

Racial and Ethnic Disparities in Travel for Head and Neck Cancer Treatment and the Impact of Travel Distance on Survival

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BACKGROUND: Patients who travel a long distance (≥50 miles) for cancer care have improved outcomes. However, to the authors' knowledge, the prevalence of long travel distances for treatment by patients with head and neck squamous cell carcinoma (HNSCC), and the effect of travel distance on overall survival (OS), remains unknown. METHODS: The authors used the National Cancer Data base from 2004 through 2013 to identify patients with HNSCC undergoing definitive treatment. Travel distance for treatment was categorized as short (<12.5 miles), intermediate (12.5-49.9 miles), and long (50-249.9 miles). The primary outcome, OS, was evaluated using Cox shared-frailty modeling. A secondary outcome, factors associated with intermediate and long travel distances, was evaluated using multivariable hierarchical logistic regression. RESULTS: Among 118,000 patients with HNSCC, 62,753 (53.2%), 40,644 (34.4%), and 14,603 (12.4%) patients, respectively, traveled short, intermediate, and long distances for treatment. After adjusting for relevant covariates, long travel distance was associated with treatment at academic and high-volume centers. Patients of black race. of Hispanic ethnicity, with Medicaid insurance, and who were treated with nonsurgical treatment were less likely to travel long distances for treatment (P<.001). Traveling a long distance for treatment was associated with improved OS on multivariable analysis (adjusted hazard ratio, 0.93; 95% confidence interval, 0.89-0.96) compared with a short distance. CONCLUSIONS: Traveling a long distance for HNSCC treatment is associated with improved survival, especially for patients receiving nonsurgical management. Racial and ethnic disparities in travel for HNSCC treatment exist. As regionalization of care continues, future work should identify and address reasons for racial and ethnic disparities in travel that may prevent access to care at high-volume facilities. Cancer 2018;124:3181-91. © 2018 American Cancer Society.

KEYWORDS: head and neck cancer, health services, quality of care, racial disparities, travel distance.

INTRODUCTION

Numerous studies have demonstrated a positive relationship between treatment at a high-volume facility and improved outcomes such as survival for a variety of malignancies, ^{1,2} including head and neck squamous cell carcinoma (HNSCC). ³⁻⁵ Although it is responsible for approximately 12,000 deaths annually in the United States, ⁶ HNSCC is relatively uncommon and requires complex, multidisciplinary management for optimal outcomes. ⁷ Therefore, some have advocated for the regionalization of HNSCC care to high-volume centers. ⁸ Receiving care at high-volume centers, which generally are located in large urban areas, may require some patients to travel greater distances. ⁹ Although traveling a greater distance for cancer treatment has been associated with improved outcomes for patients with prostate, colon, esophageal, liver, and pancreatic malignancies, ¹⁰⁻¹² to our knowledge the effect of travel distance on survival in patients with HNSCC remains unknown.

Prior work has demonstrated the existence of racial disparities in travel for cancer care, ^{10,13} especially for travel to high-volume facilities. ^{12,14} The increasing travel requirements that have occurred contemporaneously with the progressive regionalization of cancer care may be creating another barrier to equitable, quality cancer care and contributing toward worsening racial and ethnic disparities in cancer outcomes. ¹⁴ However, to our knowledge, the prevalence of long travel distances for treatment by patients with HNSCC has not been described and the question of whether racial disparities in patterns of travel for HNSCC exist has not been investigated to date.

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Given the areas of uncertainty in knowledge regarding travel patterns among patients with HNSCC, we sought to answer the following questions: 1) what effect does travel distance have on overall survival (OS)?; 2) how frequently do patients travel a long distance (50-249.9 miles) for the treatment of HNSCC?; and 3) which patients are likely to travel long distances for HNSCC treatment?

MATERIALS AND METHODS

Data Source

The National Cancer Data Base (NCDB) is a hospital-based cancer registry that is a joint program of the American College of Surgeons Commission on Cancer (CoC) and the American Cancer Society. The NCDB annually collects high-quality and internally appraised data from >1500 CoC-accredited hospitals in the United States. 15

Study Cohort

The Medical University of South Carolina institutional review board deemed this study exempt from review. This article was reviewed and approved by the American College of Surgeons CoC. The NCDB from 2004 through 2013 was used to examine travel distance among patients with HNSCC undergoing curative-intent treatment. HNSCC diagnoses were filtered using International Classification of Disease for Oncology, Third Edition codes (Supporting Table 1) and SCC histology codes. A total of 131,147 patients were identified as undergoing treatment at the reporting facility (to ensure correct correlation of travel distance for treatment). Patients with an American Joint Committee on Cancer (AJCC) clinical M classification of 1 or unknown (4350 patients), those who received palliative-intent treatment (2431 patients), those with a time to treatment initiation \geq 180 days (3567 patients), and those with a travel distance ≥250 miles or unknown (2799 patients)^{10,13} were excluded, producing a cohort of 118,000 patients.

Study Variables

The primary outcome measure was OS. Travel distance to the reporting facility is provided by the NCDB as the greatest circle distance (in miles) between the patient's and hospital's ZIP code centroid. ¹⁶ In this study, travel distance was categorized into short (<12.5 miles), intermediate (12.5-49.9 miles), and long (50-249.9 miles) based on prior work. ^{10,13,17,18}

Patient-level covariates included age, sex, race, ethnicity, educational attainment, income, rurality, and insurance. Race and ethnicity were presented according to

the Standards for the Classification of Federal Data on Race and Ethnicity as approved by the Office of Management and Budget. ¹⁹ Clinical factors included severity of comorbidity, oncologic characteristics (tumor site, AJCC clinical stage), treatment year, and treatment modality. Treatment modality was categorized as surgery, surgery plus adjuvant therapy (radiotherapy [RT] or chemoradiation [CRT]), primary RT, and primary CRT. Hospitallevel covariates included treatment facility type and annual facility volume, ¹² which was grouped in quartiles (1-9, >9 to 17, >17 to 43, and >43 cases/year). ⁴ Other categorical variables were grouped as previously described. ²⁰

Statistical Analysis

Descriptive statistics were generated to characterize patient-level and hospital-level characteristics and bivariate analyses were conducted to evaluate their relationship with travel distance. Differences between travel distance groups were analyzed using chi-square tests. Multivariable hierarchical, multinomial logistic regression analysis was performed to analyze the relationship between covariates and travel distance for treatment. A 2-level hierarchical regression model (patient-level/hospital-level characteristics and US geographic region) was chosen to assess the relationship between patient-level/hospital-level variables with travel distance because the data likely are correlated.¹³ That is, cancer care provider density is heterogeneous across the United States as a whole but more homogenous within regions of the United States, and therefore patients within the same region face similar travel distance choices when accessing care.¹⁷

The relationship between travel distance and survival was analyzed using a multivariable Cox shared-frailty model for the same reasons that a hierarchical logistic regression was performed. Associations between covariates were investigated using the variable inflation factor before modeling. The overall variable inflation factors for all variables were <5 (except insurance, for which it was 6), and therefore none were deemed collinear. Variables significant at an α level of .05 on univariable analysis were entered into the multivariable Cox shared-frailty model. Possible interaction effects between travel distance, survival and race, subsite, AJCC clinical stage, treatment modality, facility volume, and facility type were examined. Interaction terms for race, subsite, and treatment modality were significant and were included in a separate survival model.

Multiple sensitivity analyses were performed to confirm the robustness of the results. To ensure that the relationship between travel distance and survival was not an artifact of the categorization strata, the multivariable Cox

TABLE 1. Sociodemographic, Oncologic, and Treatment Characteristics Stratified by Travel Distance for Treatment

Characteristic	Short Distance N = 62,753	Intermediate Distance $N = 40,644$	Long Distance N = 14,603	P
Variable	No. (%)	No. (%)	No. (%)	
Age, y		(**)	(, ,	<.001
<50	10,247 (16.3)	7474 (18.4)	2812 (19.3)	
50-59	20,262 (32.3)	13,543 (33.3)	4837 (33.1)	
60-69	17,225 (27.4)	11,597 (28.5)	4062 (27.8)	
≥70	15,019 (23.9)	8030 (19.8)	2892 (19.8)	
Sex	-, (,	, ,	(3 3)	300.
Male	47,090 (75.0)	30,696 (75.5)	10,843 (74.3)	
Female	15,663 (25.0)	9948 (24.5)	3670 (25.7)	
Race	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	,	,	<.001
White	51,496 (82.1)	36,766 (90.5)	13,295 (91.0)	
Black	8894 (14.2)	2622 (6.5)	841 (5.8)	
Other	2363 (3.7)	1256 (3.0)	467 (3.2)	
Ethnicity	2000 (0.1.)	.200 (0.0)	(0.2)	<.001
Non-Hispanic	55,710 (88.8)	36,973 (91.0)	13,406 (91.8)	<.001
Hispanic	3195 (5.1)	954 (2.3)	306 (2.1)	
Other/unknown	2363 (3.7)	1256 (3.0)	467 (3.2)	
Charlson/Deyo Comorbidity Score	2003 (0.1)	1250 (5.0)	407 (3.2)	<.001
0	49,087 (78.2)	32,087 (78.9)	11,506 (78.8)	<.00
1				
	10,407 (16.6) 3259 (5.2)	6699 (16.5) 1858 (4.6)	2457 (16.8) 640 (4.4)	
≥2 Incurrence	3259 (5.2)	1656 (4.6)	640 (4.4)	< 001
Insurance	00 154 (41 7)	10.050 (45.0)	COOO (44 4)	<.001
Private	26,154 (41.7)	18,358 (45.2)	6002 (41.1)	
Medicare	23,775 (37.9)	14,677 (36.1)	5327 (36.5)	
Medicaid	6801 (10.8)	3511 (8.6)	1491 (10.2)	
Uninsured	3841 (6.1)	2467 (6.1)	997 (6.8)	
Other	2182 (3.5)	1631 (4.0)	786 (5.4)	
Education, quartiles		()		<.001
Highest	14,671 (23.5)	8193 (20.2)	1835 (12.6)	
Second highest	19,699 (31.4)	13,258 (32.6)	4781 (32.7)	
Second lowest	16,767 (26.7)	11,757 (28.9)	4944 (33.9)	
Lowest	11,502 (18.3)	7401 (18.2)	3024 (20.7)	
Unknown	24 (0.0)	25 (0.1)	19 (0.1)	
Income, quartiles				<.001
Highest	18,638 (29.7)	12,199 (30.0)	1556 (10.7)	
Second highest	16,228 (25.9)	11,235 (27.6)	3253 (22.3)	
Second lowest	14,393 (22.9)	10,195 (25.1)	5383 (36.9)	
Lowest	13,441 (21.4)	6952 (17.1)	4373 (29.9)	
Unknown	53 (0.1)	63 (0.2)	38 (0.3)	
County type				<.001
Metropolitan	58,821 (93.7)	28,752 (70.8)	6334 (43.4)	
Urban	2632 (4.2)	9832 (24.2)	6807 (46.6)	
Rural	47 (0.1)	1265 (3.1)	1157 (7.9)	
Unknown	1253 (2.0)	794 (2.0)	305 (2.1)	
Site				<.001
Oral cavity	14,072 (22.4)	11,516 (28.3)	5980 (41.0)	
Oropharynx	23,563 (37.5)	15,178 (37.3)	4603 (31.5)	
Hypopharynx	3374 (5.4)	1744 (4.3)	504 (3.5)	
Larynx	21,744 (34.7)	12,206 (30.0)	3516 (24.1)	
AJCC clinical stage grouping	, , ,	, , ,	, ,	<.001
I	15,639 (24.9)	10,229 (25.2)	3289 (22.5)	
II	9630 (15.3)	6274 (15.4)	2363 (16.2)	
 III	11,704 (18.7)	7299 (18.0)	2462 (16.9)	
IV	25,780 (41.1)	16,842 (41.4)	6489 (44.4)	
Treatment modality	25,155 (1111)	. 5,5 .2 ()	0.00 ()	<.001
Surgery	12,147 (19.4)	10,703 (26.3)	6044 (41.4)	<.501
Surgery plus adjuvant	12,490 (19.9)	8707 (21.4)	3864 (26.5)	
Radiotherapy	20,998 (33.5)	10,575 (26.0)	1781 (12.2)	
Chemoradiation	17,118 (27.3)	10,673 (26.0)	2914 (20.0	
	11,110 (21.3)	10,009 (20.2)	2914 (20.0	- 00-
Facility type	6100 (0.0)	2409 (9.4)	205 (0.0)	<.001
Community	6190 (9.9)	3408 (8.4)	325 (2.2)	
Comprehensive community	25,535 (40.7)	14,776 (36.4)	2661 (18.2)	
Academic	22,918 (36.5)	17,869 (44.0)	10,179 (69.7)	
Integrated network	6793 (10.8)	3547 (8.7)	972 (6.7)	
Unknown	1317 (2.1)	1044 (2.6)	466 (3.2)	

TABLE 1. Continued

Characteristic	Short Distance $N = 62,753$	Intermediate Distance $N = 40,644$	Long Distance $N = 14,603$	Р
1-9	18,505 (29.5)	7920 (19.5)	1022 (7.0)	
>9 to 17	17,808 (28.4)	10,839 (26.7)	1927 (13.2)	
>17 to 43	15,817 (25.2)	10,164 (25.0)	3069 (21.0)	
>43	10,623 (16.9)	11,721 (28.8)	8585 (58.8)	
Region of United States				<.001
New England	3915 (6.4)	2118 (5.3)	210 (1.5)	
Middle Atlantic	9980 (16.2)	5035 (12.7)	1164 (8.2)	
South Atlantic	13,883 (22.6)	9826 (24.8)	3097 (21.9)	
East North Central	13,785 (22.4)	7882 (19.9)	2341 (16.6)	
East South Central	3904 (6.4)	4031 (10.2)	1911 (13.5)	
West North Central	3854 (6.3)	3171 (8.0)	2400 (17.0)	
West South Central	3694 (6.0)	2962 (7.5)	1207 (8.5)	
Mountain	2291 (3.7)	1394 (3.5)	780 (5.5)	
Pacific	6130 (3.7)	3181 (8.0)	1027 (7.3)	
Year of diagnosis				<.001
2004-2005	10,120 (16.1)	5902 (14.5)	2065 (14.1)	
2006-2007	10,838 (17.3)	6603 (16.2)	2367 (16.2)	
2008-2009	13,032 (20.8)	8183 (20.1)	2943 (20.2)	
2010-2011	13,809 (22.0)	9311 (22.9)	3380 (23.1)	
2012-2013	14,953 (23.8)	10,645 (26.2)	3848 (26.4)	

Abbreviations: AJCC. American Joint Committee on Cancer.

shared-frailty model was repeated using travel distance as a continuous variable with adjusted hazard ratios (aHRs) reported for 10-mile travel increments. 10,13 Although prior studies analyzing travel distance excluded patients traveling >250 miles, 10,13,17,18 we performed an additional sensitivity analysis including those patients who traveled >250 miles. Because of the biological and prognostic differences between carcinogen-mediated and human papillomavirus (HPV)-related HNSCC, 21 we performed a sensitivity analysis excluding patients with HPV-related oropharyngeal SCC (using Collaborative Stage Site-Specific Factor 10 codes 020-060). Because the NCDB did not record HPV status until 2010, but many patients from 2004 through 2010 likely had HPV-related cancers, an additional sensitivity analysis was performed excluding all patients with oropharyngeal SCC.

Data analysis was performed using SAS statistical software (version 9.4; SAS Institute Inc, Cary, North Carolina) and R packages (R Foundation, Vienna, Austria). All statistical tests were 2-sided with statistical significance set at a *P* value of .05. Measures of precision of point estimates are presented as 95% confidence intervals (95% CIs).

RESULTS

Demographic, Clinicopathologic, and Treatment Characteristics

A total of 118,000 patients with HNSCC undergoing treatment from 2004 through 2013 were included in the analysis. The patient demographic, clinicopathologic,

and treatment characteristics and their relationship to travel distance for treatment are presented in Table 1. Overall, 53, 34, and 12% of patients traveled short, intermediate, and long distances, respectively, for treatment of HNSCC. The mean travel distance was 5, 115, 94 miles for short, intermediate, and long distances, respectively.

Factors Associated With Increasing Travel Distance

A hierarchical multinomial logistic regression analysis was performed to identify factors associated with intermediate and long travel distances for treatment (Table 2). African American patients had 69% lower odds of traveling a long distance (adjusted odds ratio [aOR], 0.31; 95% CI, 0.28-0.34) compared with white patients. Hispanic individuals traveled a long distance less frequently than non-Hispanic individuals (aOR, 0.54; 95% CI, 0.46-0.62). Those patients with Medicaid (aOR, 0.75, 95% CI, 0.69-0.82) or no insurance (aOR, 0.82; 95% CI, 0.74-0.91) had a lower odds of traveling long distances compared with those with private insurance. Compared with patients undergoing surgical management, patients treated with RT (aOR, 0.21; 95% CI, 0.20-0.23) or CRT (aOR, 0.29; 95% CI, 0.27-0.32) were less likely to travel long distances for treatment.

Hospital-level characteristics were found to be associated with the odds of long travel distances for treatment. Patients were 12-fold more likely to travel a long distance

TABLE 2. Multivariable Hierarchical Regression Analysis of Factors Associated With Intermediate and Long Travel Distances for Treatment

Variable	Intermediate Versus Short Distance		Long Versus Short Distance	
	Adjusted OR (95% CI)	P	Adjusted OR (95% CI)	P
Age, y	.,		.,	•
<50	1.00 (reference)		1.00 (reference)	
50-59	0.93 (0.89-0.97)	.001	0.94 (0.87-1.01)	.07
60-69	0.93 (0.89-0.98)	.006	0.96 (0.88-1.04)	.32
≥70	0.72 (0.68-0.76)	<.001	0.73 (0.66-0.81)	<.00
Sex	(* * * * * * * * * * * * * * * * * * *		,	
Male	1.00 (reference)		1.00 (reference)	
Female	0.95 (0.92-0.99)	.007	0.88 (0.83-0.94)	<.00
Race	()		(,	
White	1.00 (reference)		1.00 (reference)	
Black	0.44 (0.41-0.46)	<.001	0.31 (0.28-0.34)	<.00
Other	0.69 (0.63-0.75)	<.001	0.72 (0.62-0.84)	<.00
Ethnicity	0.00 (0.00 0.1.0)	(100)	0.12 (0.02 0.0 1)	ν.σσ
Non-Hispanic	1.00 (reference)		1.00 (reference)	
Hispanic	0.48 (0.45-0.53)	<.001	0.54 (0.46-0.62)	<.00
Charlson/Deyo Comorbidity Score	0.40 (0.45-0.55)		0.54 (0.40-0.02)	<.00
	1 00 (reference)		1.00 (reference)	
0	1.00 (reference)	17	1.00 (reference)	- 00-
1	0.97 (0.93-1.01)	.17	0.87 (0.82-0.93)	<.00
≥2 !nouvenee	0.90 (0.83-0.95)	<.001	0.73 (0.65-0.82)	<.00
Insurance	4.00 (.6)		100 (6	
Private	1.00 (reference)		1.00 (reference)	
Medicare	0.98 (0.94-1.02)	.26	0.96 (0.89-1.03)	.22
Medicaid	0.82 (0.78-0.87)	<.001	0.75 (0.69-0.82)	<.00
Uninsured	0.88 (0.82-0.94)	<.001	0.82 (0.74-0.91)	<.00
Other	1.11 (1.03-1.21)	.009	1.50 (1.33-1.69)	<.00
Education, quartiles				
Highest	1.00 (reference)		1 (reference)	
Second highest	1.46 (1.40-1.53)	<.001	1.31 (1.21-1.42)	<.00
Second lowest	1.90 (1.81-2.01)	<.001	1.14 (1.03-1.25)	.008
Lowest	2.37 (2.22-2.53)	<.001	1.02 (0.91-1.14)	.77
Income, quartiles				
Highest	1.00 (reference)		1.00 (reference)	
Second highest	0.73 (0.70-0.76)	<.001	1.81 (1.66-1.97)	<.00
Second lowest	0.49 (0.46-0.51)	<.001	2.17 (1.98-2.39)	<.00
Lowest	0.27 (0.26-0.29)	<.001	1.74 (1.56-1.94)	<.00
County type				
Metropolitan	1.00 (reference)		1.00 (reference)	
Urban	14.97 (14.15-15.83)	<.001	65.59 (60.90-70.65)	<.00
Rural	118.4 (87.1-161.0)	<.001	943.6 (685.8->999)	<.00
Site	(*		, , , , , , , , , , , , , , , , , , , ,	
Oral cavity	1.00 (reference)		1.00 (reference)	
Oropharynx	0.93 (0.89-0.98)	.004	0.83 (0.77-0.89)	<.00
Hypopharynx	0.88 (0.81-0.95)	<.001	0.71 (0.62-0.81)	<.00
Larynx	0.94 (0.90-0.98)	<.008	0.80 (0.75-0.86)	<.00
AJCC clinical stage grouping	0.54 (0.56 0.56)	<.000	0.00 (0.70 0.00)	<.00
I	1.00 (reference)		1.00 (reference)	
II	1.11 (1.06-1.17)	<.001	1.40 (1.29-1.52)	<.00
III	• • • • • • • • • • • • • • • • • • • •	<.001		<.00
IV	1.11 (1.05-1.17)		1.52 (1.40-1.66)	
	1.19 (1.13-1.24)	<.001	1.83 (1.69-1.98)	<.00
Treatment modality	1.00 / (4 (6)	
Surgery	1.00 (reference)	224	1 (reference)	
Surgery plus adjuvant	0.77 (0.74-0.81)	<.001	0.53 (0.49-0.57)	<.00
Radiotherapy	0.61 (0.58-0.64)	<.001	0.21 (0.20-0.23)	<.00
Chemoradiation	0.67 (0.63-0.71)	<.001	0.29 (0.27-0.32)	<.00
Facility type				
Community	1.00 (reference)		1.00 (reference)	
Comprehensive community	1.39 (1.30-1.49)	<.001	3.90 (3.32-4.56)	<.00
Academic	1.71 (1.58-1.85)	<.001	12.34 (10.36-14.70)	<.00
Integrated network	1.00 (0.92-1.10)	.92	3.83 (3.32-4.56)	<.00
Facility annual volume, quartiles				
1 to 9	1.00 (reference)		1.00 (reference)	
>9 to 17	1.72 (1.64-1.81)	<.001	1.63 (1.46-1.82)	<.00
>17 to 43	2.02 (2.91-2.13)	<.001	2.95 (2.62-3.31)	<.00
>43	3.68 (3.47-3.91)	<.001	13.86 (12.26-15.67)	<.00

Abbreviations: CI, confidence interval; AJCC, American Joint Committee on Cancer; OR, odds ratio.

for care at an academic center (aOR, 12.3; 95% CI, 10.4-14.7) and 14-fold more likely to travel for care at a high-volume facility (aOR, 13.9; 95% CI, 12.3-15.7).

Subgroup Analysis Examining Reasons for Racial and Ethnic Differences in Travel

Given the observed racial and ethnic disparities in the likelihood of traveling for treatment, subset analyses in African American and Hispanic patients were performed to characterize determinants of travel in these groups. In the subset of African American individuals, any nonprivate form of insurance and residence within a ZIP code with lower income levels were associated with a decreased likelihood of traveling a long distance for HNSCC care (Supporting Table 2). Similar to the entire cohort, African American patients were less likely to travel for RT (aOR, 0.14; 95% CI, 0.10-0.19) and CRT (aOR, 0.19; 95% CI, 0.14-0.26) but more likely to travel a long distance for treatment at an academic (aOR, 10.4; 95% CI, 4.6-23.3) or highvolume (aOR, 10.4; 95% CI, 6.1-17.6) facility on hierarchical logistic regression modeling. In the subset of patients of Hispanic ethnicity (Supporting Table 3), uninsured patients (aOR, 0.47; 95% CI, 0.14-0.81) and those residing within a ZIP code with lower educational levels had lower odds of traveling a long distance. Hispanic individuals were more likely to travel a long distance for treatment at an academic (aOR, 12.5; 95% CI, 3.97-39.44) or high-volume (aOR, 9.59; 95% CI, 4.65-19.80) facility. There was no interaction observed between race, ethnicity, and the odds of traveling a long distance for HNSCC care (P = .07).

Association Between Travel Distance and Survival

In the multivariable Cox shared-frailty model adjusting for relevant covariates (Fig. 1), patients who traveled a long distance for treatment had improved OS compared with patients who traveled a short distance (aHR, 0.93; 95% CI, 0.89-0.96). Other covariates found to be associated with OS included age, race, insurance, comorbidity, subsite, AJCC clinical stage, treatment modality, facility type, and facility volume. In a subset analysis of African American patients, Cox shared-frailty modeling demonstrated that the risk of mortality for a long distance compared with a short distance was unchanged in terms of effect size (aHR, 0.92; 95% CI, 0.82-1.03).

Subgroup Analysis of Interaction Effects of Travel Distance With Other Covariates and Survival

Interaction effects between travel distance, survival and race, subsite, AJCC clinical stage, treatment modality, facility volume, and facility type were examined. Significant interactions were found for race, subsite, and treatment modality (P<.001), but not for disease stage (P = .58), facility volume (P = .95), or facility type (P = .80). A subgroup effects model demonstrating the interaction between travel distance and race, subsite, and treatment modality was developed (Supporting Table 4). Figure 2 shows the interaction between travel distance and race (Fig. 2A), subsite (Fig. 2B), and treatment modality (Fig. 2C). The subgroup analysis shows that for treatment modality, the improved OS observed with increased travel distance was primarily due to the benefit of longer travel for RT and CRT. Interaction testing for subsite demonstrated that the effect of travel distance on OS is mediated through oropharyngeal and oral cavity cancers.

Sensitivity Analyses

Numerous sensitivity analyses were performed to confirm the robustness of the relationship between longer travel and survival. Increasing travel distance remained associated with improved OS when analyzed as a continuous variable; the risk of death decreased by 1% for every 10-mile increase in travel distance (aHR, 0.99; 95% CI, 0.98-0.99). Including patients who traveled ≥250 miles for treatment did not change the improvement in survival noted with longer travel distance (aHR, 0.92; 95% CI, 0.89-0.95) (Supporting Table 5). Excluding patients with high-risk HPV-positive oropharyngeal SCC (aHR, 0.89; 95% CI, 0.86-0.92) (Supporting Table 6) and all patients with oropharyngeal SCC (aHR, 0.93; 95% CI, 0.90-0.97) (Supporting Table 7) did not alter the relationship between longer travel distance and improved survival. An additional sensitivity analysis excluding patients with oral cavity cancer was performed because these individuals were more likely to travel a long distance for treatment and had improved survival. The association between long travel distance and survival remained unchanged (aHR, 0.89; 95% CI, 0.85-0.93) (Supporting Table 8).

DISCUSSION

In the current study, we demonstrated that patients who traveled a long distance (50-249.9 miles) for HNSCC treatment had a decreased risk of death compared with those who traveled a short distance (<12.5 miles). To our knowledge, the current study is the first to demonstrate a

Forest Plot of Factors Associated with Survival

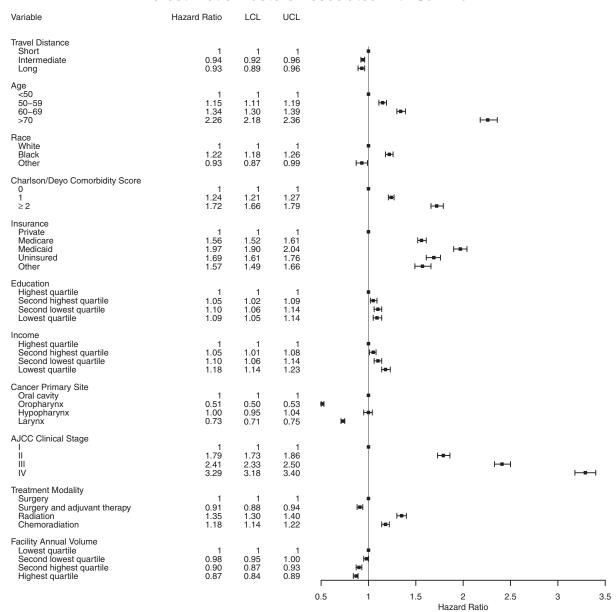


Figure 1. Multivariable Cox shared-frailty model demonstrating the effect of travel distance for treatment and other covariates on survival among 118,000 patients treated for head and neck squamous cell carcinoma. Estimated adjusted hazard ratios are shown as black circles; the 95% confidence intervals are represented by horizontal lines. AJCC indicates American Joint Committee on Cancer; LCL, lower confidence limit; UCL, upper confidence limit.

significant association between increasing travel distance and improved OS among patients with HNSCC. The association between travel distance and survival has been examined for other cancer sites and the current study findings are consistent with these prior studies. ¹⁰⁻¹² The survival benefit observed from increasing travel distance is partially a consequence of the regionalization of care to high-volume centers with resultant improvements in

oncologic outcomes, ^{11,12,14} because patients in the current study who traveled a long distance for treatment were found to be significantly more likely to be treated at high-volume and academic centers. In HNSCC, ³⁻⁵ as in other malignancies, ^{1,2} there is evidence to support a relationship between higher patient volumes and improved outcomes. However, the current study data support the association between long travel distance and survival independent of

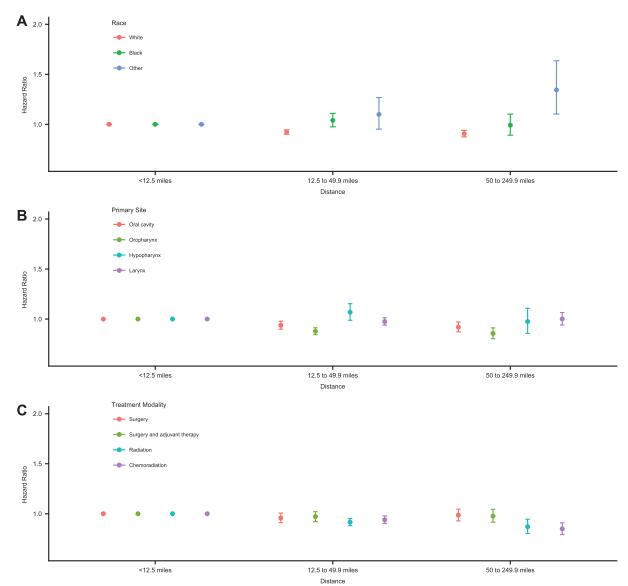


Figure 2. Subgroup analysis demonstrating interactions between covariates and travel distance for treatment for those covariates for which significant interactions existed (race, primary site, and treatment modality). Each hazard ratio is the combined effect of the covariate and travel distance for treatment compared with a reference travel distance of <12.5 miles. Survival according to travel distance varied by (A) race (white, black, or other), (B) primary head and neck subsite (oral cavity, oropharynx, hypopharynx, or larynx); and (C) treatment modality (surgery, surgery and adjuvant therapy, radiotherapy, or chemoradiation).

facility type, facility volume, age, race, insurance status, treatment modality, and a variety of relevant clinical and sociodemographic variables. Similar effect sizes were noted in the African American patient subset as well as in numerous sensitivity analyses. Nevertheless, there is a likely role for unmeasured confounding patient and treatment facility characteristics in this association between longer travel distance and improved survival. Given that long travel distances generally are regarded as a burden to cancer care, 17 patients who can overcome the barrier

associated with travel distance possibly are more highly motivated, supported socially, health-seeking in their behaviors, and adherent to treatment recommendations. ^{10,11} Future research should identify these additional unmeasured variables that in part mediate the relationship between travel distance and survival and focus interventions on increasing these health-promoting behaviors among all patients with HNSCC.

Given the independent association between long travel distances and survival in patients with HNSCC,

identifying which patients are least likely to travel long distances for treatment is important to ensure equitable care. African American patients had a >3-fold decrease in the odds of traveling long distances for treatment compared with white patients, and Hispanic patients had a 2fold decreased odds of traveling a long distance compared with non-Hispanic patients. To the best of our knowledge, the reasons that African American and Hispanic patients are less likely to travel long distances for the treatment of HNSCC compared with white and non-Hispanic patients are not known but are likely multifactorial in nature. Racial and ethnic disparities in travel for cancer treatment have been documented for patients with non-HNSCC12,22; the results of the current study add to the growing literature. Lack of insurance, lack of access to an automobile or someone to drive patients to treatment, and financial toxicity as well as cultural beliefs regarding health care have been shown to contribute toward racial and ethnic differences in travel patterns. 22-24 The results of the current study generally are in agreement, because we also observed that certain social determinants of health (insurance, education, and income) contributed toward low rates of traveling long distances for HNSCC care in African American and Hispanic patients. Two exceptions included the finding that higher educational attainment at census tract quartile levels was inversely associated with a higher likelihood of long travel distance in African American patients and that lower census tract income quartiles were not associated with decreased odds of traveling longer for HNSCC care. Whether these racial and ethnic differences in income and education represent true associations or are artifacts of the data collection (ZIP code-level quartiles) is unknown.

Racial and ethnic disparities in HNSCC survival were present in the current study and are well documented in other studies.^{25,26} These disparities are due in part to inequities in stage of disease at the time of presentation, timely care, guideline-concordant care, and access to care. 25,26 The exact roles that racial and ethnic disparities in travel distance play in racial and ethnic disparities in outcomes is unknown. The independent improvement in survival associated with long travel distances for treatment in the African American subset was of the same magnitude as observed in the overall cohort. Although African American patients were more likely to be treated at academic centers and high-volume centers when they traveled a long distance for treatment, the benefit of longer travel persisted independent of these factors for African American individuals as it did for the cohort at large. As cancer care continues to be centralized in high-volume institutions and travel distances for the treatment of HNSCC care increase, ¹⁴ racial and ethnic differences in travel for HNSCC care may exacerbate existing racial and ethnic disparities in outcomes. Further work is necessary to identify and address barriers related to travel for HNSCC care for African American and Hispanic patients to develop strategies to improve the equity and quality of HNSCC care.

We also found that patients who underwent nonsurgical management were 3-fold less likely to travel long distances for treatment, even though traveling a long distance for nonsurgical management correlated with improved survival. The reasons for discrepancies in willingness to travel for surgical versus nonsurgical care for HNSCC are unknown. However, the fact that patients are more willing to travel for surgery than (chemo)radiation has been documented for other malignancies, 10 despite studies demonstrating a volume-outcome relationship for RT.²⁷ Differential travel patterns for the surgical and nonsurgical treatment of HNSCC have implications for multidisciplinary evaluation and management, processes of care that improve survival. As cancer care continues to regionalize to high-volume centers, identifying and addressing the barriers to travel for patients receiving nonsurgical modalities will be a critically important part of elevating the quality of care for all patients with HNSCC.

In addition to race, ethnicity, and treatment modality, other factors were found to be associated with the likelihood of traveling a long distance for HNSCC care. As noted in other oncologic sites, ^{10,11,18} patients with more severe comorbidities were less likely to travel, presumably reflecting on the need to be sufficiently healthy to withstand the physical demands of long travel.

Limitations

The current study had important limitations. Because it was a retrospective database study, reasons for the choice of travel distance could not be discerned. These may include factors related to patient motivation, insurance network restrictions, local referral patterns, travel cost, health-seeking behaviors, and social support. The calculation of travel distance to all surrounding hospitals was not possible. Therefore, we were unable to ascertain whether patients voluntarily traveled greater distances to seek care (bypassing a possible treatment facility) when they could have sought care closer to home or know where they stayed during treatment. Provider density, which is known to vary across the United States, 17 was not assessed. Hierarchical regression modeling was used to control for this, but whether this

technique fully addressed this concern is unknown. Treatment biases inherent in the retrospective observational study design may affect survival. Multilevel Cox models were used to control for this source of bias, but statistical analysis cannot control for relevant variables not captured in the NCDB. In addition, the survival benefit observed with surgery compared with RT, although consistent with other studies using the NCDB,²⁸ was not fully explored and likely is related to the use of RT for oral cavity cancer, differences in the frequency of AJCC stage 1 disease, treatment at academic centers, and treatment at high-volume centers.³ Individual, patient-level socioeconomic information is not available in the NCDB. Adjustments were made for ZIP code-level income and education, but these may be inadequate. Despite these limitations, the current study possesses numerous methodological strengths. It captured patients of all adult ages, with a variety of insurance types, and different treatment modalities; had a national scope, large sample size, and relevant oncologic details; and analyzed treatment at different types of hospitals.

Conclusions

Traveling a long distance for the treatment of HNSCC is associated with improved OS, especially for patients receiving nonsurgical management. Racial and ethnic disparities in travel for HNSCC treatment exist. As regionalization of care continues, future work should identify and address the reasons for racial and ethnic disparities in travel that may prevent access to care at high-volume facilities.

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CONFLICT OF INTEREST DISCLOSURES

The authors made no disclosures.

AUTHOR CONTRIBUTIONS

Evan M. Graboyes: Conceptualization, formal analysis, methodology, writing-original draft, and writing-review and editing. Mark A. Ellis: Conceptualization, data curation, formal analysis, and writing-original draft. Hong Li: Conceptualization, formal analysis, methodology, and writing-review and editing. John M. Kaczmar, Anand K. Sharma, Eric J. Lentsch, Terry A. Day, and Chanita Hughes Halbert: Conceptualization, methodology, and writing-review and editing.

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