003	Race & Ethnicity						
Race & Ethnicity					Black	13(72%)	38(54%)	2.26(0.73-7)	0.15195		
Black	51(57%)				White	4(22%)	22(31%)	0.64(0.19-2.15)	0.46522		
White	26(29%)				Asian	1(6%)	10(14%)	0.36(0.04-3)	0.32610		
Asian	11(12%)				Other			0(0-0)	1.00000		
Other	1(1%)	O(E0())	0(00()		Hispanic		2(3%)	0(0-0)	1.00000		
	2(2%)	2(5%)	0(0%)			\$210,969	\$341,743		0.23130		
Privato	35(39%)	18(47%)	17(33%)	0 17000	Private	3(17%)	32(15%)	0.24(0.06-0.92)	0.02756		
Iningured	11(12%)	8(21%)	3(6%)	0.17999	Uninsured	5(28%)	6(8%)	4 17(1 1-15 72)	0.02730		
Non-Medicaid	11(12/0)	0(2170)	0(070)	0.00140	Non-Medicaid	0(2070)			0.02000		
Public	38(43%)	12(32%)	26(51%)	0.00639	Public	10(56%)	28(39%)	1.92(0.68-5.46)	0.21688		
Medicaid	5(6%)	0(0%)	5(10%)	0.17651	Medicaid	0(0%)	5(7%)	0(0-0)	1.00000		
Onset of symptoms					Onset of symptoms						
prior to presentation	3.6	3.9	3.3	0.24388	prior to presentation	2.7	3.8		0.08505		
(days)					(days)						
1st symptom	70/700/)	00/700/)	44(000())		1st symptom	40(000()	F 4 (7 00())		0.00504		
Rash	70(79%)	29(76%)	41(80%)	0.64249	Rash	16(89%)	54(76%)	2.52(0.53-12.08)	0.23534		
Systemic	/(8%)	2(5%)	5(10%)	0.43121	Systemic	1(6%)	6(8%)	0.64(0.07-5.66)	0.68361		
Boin Time to dermatology	12(13%)	7(10%)	5(10%)	0.23906	Time to dermatology	1(0%)	11(15%)	0.32(0.04-2.00)	0.27022		
consult (davs)	0.6	0.3	0.8	0.02573	consult (days)	1.0	0.4		0.04241		
Biopsy performed	86(97%)	37(97%)	49(96%)	0.73873	Biopsy performed	16(89%)	70(99%)	0.11(0.01-1.34)	0.04163		
Initial BSA%					Initial BSA%		1				
<10%	28(31%)	14(37%)	14(27%)	0.34531	<10%	2(11%)	26(37%)	0.22(0.05-1.02)	0.03738		
10-30%	33(37%)	13(34%)	20(39%)	0.62871	10-30%	3(17%)	30(42%)	0.27(0.07-1.03)	0.04471		
>30%	28(31%)	11(29%)	17(33%)	0.65939	>30%	13(72%)	15(21%)	9.71(2.99-31.54)	0.00003		
SCORETEN	2.5	2.2	2.8	0.04865	SCORETEN	3.8	2.2		0.00000		
Age≥40	64(72%)	26(68%)	38(75%)	0.39732	Age≥40	17(94%)	48(68%)	8.15(1.02-65.02)	0.02192		
BSA≥10%	61(69%)	24(63%)	37(73%)	0.34531	BSA210%	T6(89%)	45(63%)	4.62(0.98-21.72)	0.03738		
Malignancy	13(15%)	10(26%)	3(0%)	0.00694	Heart Pate (>120)	3(20%)	O(11%)	3.03(0.00-10.75)	0.07646		
Heart Rate (≥ 120)	$\frac{19(21\%)}{21(25\%)}$	4(11%) 7(190/)	13(29%)	0.03150	$\frac{\text{Reall Rate (\geq 120)}}{\text{Bicarbonato (} < 20)}$	0(44 %)	11(15%)	4.30(1.41-13.31) 4.01(1.36,11,70)	0.00742		
$\frac{\text{Bicarbonate}(<20)}{\text{Clusses}(>250)}$	2(2%)	1(3%)	24(47%)	0.00000	$\frac{\text{Bicarbonale}(<20)}{\text{Glucose}(>250)}$	1(6%)	20(2076)	4.01(1.30-11.79) 4.12(0.24-60.22)	0.00079		
$\frac{\text{BLIN}(>28)}{\text{BLIN}(>28)}$	40(45%)	13(34%)	27(53%)	0.03273	BUN (>28)	15(83%)	25(35%)	9(2 37-34 12)	0.00030		
Risks	+0(+070)	10(0+70)	21(0070)	0.00+73	Risks		20(0070)	0(2.07 01.12)	0.00000		
Age	54.0	53.6	54.3	0.88518	Age	61.2	52.2		0.11228		
Malignancy	13(15%)	10(26%)	3(6%)	0.00694	Malignancy	5(28%)	8(11%)	9(2.37-34.12)	0.07648		
Heart Rate	99.9	94.1	104.2	0.01837	Heart Rate	114.3	96.2		0.00046		
Bicarbonate	21.9	22.8	21.2	0.07046	Bicarbonate	20.1	22.4		0.03667		
Glucose	132.5	130.0	134.4	0.68606	Glucose	153.3	127.2		0.04520		
BUN	33.1	28.1	37.0	0.13722	BUN	55.3	27.4		0.00007		
Cr	1.7	1.4	2.0	0.03354	Cr	2.5	1.5		0.01104		
eGFR	45.6	50.6	41.9	0.03269	eGFR	34.7	48.3		0.00592		
HIV+	9(10%)	2 (5%)	7(14%)	0.19027	HIV+	2(11%)	7(10%)		0.87496		
Hospital acquired infection					Hospital acquired infection	40(4000())					
(bacteremia, pneumonia, UTI, or	35(40%)	11(29%)	24(47%)	0.08359	monia, UTI, or fungemia)	18(100%)	14(20%)		<.00001		
Turigemia) Offending Agent	No etc	l atistically sign	ificant differe	nce	Offending Agent	No	No statistically significant difference				
Therapeutic					Therapeutic	No statistically significant difference			<u>م</u>		
Intervention	No sta	atistically sign	ificant differe	nce							
Length of admission (days)	16.9	15.3	18.1	0.46791	Length of admission (days)	15.1	17.4		0.62513		
Deceased	18(20%)	5(13%)	13(25%)	0.15195	J Table 2. Sub-an	alvses of r	lata haser	on mortality	(Ava		

Table 1: Patient characteristics and outcome measures in Black to Non-Black patients.

Cases	Correct assignments	In percent			Coefficient B	SE	Z	p-value	OR	95% CI
89	79	88.76%		Non-Medicaid Public Insurance	0.6	0.9	0.67	0.501	1.83	0.31 - 10.65
				Medicaid Insurance	-18.49	6299.13	0	0.998	0	0 - ∞
	Predicated Alive	Predicted Dead	Correct	Uninsured	2.69	1.15	2.34	0.019	14.77	1.55 - 140.32
Observed Alive	67	4	94.37%	SCORETEN	1.51	0.46	3.27	0.001	4.53	1.83 - 11.23
Observed Dead	6	12	66.67%	GFR	0	0.02	0.17	0.866	1	0.97 - 1.04
Total			88.76%	Time to consult	0.47	0.31	1.52	0.128	1.61	0.87 - 2.96
				Constant	-7.16	2.25	3.18	0.001		
-2 Log-Likelihood	Cox & Snell R ²	Nagelkerke R ²	McFadden's R ²							
53.53	0.33	0.53	0.4							

Table 3: Logistic regression analysis of insurance type, SCORTEN, GFR and time to dermatology consult on mortality.

average; OR, odds ration; 95% CI, 95% confidence interval).

- regression analyses.

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CONCLUSIONS

• Black patients presented with higher SCORTEN, had a longer time to Dermatology consultation, and were more likely to be uninsured.

• Mortality (25% vs 13%, p = 0.15) and hospital acquired infection (47% vs 29%, p = 0.08) trended higher in Black patients vs non-Black patients, but did not reach significance • Privately insured patients were significantly more likely to receive a dermatology consult on the same day of presentation (data not shown). This difference was observed despite that privately insured patients had significantly lower SCORTEN.

• Uninsured status and higher SCORETEN on presentation were associated with increased mortality in univariate and logistic regression analyses.

• Patient race/ethnicity was not significantly associated with increased mortality in logistic

• While delay in Dermatology consultation was associated with increased mortality in univariate analysis, this was was not observed in logistic regression analyses.

REFERENCES

Zimmerman D, Dang NH. Stevens–Johnson Syndrome (SJS) and Toxic Epidermal Necrolysis (TEN). Oncologic Critical Care. Published online October 13, 2019:267-280. doi:https://doi.org/10.1007/978-3-319-74588-6_195

Micheletti RG, Chiesa-Fuxench Z, Noe MH, et al. Stevens-Johnson Syndrome/Toxic Epidermal Necrolysis: A Multicenter Retrospective Study of 377 Adult Patients from the United States. The Journal of Investigative Dermatology. 2018;138(11):2315-2321. doi:https://doi.org/10.1016/j.jid.2018.04.027

Hsu DY, Brieva J, Silverberg NB, Silverberg JI. Morbidity and Mortality of Stevens-Johnson Syndrome and Toxic Epidermal Necrolysis in United States Adults. *Journal of Investigative Dermatology*. 2016;136(7):1387-1397.

Lu N, Rai SK, Terkeltaub R, Kim SC, Menendez ME, Choi HK. Racial Disparities in the Risk of Stevens-Johnson Syndrome and Toxic Epidermal Necrolysis as Urate-Lowering Drug Adverse Events in the US. Seminars in arthritis and rheumatism. 2016;46(2):253-258. doi:https://doi.org/10.1016/j.semarthrit.2016.03.014

Ploysyne Rattanakaemakorn, Pasita Palakornkitti, Prinpat Pinyowiwat, Phatphitcha Jedee, Kunlawat Thadanipon. Chronic kidney disease is potentially an independent prognostic factor for death in Stevens-Johnson syndrome and toxic epidermal necrolysis patients. Frontiers in Medicine. 2022;9. doi:https://doi.org/10.3389/fmed.2022.939210