# The Latinx Disparity in Surgery for Kidney Cancer: Data from The South Texas Region

Furkan Dursun, MD,¹ Rahul S. Patel, MS,¹ Dawn S. Hui, MD,² Hanzhang Wang, MD,¹ Ahmed M. Mansour MD,¹,³ Deepak K. Pruthi, MD,¹,³ David G. Alonzo, MD,⁴ Lalithapriya Jayakumar, MD,⁵ Ronald Rodriguez, MD,¹,³ Robert S. Svatek, MD,¹,³ Michael A. Liss, MD,¹,³ Dharam Kaushik, MD.¹,³

- 1. Department of Urology, University of Texas Health San Antonio, TX, USA
- 2. Department of Cardiothoracic Surgery, University of Texas Health San Antonio, TX, USA
- 3. University of Texas Health San Antonio/ MD Anderson Mays Cancer Center, TX, USA
- 4. Department of Urology, University of Texas Rio Grande Valley, TX, USA
- 5. Department of Surgery, Division of Vascular & Endovascular Surgery, University of Texas Health San Antonio, TX, USA doi.org/10.52733/KCJ20n1-a1

### ABSTRACT

The South Texas region, with a predominantly Latinx population, has a very high incidence of renal cell carcinoma (RCC), including those with tumor extending into the major blood vessels called venous tumor thrombus (VTT). There is currently no data on outcomes of Latinx patients with VTT as most published studies are from predominantly Caucasian population. Therefore, we performed this study to fill an urgent, unmet need. We reviewed patients who underwent radical nephrectomy with removal of VTT (called tumor thrombectomy) between 2015 and 2020. We collected data on demographics, clinical, pathological characteristics and outcomes of patients. Univariate and multivariate Cox regression analyses were used to evaluate the associations between ethnicity and disease progression or survival. We identified 112 patients, of which 67 (62%) were Latinx, and 41 (38%) were non-Latinx. Approximately 60% of patients had Level II-IV VTT; Latinx presented with a higher level of tumor thrombus (p=0.046). Latinx patients had a higher rate of no insurance (11% vs. 27%, p=0.04) and were more likely being lost to follow-up after surgery (22.4% vs. 13.3%, p=0.23) compared to non-Latinx. Fewer Latinx received systemic therapy (28% vs. 42%; p=0.13). Ninety-day mortality for the entire cohort was 3.8%. The Latinx population in the South Texas region present late, with advanced thrombus level, and do not have access to systemic therapy. Given symptomatic disease, surgical treatment, if feasible, is their only option. Our results highlight disparate treatment patterns which require further investigation and health-care policy changes.

# **KEYWORDS:** Latinx; Renal cell carcinoma; Tumor thrombectomy; Venous tumor thrombus

### INTRODUCTION

idney cancer is the 8<sup>th</sup> most common malignancy in the United States with an estimated 73,750 new cases and approximately 15,000 mortalities in 2020.1 Renal cell carcinoma (RCC) has a tendency for vascular invasion: 10-15% of patients have venous tumor thrombus (VTT) and 1-3% of cases have tumor thrombus extending into the right atrium.<sup>2,3</sup> Inferior vena cava involvement or obstruction by tumor thrombus can produce various clinical manifestations, including gross hematuria, lower extremity edema, varicocele, abdominal distension with hepatic dysfunction (possibly related to a Budd-Chiari syndrome), fatigue, malabsorption, shortness of breath, and pulmonary emboli.4

Even in the era of targeted therapy and subsequent immune checkpoint inhibitors, radical nephrectomy (RN) with tumor thrombectomy remains the mainstay of treatment for RCC patients with VTT. This aggressive surgery has significant morbidity and mortality, but the 5-year survival rates can be significant as up to 70%, especially for patients without metastatic disease.<sup>5</sup>

Contemporary management of RCC with VTT requires an interdisciplinary approach and resources to provide the best curative or palliative care, which is mostly available in tertiary care centers. Unfortunately, racial disparities exist both in the access

<sup>\*</sup> Corresponding Author: Dharam Kaushik M.D. Department of Urology, University of Texas Health, San Antonio, TX 78229-3900 Tel: 210-567-5676. Email: kaushik@uthscsa.edu; drdkuro@gmail.com

DEMOGRAPHICS		NON-LATINX	LATINX	TOTAL	P VALUE
Age	Age median (IQR)	63 (56-68.5)	58 (50-67)	61 (52-68)	0.08
Gender	Male Female	28 (62.22%) 17 (37.78%)	50 (74.63%) 17 (25.37%)	78 (69.64%) 34 (30.36%)	0.16
ВМІ	mean/median, median (IQR)	30.7 (25.6-33.8)	31 (26.3-34.3)	30.8 (26-33.8)	0.8
Smoking status	No Yes Former	29 (64.44%) 5 (11.11%) 11 (24.44%)	40 (59.7%) 12 (17.91%) 15 (22.39%)	69 (61.61%) 17 (15.18%) 26 (23.21%)	0.62
Chronic Conditions	DM HTN Hypercholesterolemia	16 (35.56%) 31 (68.89%) 16 (35.56%)	33 (50%) 41 (62.12%) 20 (30.3%)	49 (44.14%) 72 (64.86%) 36 (32.43%)	0.13 0.46 0.56
ASA Score	2 3 4	0 (0%) 23 (52.27%) 21 (47.73%)	1 (1.54%) 38 (58.46%) 26 (40%)	1 (0.92%) 61 (55.96%) 47 (43.12%)	0.54
	Preop Creatinine	1.2 (0.9-1.41)	0.95 (0.8-1.32)	1.05 (0.86-1.38)	0.06
Symptoms	Gross Hematuria Flank/back pain Weight loss Pulmonary Emboli Deep vein thrombus	18 (40.91%) 18 (40%) 19 (42.22%) 3 (6.82%) 0 (0%)	34 (51.52%) 29 (43.94%) 27 (40.91%) 2 (3.23%) 2 (3.23%)	52 (47.27%) 47 (42.34%) 46 (41.44%) 5 (4.72%) 2 (1.89%)	0.28 0.68 0.89 0.39 0.23
Insurance status	Medicare/Medicaid Private None County Insurance	18 (40%) 21 (46.67%) 5 (11.11%) 1 (2.22%)	25 (37.31%) 18 (26.87%) 18 (26.87%) 6 (8.96%)	43 (38.39%) 39 (34.82%) 23 (20.54%) 7 (6.25%)	0.04
Distance to hospital median (IQR)	·	51 (26-108)	54 (16-190)	52 (19-165.5)	0.79
Income	<\$40,227 \$40,227 - \$50,353 \$50,354 - \$63,332 \$63,333 +	4 (8.89%) 12 (26.67%) 5 (11.11%) 24 (53.33%)	33 (50%) 16 (24.24%) 8 (12.12%) 9 (13.64%)	37 (33%) 28 (25.23%) 13 (11.71%) 33 (29.73%)	<0.001
Education Level*	>17.6% 10.9% - 17.5% 6.3% - 10.8% <6.3%	14 (31.11%) 11 (24.44%) 9 (20%) 11 (24.44%)	49 (74.24%) 10 (15.15%) 4 (6.06%) 3 (4.55%)	63 (56.76%) 21 (18.92%) 13 (11.71%) 14 (12.61%)	<0.001
Clinical size	Clinical max size, median (IQR)	101 (77-127)	90 (70-120)	96 (74-123)	0.59
Thrombus level	0 1 2 3 4	10 (22.73%) 12 (27.27%) 6 (13.64%) 11 (25%) 5 (11.36%)	15 (22.39%) 7 (10.45%) 24 (35.82%) 16 (23.88%) 5 (7.46%)	25 (22.52%) 19 (17.12%) 30 (27.03%) 27 (24.32%) 10 (9.01%)	0.046
Clinical LN involvement	•	13 (28.89%)	22 (33.33%)	35 (31.53%)	0.15
Clinical Metastasis		12 (26.67%)	16 (24.24%)	28 (25.23%)	0.79

Table 1 | Baseline characteristics and disease characteristics of Latinx and Non-Latinx \*Education level represents the percentage of adults in the patient's zip code who did not graduate from high school. Higher number signifies area with a greater number of people who did not complete high-school graduation.

administered in the U.S.<sup>6,7</sup> Most of racial disparity studies have focused on differences between African Americans and non-Latinx Whites (NLW), there is paucity of data in RCC regarding disparities in care among the Latinx population, which is the largest minority with 18.5% of the total U.S. population.<sup>8</sup>

to health care and the quality of care population, the South Texas region on tumor thrombus patients is has a very high incidence of RCC, in the predominantly Caucasian including those with VTT.9 Of all population with insurance, we the South Texas Latinx, 31-61% are performed this study to evaluate uninsured.<sup>10</sup> This disparity results in disparities between Latinx and delayed presentation of patients with non-Latinx with RCC and VTT in RCC. However, there is a paucity of South Texas region. Our institution data on outcomes of Latinx patients is a tertiary referral center with who underwent extirpative surgery a catchment area of 2.5 million. their symptomatic disease. We are uniquely positioned as we With a predominantly Latinx Since most published literature accept all South Texas patients for

SURGICAL CHARACTERISTICS	NON-LATINX	LATINX	TOTAL	P VALUE
Anesthesia time (min), median (IQR) Operative time (min), median (IQR) Estimated blood loss (ml), median (IQR) Blood transfusion (ml), median (IQR) Length of Hospital Stay (day), median (IQR)	465 (350-575) 349 (253-446) 1200 (500-3000) 825 (0-1650) 7 (5-11)	452 (358-594) 335 (249-426) 1000 (437.5-2050) 500 (0-1200) 6 (4-9)	459 (356-584) 340 (251-432) 1000 (500-2375) 555 (0-1330) 6 (4-10)	0.95 0.84 0.54 0.67 0.06
SURGICAL APPROACH Open Laparoscopic Robotic	39 (95.12%) 2 (4.88%) 0 (0%)	55 (93.22%) 2 (3.39%) 2 (3.39%)	94 (94%) 4 (4%) 2 (2%)	0.47
SURGICAL INCISION Chevron Midline Thoracoabdominal Subcostal	28 (73.68%) 5 (13.16%) 2 (5.26%) 3 (7.89%)	34 (64.15%) 7 (13.21%) 8 (15.09%) 3 (5.66%)	62 (68.13%) 12 (13.19%) 10 (10.99%) 6 (6.59%)	0.54
Sternotomy	11 (26.83%)	12 (22.22%)	23 (24.21%)	0.60
Cytoreductive Nephrectomy	9 (22.5%)	17 (29.82%)	26 (26.8%)	0.42
BYPASS  No bypass  Cardiopulmonary  Veno-venous	34 (82.93%) 3 (7.32%) 4 (9.76%)	44 (80%) 8 (14.55%) 3 (5.45%)	78 (81.25%) 11 (11.46%) 7 (7.29%)	0.43
Intraoperative TEE	25 (62.5%)	30 (57.69%)	55 (59.78%)	0.64
Liver Mobilization	16 (41.03%)	33 (63.46%)	49 (53.85%)	0.03
Pringle Maneuver	3 (7.69%)	4 (7.69%)	7 (7.69%)	1
Lymph Node Dissection	27 (62.79%)	45 (68.18%)	72 (66.06%)	0.56
CLAVIEN DINDO CLASSIFICATION OF COMPLICATIONS  2 3a 3b 4 5	1 (2.27%) 0 (0%) 6 (13.64%) 2 (4.55%) 1 (2.27%)	3 (4.92%) 3 (4.92%) 8 (13.11%) 2 (3.28%) 3 (4.92%)	4 (3.81%) 3 (2.86%) 14 (13.33%) 4 (3.81%) 4 (3.81%)	0.30
Postop Creatinine	1.23 (1.05-1.68)	1.2 (1-1.49)	1.2 (1.02-1.6)	0.41
30-day readmission rate	5 (11.9%)	4 (6.78%)	9 (8.91%)	0.37
90-day mortality rate	1 (2.22%)	3 (4.48%)	4 (3.57%)	0.39

Table 2 | Surgical characteristics and complications of Latinx and Non-Latinx.

surgical services, regardless of their diagnosis, insurance status.

# **METHODS Patient Selection**

After approval by the Institutional Research Ethics Board (IRB number: HSC2021108E), we reviewed the medical records of our prospectively maintained database of patients who underwent RN with tumor thrombectomy for RCC between 2015 and 2020 at our institution.

#### **Covariates**

We abstracted patient-level characteristics including age at

sex, ethnicity, body mass index (BMI). American Anesthesiologists Society of (ASA) classification, smoking status. insurance status. and comorbidities such as diabetes mellitus (DM). hypertension and hypercholesterolemia. Socioeconomic variables including annual household income. education level and distance from home to hospital were estimated from county of residence by using patients' self-reported zip code. Ethnicity and race were determined by patient self-report on their electronic medical records (EPIC).

Disease characteristics included clinical TNM stage,<sup>11</sup> thrombus level, maximum clinical tumor size and preoperative lab values. Mayo Clinic RCC tumor thrombus level classification system was used for thrombus level.<sup>12</sup>

Operative data included total surgical time, surgical approach and incision type, type of bypass, use of intraoperative transesophageal echocardiography (TEE), utilization of liver mobilization or Pringle maneuver, resection of IVC, estimated blood loss (EBL), blood transfusion during surgery, and perioperative complications.

COMPLICATION	ETHNICITY	TUMOR THROMBUS LEVEL				
COMPLICATION		0	1	2	3	4
Intraoperative death	Latinx				1	
miraoperative death	Non-Latinx					
Perioperative death	Latinx				1	1
renoperative death	Non-Latinx		1			
Haemorrhage	Latinx				1	
паетюттаде	Non-Latinx			1		
Pulmonary embolus	Latinx			1		
Pulmonary embolus	Non-Latinx				1	
Acute renal failure	Latinx			2		
Acute renai fallure	Non-Latinx					1
Ileus	Latinx			2		
neus	Non-Latinx		1			
Pneumothorax	Latinx	1				
Frieumothorax	Non-Latinx					
Bile leak	Latinx	1				
Dile leak	Non-Latinx					
Wound dehiscence	Latinx		1			
vvouria deniscence	Non-Latinx					
Other complications	Latinx	2	1	1	4	
	Non-Latinx		1	1	1	1

Table 3 | Complications by tumor thrombus level and ethnicity

**Pathological** characteristics the tumor were collected from Early the pathology report. postoperative complications (< 90 days) were classified according to the Clavien-Dindo classification system. 30-day readmission rates, 90-day mortality rate, patient follow-up data including local recurrence, disease progression, and survival, as well as receipt of neoadjuvant or adjuvant chemotherapy, were recorded. Progression of disease was classified as attenuated (not until at least 6 months after surgery) or rapid (progression or death within 6 months of surgery).

### **Statistical Analysis**

associations between The ethnicity (Latinx or Non-Latinx) and demographic clinical characteristics assessed were with Fisher's exact test and the two-sample Wilcoxon rank-sum test for continuous variables. Chisquare test was used for assessing categorical variables. Univariate and multivariate Cox regression models were used to evaluate the

of associations between ethnicity and misease progression or survival. Significance was set at 0.05 and all p values were two-tailed. Statistics was performed using STATA 15.0 m. (StataCorp, College Station, Texas).

### **RESULTS**

We identified a total of 112 patients with bulky advanced RCC and VTT who underwent RN and tumor thrombectomy, including 67 (60%) Latinx and 45 (40%) non-Latinx patients. There were no significant differences observed between Latinxs and non-Latinx across age, sex, accompanying chronic conditions, ASA classification and BMI (Table 1). Most of the patients were male (70%) and more than half were obese with a median BMI of 30.8 (IQR 26-33.8). All but one patient was classified as ASA III (56%) or IV (43%). They had accompanying chronic conditions such as diabetes mellitus (44%), hypertension (65%), and hypercholesterolemia (32%). 15% of patients were active smokers. More than three-quarters (78%) of the patients had at least one complaint of constitutional symptoms; weight loss (41%), gross hematuria (47%), and/or flank pain (42%).

Latinx patients had significantly higher rate of no insurance compared to non-Latinx (26.9% vs. 11.1%, p=0.04). There was no significant difference for distance from home to hospital between Latinx and non-Latinx (54 miles Vs. 51 miles, p=0.79). The Latinx population had significantly lower annual income (p<0.001) and rate of high school graduation (p<0.001).

## Clinical and Perioperative Features

The median clinical maximum tumor length at presentation was 96 mm (IQR 74-123). Thirty two

percent of the patients presented with clinical node positive disease and 25 % with metastatic disease. Sixty-seven (60%) patients had level II-IV tumor thrombus indicating advanced disease. Sixty-two (55%) patients had right-sided tumors, which were more likely to present with level III or IV tumor thrombus than left-sided tumors (40% vs. 24%; p=0.07). Latinx presented with higher level of tumor thrombus (p=0.046). Level II-IV thrombus was seen in 67% of Latinx vs 50% of non-Latinx (Table 1).

The median operation time was 340 minutes (IQR 251-432). The median EBL was 1000 ml (IQR 500-2375) and the median blood transfusion during the surgery was 555ml (IQR 0-1330). There were significant increases in the EBL during surgery as the tumor thrombus level increased (p=0.003). There was also an increasing trend in the amount of blood given during surgery as the tumor thrombus level increased (p=0.08).

Table 2 lists the differences observed between Latinx and non-Latinx across intraoperative features, perioperative complications, 30-day readmission date and 90-day mortality. Latinx patients, due to the high rate of level II-IV tumor thrombus, indicating advanced disease burden, were more

	NON-LATINX	LATINX	TOTAL	P VALUE
Tumor volume, median (IQR) (cm³)	489 (188-963)	418 (288-908)	426 (246-933)	0.68
, , , , , , , , , , , , , , , , , , , ,	100 (100 000)	110 (200 000)	120 (210 000)	0.00
Positive surgical margin	3 (6.67%)	4 (5.97%)	7 (6.25%)	0.88
HISTOLOGY Clear cell	44 (04 440()	04 (04 040()	100 (01 070()	
Papillary type 1 or 2	41 (91.11%) 3 (6.67%)	61 (91.04%) 4 (5.97%)	102 (91.07%) 7 (6.25%)	
Other	1 (2.22%)	2 (2.98%)	3 (2.67%)	0.58
FUHRMAN GRADE 1	0 (0%)	4 (5.97%)	4 (3.57%)	
2	14 (31.11%)	15 (22.39%)	29 (25.89%)	
3 4	15 (33.33%)	23 (34.33%)	38 (33.93%)	0.32
Regional lymph node involvement	16 (35.56%) 6 (13.33%)	25 (37.31%) 8 (11.94%)	41 (36.61%) 14 (12.5%)	0.47
Trogional lymph hode involvement	0 (10.0070)	0 (11.5470)	14 (12.570)	0.47
Histological tumor necrosis	27 (61.36%)	43 (64.18%)	70 (63.06%)	0.76
Sarcomatoid component	5 (11.11%)	10 (14.93%)	15 (13.39%)	0.56
Rhabdoid component	7 (15.56%)	12 (17.91%)	18 (16.96%)	0.70
Translocation carcinoma	0 (0%)	7 (10.45%)	7 (6.25%)	0.03

Table 4 | Pathological outcomes of Latinx and Non-Latinx

likely to have a thoracoabdominal or by ethnicity (p=0.06). 30-day incision compared to non-Latinx readmission rate was 9% and there (15% vs. 5% p=0.21). Also, Latinx was no underwent more complex surgical maneuvers such as complete hepatic mobilization during surgery than ethnicity (p=0.37). non-Latinx (63% vs. 41%, p=0.03). The radical nephrectomy was done for cytoreductive purposes for 27% of the patients, and retroperitoneal lymph node dissection was utilized for 66% of the patients. Given advanced disease states, 60% had intraoperative TEE to evaluate in real time the level of thrombus during the surgery.

The 90-day surgical complications are summarized by tumor thrombus level and ethnicity in Table 3. One patient died during the surgery and three patients died in the postoperative 90 days (2.7%), with a total perioperative mortality of 3.6%. The median length of hospital stay was six (IQR 4-10) days. There was no statistically significant difference in the duration of hospitalization by tumor thrombus level (p=0.11)

statistically significant difference in the readmission rate by thrombus level (p=0.56) or by

### Tumor Histopathology and **Survival Outcomes**

The median tumor volume was 426 cm3 (IQR 246-933) on the pathological evaluation. Fourteen (12.5%)patients had regional lymph node involvement, seven patients (6.3%) had a positive surgical margin. Among 112 patients evaluated, 102 (91.1%) had clear cell RCC, and 7 (6.3%) had papillary RCC. There was no significant difference in the tumor histology by ethnicity (p=0.58). Pathologic features are listed in Table 4. More than 70% of the patients were marked as grade 3 or 4 within the Fuhrman grading system- an indicator of very aggressive cancer. Additionally, 63.1% of the patients had histological tumor necrosis, 13.4% had sarcomatoid differentiation, 17%

had rhabdoid differentiation, and 6.3% had translocation carcinoma. Latinx patients had higher incidence of very aggressive variant of RCC called translocation carcinoma: seven patients were reported with translocation renal cell carcinoma, and all of them were Latinx (p=0.03).

The median follow-up time was 453 days (IQR 129-1125). The Latinx was more likely had no postoperative follow-up compare to non-Latinx. (22.4% vs. 13.3%, p=0.23). There were no significant differences in 2-year and 3-year survival for Latinx and Non-Latinx patients (82% vs 83%, and 75% vs 77%, respectively, p=0.81 and 0.98). Forty-eight patients (43%) had progression, and twenty-one (18.8%) patients died of advanced RCC. Patients with higher Fuhrman grade were more likely to have disease progression (HR=2.08 p=0.001). Of those patients with metastatic disease, 33% received systemic therapy. We observed less utilization of systemic therapy in the Latinx group compared to the non Latinx

	Hazard Radio with 95% Cl	p-value
Survival*	0.9 (0.32, 2.51)	0.84
Disease progression	0.75 (0.36, 1.58)	0.45
Rapid progression	0.71 (0.24, 2.13)	0.54
Attenuated progression	1.11 (0.47, 2.66)	0.81

Table 5. Cox regression models for survival outcomes comparing Latinx vs Non-**Latinx:** \*Model adjusted for thrombus level, clinic stage, age, BMI, and gender.

group (28% vs. 42%; p=0.13). There procedure in a predominantly differences in 2- and 3-year survival single-institution, thrombus (2- year survival: 83% disease progression and survival (HR 0.75, CI 0.36–1.58, p= 0.45 and HR 0.9, CI 0.32-2.51, p= 0.84 respectively) (Table 5).

#### DISCUSSION

Texas is home to approximately 19% of Latinx population in the US, majority of which are concentrated increasing in population size which in the South Texas region. Since kidney cancer is the 4<sup>th</sup> leading cause of death among Latinx males, we attempted to study the disparity and variation in a highly complex the cancer incidence is not higher presentation- kidney cancer with tumor thrombus. The idea behind performing this study was to understand disparities to make system-based changes. Our goal was to twofold: to identify the disparity in presentation of advanced kidney cancer and identify differences in quality of care exist between Latinx and non-Latinx undergoing radical nephrectomy with tumor thrombectomy. In addition, we sought to share our experience the highest incidence.9 Two recent with this challenging, high-risk population-based studies reported

were no statistically significant Latinx cohort. In this high volume, retrospective rates when compared patients with study, we found that the Latinx in level o-II tumor thrombus and the South Texas region presented patients with level III-IV tumor late, with advanced thrombus level (67% vs. 50%, p=0.046). The 79% p= 0.687, and 3-year survival: uninsurance rate was significantly 76% - 74% p=0.454, respectively). higher in Latinx than non-Latinx In the multivariate cox regression (11% vs. 27% p=0.04), impacting the model adjusted for thrombus level, number of patients lost to followclinic stage, age, BMI, gender; the up and receiving systemic therapy. ethnicity was not found to be a Our results highlight disparate significant prognostic factor for treatment patterns across patients. Despite delayed presentation, our study shows that rapid coordination of extirpative surgery enhances the outcome for these patients with equivalent perioperative outcomes.

> Latinx is the largest and youngest racial/ethnic minority group in the US and rapidly is expected to be doubled over the next four decades.<sup>13</sup> Unlike non-Latinx, cancer is the leading cause of death among Latinx, even though than non-Latinx.<sup>14</sup> Latinx are more likely to present with an advanced disease which partly contributes to higher mortality rates, especially for colorectal, endometrial, liver, and stomach cancers.<sup>15</sup> The incidence and mortality of kidney cancer for Latinx living in the US is similar to all US populations.¹ But the incidence of kidney cancer in Texas, especially in South Texas, is higher than in the U.S., with Latinx significantly having

data on racial disparities in RCC. Callahan et al. assessed the reasons for the incidence disparity of RCC in African Americans and Whites, but Latinx were not evaluated in this study.<sup>16</sup> They reported that interventions that prevent hypertension and chronic kidney disease among African Americans could potentially eliminate the racial disparity in RCC incidence.<sup>16</sup> Utilizing the SEER database, Whitson et al. evaluated the prognostic factors of RCC patients with VTT, and they reported race was not associated with mortality (HR 1.2 CI 0.5-2.9, p=0.63).<sup>17</sup> Race or ethnicity was not reported in the prior case series for RCC and VTT. 18-21 Our study is unique, evaluating the disparities in patient presentation, treatment approaches, and outcomes for Latinx and Non-Latinx patients with RCC and VTT. It provides a contemporary real world scenario from South Texas- a region with very high incidence of kidney cancer, low insurance, diverse cultural and ethnic background.

with thrombectomy has proven long-term disease-free survival, especially for patients with the non-metastatic disease. 22, 23, 18, 20 The clear cell carcinoma is the predominant histology reported, changing between 70 to 93% in different case series. 19, 20, 21, 24 Neither thrombus level nor histological type was associated with progression and survival.<sup>18-24</sup> Likewise, most of our cases were clear cell RCCs (91%) and the histologic type and thrombus level were not associated with disease progression or survival.

Surgical management of RCC with VTT is challenging and requires multidisciplinary perioperative care. The surgical approach may involve full liver mobilization and thoracotomy to allow access to the intrahepatic and intrapericardial tumor thrombus especially for patients with level II-IV VTT. The intraoperative and postoperative death in 30 or 90 days was reported between 2.9 - 6.1% in the largest series. 18-21, 25 In the same series with available data, the perioperative death rate for level III-IV tumor thrombus patients was reported between 5.6-14.6%. 18-20, 24 Similar to those studies, the perioperative death rates of our cohort and for the subgroup of patients with level III-IV tumor thrombus was 3.6% and 8.1%, respectively.

Xp11.2 translocation RCC is caused by increased expression of transcription factor for immunoglobulin heavy chain enhancer 3 (TFE3) as a result of translocation of the TFE3 gene on chromosome Xp11.2. It mainly occurs in children and young adults, accounting for 20-40% of RCC cases in children. The incidence is low in adults accounting for approximately 1% of RCC and there are only 7 cases reported over the age of 65 years. 26,27 The prognosis is less aggressive in children and adolescents, but shows poor prognosis in adults.<sup>28</sup> Because of the rarity, our understanding of its pathogenesis is incomplete and there is no data about the ethnic differences in the incidence of Xp11.2 translocation RCC. We reported patients with translocation carcinoma and all of them were Latinx (p=0.03) with a median age of 50 years (range 21 to 67). Latinx patients had higher incidence of this aggressive variant of RCC which needs further investigation.

This study has several limitations. Foremost. this retrospective single-institution study is limited by inherent biases in patient selection, data collection, and analysis. The utilization of preoperative embolization, lymph node dissection, bypass, and surgical approach was based on the surgeon's clinical judgment and experience which may be a source of additional bias. The generalizability of our results is limited due to the high Latinx population in the South Texas region. However, the fundamental strength of this study lies in the analysis of a large sample size. To our knowledge, this is the first study assessing disparities in the presentation and quality of care between Latinx and non-Latinx RCC patients with venous tumor thrombus.

In conclusion, compared to Non-Latinx, the Latinx population in the South Texas region presents late, with advanced thrombus level, and does not have access to systemic therapy. Surgical treatment, if feasible, is their only option. Our results highlight disparate treatment patterns. A better understanding of the factors that cause such a high rate of uninsurance in the Latinx is crucial to creating more appropriate and effective reform strategies.

#### **FUNDING**

None

### **AUTHOR CONTRIBUTION**

DK planned the concept of the study and DK, FD planned the study design. FD, RP, and DK collected and verified the data. HW analyzed the data and DK, FD, RR and HW were involved in data interpretation. FD and RP wrote the first version of the manuscript. FD, RP, DH, HW, AM, DP, DA, LJ, RR, RS, ML and DK contributed to interpretation of the results and revision of the manuscript and approved the final version of the manuscript. Project administration was conducted by DK.

### **Declaration of competing interests**

The authors have no conflicts of interest.

### **ACKNOWLEDGEMENTS**

Furkan Dursun has received research support through the University of Texas Health Science Center at San Antonio Cancer Research Training Program supported by the CPRIT Research Training Award (*RTA*; *RP170345*). Dharam Kaushik is supported by Stanley and Sandra Rosenberg Endowment in Urologic Research

### **REFERENCES**

- 1. Siegel, R. L., Miller, K. D. and Jemal, A. (2020) 'Cancer statistics, 2020', CA: A Cancer Journal for Clinicians, 70(1), pp. 7–30. doi: 10.3322/caac.21590.
- 2. Glazer, A. A. and Novick, A. C. (1996) 'Long-term followup after surgical treatment for renal cell carcinoma extending into the right atrium', Journal of Urology, 155(2), pp. 448–450. doi: 10.1016/

S0022-5347(01)66415-2.

- 3. Kaushik D, Linder BJ, Thompson RH, Eisenberg MS, Lohse CM, Cheville JC, Leibovich BC, Boorjian SA. The impact of histology on clinicopathologic outcomes for patients with renal cell carcinoma and venous tumor thrombus: a matched cohort analysis. Urology. 2013 Jul;82(1):136-41. doi: 10.1016/j.urology.2013.02.034. Epub 2013 May 1. PMID: 23642851.
- 4. Sunela KL, Kataja MJ, Kellokumpu-Lehtinen PL. Changes in symptoms of renal cell carcinoma over four decades. BJU Int. 2010 Sep;106(5):649-53. doi: 10.1111/j.1464-410X.2010.09241.x. Erratum in: BJU Int. 2013 Dec;112(8):E436. PMID: 20151959.
- 5. Marshall, F. F. et al. (1988) 'Surgical management of renal cell carcinoma intracaval with neoplastic extension above hepatic veins', Journal of Urology, 139(6), pp. 1166-1171. doi: 10.1016/ s0022-5347(17)42848-5.
- 6. Meyer, P. A. et al. (2013) 'Conclusion and future directions: CDC Health Disparities and Inequalities Report United States, 2013.', Morbidity and mortality weekly report. Surveillance summaries (Washington, D.C.: 2002), 62 Suppl 3(3), pp. 184–186. Available at: https://pubmed.ncbi.nlm.nih.gov/24264513/ (Accessed: 5 April 2021).
- 7. DeSantis, C. E. et al. (2016) 'Cancer statistics for African Americans, 2016: Progress and opportunities in reducing racial disparities', CA: A Cancer Journal for Clinicians, 66(4), pp. 290–308. doi: 10.3322/caac.21340. 8. 2019 Population Estimates by Age, Sex, Race and Hispanic Origin (no date). Available at: https://www.census.gov/newsroom/press-kits/2020/population-estimates-detailed.html (Accessed: 5 April 2021).
- 9. Perez, C. et al. (2019) 'Abstract 4187: Racial disparities in kidney cancer incidence across South Texas, Texas, and United States using TCR, SEER and NPCR registries (2004-2014)', in Cancer Research. American Association for Cancer Research (AACR), pp. 4187–4187. doi: 10.1158/1538-7445.am2019-4187.
- 10. Quick Statistics on the Uninsured in Texas and the U.S. (no date). Available at: https://www.

texmed.org/template.aspx?id=5519 (Accessed: 5 April 2021).

Amin MB, Greene FL, Edge 11. SB. Compton CC, Gershenwald JE, Brookland RK, Meyer L, Gress DM, Byrd DR, Winchester DP. The Eighth Edition AJCC Cancer Staging Manual: Continuing to build a bridge from a population-based to a more "personalized" approach to cancer staging. CA Cancer J Clin. 2017 Mar;67(2):93-99. doi: 10.3322/ caac.21388. Epub 2017 Jan 17. PMID: 28094848.

Leibovich, B. C. et al. (2020) 12. 'Renal Cell Carcinoma with Inferior Vena Cava Extension: Can Classification Be Optimized to Predict Perioperative Outcomes?', Kidney Cancer, 4(2), pp. 111-115. doi: 10.3233/kca-190070.

Vespa, J., Medina, L. and 13. Armstrong, D. M. (no date) 'Demographic Turning Points for the United States: Population Projections for 2020 to 2060 Population Estimates and Projections Current Population Reports'. Available at: www.census.gov/programs-surveys/ popproj (Accessed: 23 July 2021).

Yanez, B. et al. (2016) 'Cancer Outcomes in Hispanics/Latinos in the United States: An Integrative Review and Conceptual Model of Determinants Health', Journal of Latina/o psychology, 4(2), p. 114. doi: 10.1037/ LAT0000055.

Pinheiro, P. S. et al. (2017) 'Cancer mortality in hispanic ethnic Cancer **Epidemiology** groups', Biomarkers and Prevention, 26(3), pp. 376–382. doi: 10.1158/1055-9965. EPI-16-0684.

Callahan, C. L. et al. (2020) 16. 'Understanding racial disparities in renal cell carcinoma incidence: estimates of population attributable risk in two US populations', Cancer Causes and Control, 31(1), pp. 85-93. doi: 10.1007/s10552-019-01248-1.

and Meng, M. V. (2013) 'Population of Surgery, 168(2), pp. 205-209. doi: based analysis of survival in patients 10.1016/S0002-9610(94)80069-3. with renal cell carcinoma and venous tumor thrombus', Urologic Oncology: 'Oncologic outcomes following surgical Seminars and Original Investigations, 31(2), pp. 259-263. doi: 10.1016/j. inferior vena caval thrombus extending urolonc.2010.11.017.

cell carcinoma with extension of tumor thrombus into the vena cava: Surgical strategy and prognosis', Journal of Vascular Surgery, 33(4), pp. 789–796. venous tumour thrombectomy managed doi: 10.1067/mva.2001.111996.

Mayo-Clinic experience with surgical Association Journal, 12(9), pp. E391management complications and outcome E397. doi: 10.5489/cuaj.5013. for patients with renal cell carcinoma and venous tumour thrombus', BJU International, 94(1), pp. 33-41. doi: Translocation Renal Cell Carcinoma: 10.1111/j.1464-410X.2004.04897.x.

term Survival in Patients Undergoing 2019 Aug 3;11(8):1110. doi: 10.3390/ Radical Nephrectomy and Inferior cancers11081110. PMID: Vena Cava Thrombectomy: Single-Center Experience', European Urology, 27. eururo.2009.06.009.

clinicopathologic and operative International factors on short-term and long-term Journal, 9(4), pp. 182-186. doi: 10.1007/ survival in renal cell carcinoma with \$13691-020-00430-6. venous tumor thrombus extension: A multi-institutional retrospective study 'Adult-onset renal cell carcinoma in Japan', BMC Cancer, 13(1), p. 1. doi: associated with Xp11.2 translocation/ 10.1186/1471-2407-13-447.

22. the vena cava', Journal of Urology, juro.2014.03.111. 145(1), pp. 20-23. doi: 10.1016/ s0022-5347(17)38235-6.

Swierzewski, D. J., Swierzewski, M. J. and Libertino, J. A. (1994) 'Radical nephrectomy in patients with renal cell

carcinoma with venous, vena caval, and Whitson, J. M., Reese, A. C. atrial extension, The American Journal

24. Haddad, A. Q. et al. (2014) resection of renal cell carcinoma with above the hepatic veins: A contemporary Tsuji, Y. et al. (2001) 'Renal multicenter cohort', Journal of Urology, 192(4), pp. 1050–1056.

25. Pruthi, D. K. et al. (2018) 'Contemporary surgical outcomes of with intraoperative Doppler ultrasound Blute, M. L. et al. (2004) 'The for kidney cancer', Canadian Urological

26. Caliò A, Segala D, Munari E, Brunelli M, Martignoni G. MiT Family from the Early Descriptions to the Ciancio, G. et al. (2010) 'Long- Current Knowledge. Cancers (Basel). PMCID: PMC6721505.

Masago, T. et al. (2020) 'Xp11.2 57(4), pp. 667-672. doi: 10.1016/j. translocation renal cell carcinoma with TFE3 gene fusion in the elderly: Hirono, M. et al. (2013) 'Impacts case report and literature review', Cancer Conference

28. Gong, P. et al. (2018)TFE3 gene fusion 3 case reports Hatcher, P. A. et al. (1991) and review of literature', Medicine 'Surgical management and prognosis (United States), 97(24). doi: 10.1097/ of renal cell carcinoma invading MD.000000000011023.10.1016/j.