

Nigerian Medical Journal

**Original Article** 

# Pilot Study on the Locoregional Demographics of Prostate Cancer in River State, Nigeria \*John E. Raphael, Okigbeye Danagogo

Urology Division, Department of Surgery, University of Port Harcourt Teaching Hospital, Port Harcourt, Rivers State, Nigeria

#### Abstract

**Background**: The aetiology of prostate cancer is a subject of intense research and has been attributed to several risk factors. Geographical variations have also been observed with high incidence in western countries and rising rates in developing countries. This study aims to evaluate the locoregional distribution of histologically confirmed prostate cancer patients and discuss the risk factors for the observed variations according to their Local Government Areas (LGA) in Rivers State.

**Methodology**: This is a descriptive retrospective study on patients with histologically confirmed prostate cancer from three hospitals that take urology referrals over 10years. The patient's hospital records were obtained, and records of their ages, permanent place of residence, prostate biopsy histopathology reports, and Gleason's scores were extracted. These data were collated and analyzed using SPSS version 20.

**Results**: There were 278 patients managed over a 10year period with a mean age of 68.39years+\_10.06. Prostate cancer was the commonest in the 60-69year age group (37.1%), followed by the 70-79year range. The high-risk, poorly differentiated cancer (Gleason 8-10; ISUP 4,5) was the commonest, followed by the intermediate-risk cancers (Gleason 7; ISUP 2,3). Port Harcourt (41.0%) and ObiaAkpo (36.0%) Local Government Areas (LGAs) in Port Harcourt City had the highest frequencies of prostate cancer. There was no relationship between age and Gleason's score in the men.

**Conclusion**: Prostate cancer appears more common in the LGAs in Port Harcourt City compared to the semi-urban and rural LGAs in Rivers State. High-grade cancers are more frequent both in rural and urban LGA. High Gleason score cancers appear more frequently in the metropolitan Port Harcourt City than rural LGAs. **Keywords**: Prostate Cancer; Demography; Distribution; Gleason's Score.

**Correspondence:** John Edoka Raphael, Urology Division, Department of Surgery, University of Port Harcourt Teaching Hospital, Port Harcourt, Rivers State, Nigeria **Email:** drraphaeljohn@gmail.com

How to cite this article: Raphael JE, Danagogo O. Pilot study on the Locoregional Demographic of prostate cancer in River State. Niger Med J 2021;62;(6): 340-345

**Quick Response Code:** 



This is an open access journal, and articles are distributed under the terms of the Creative Commons Attribution-Non-Commercial-Share Alike 4.0 License, which allows others to remix, tweak, and build upon the work non-commercially, as long as appropriate credit is given, and the new creations are licensed under the identical terms.

## Introduction

The aetiology of prostate cancer (PCa), though unknown, is a subject of intense research and has been attributed to several risk factors, including race, diet, serum androgen level, family history, and environmental factors. [1,2,3] Geographical variations have also been observed with high incidence in western countries and lower rates in developing countries.[1] Adaptation to Western influences with immigration and or urbanization has frequently been associated with an increased incidence in the same racial populations, suggesting that factors in the lifestyle change are possibly responsible for the increased risk. For example, studies have found a significant increase in the incidence of prostate cancer among African Americans who migrated to the United States of America compared to Africans living in Africa. A similar pattern was also observed among Chinese living in the USA vis-à-vis their counterparts living in China.[4,5] A study in Port Harcourt, Rivers State, Nigeria, investigated the ethnic variations among prostate cancer patients but did not put into consideration their locoregional distribution and place of residence.[6]This study aims at evaluating the locoregional distribution of patients with histologically confirmed prostate cancer and the risk factors for the observed variations according to their Local Government Areas (LGA) in Rivers State.

## Methods and Methodology

This is a descriptive retrospective study on patients with histologically confirmed prostate cancer from three hospitals that take urology referrals over a 10year period from 2011- 2020: University of Port Harcourt Teaching Hospital, Sophia Hospital, and Rosivylle Clinic and Urology Centre. The three Centers receive urology referrals from Port Harcourt and environs. Port Harcourt is a coastal city and capital of Rivers State, located in Southern Nigeria. The patients' hospital records were obtained, and their ages, permanent place of residence, prostate biopsy histopathology reports, and Gleason's scores were extracted. These data were collated using Microsoft Excel 2016 version (Microsoft Corporation, Redmond, WA, USA), and they were subjected to analysis using SPSS version 20.

#### Results

There were 278 patients managed over a 10year period with a mean age of 68.39years+\_10.06 and median age of 67.50years. Prostate cancer was most frequent in the 60-69year age group (37.1%), followed by the 70-79year range. The high-risk, poorly differentiated cancer (Gleason 8-10; ISUP 4,5) was the commonest, followed by the intermediate-risk cancers (Gleason 7; ISUP 2,3). Port Harcourt (41.0%) and Obia Akpo (36.0%) Local Government Areas in Port Harcourt City had the highest frequencies of prostate cancer. There was no relationship between age and Gleason's score in the men.

Table 1: Characteristics and Gleason's score of men with prostate cancer

Variable	Ν	%
Age group		
40-49	3	1.1
50-59	47	16.9
60-69	103	37.1
70-79	87	31.3
≥80	38	13.6
Grading (Gleason's score)		
Well-differentiated (2-6)	52	18.7
Moderately differentiated (7)	46	16.5
Poorly differentiated (8-10)	100	36.0
Unknown	80	28.8
Total	278	100.0

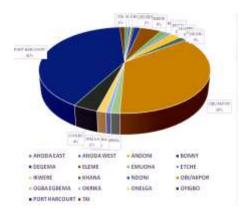


Figure 1: Distribution of prostate cancer patients by Local Government Areas in Rivers State.

				Grading	(Gleaso	n score)		
	Well			derately				
		rentiat		erentiate	Poor			
Age group	ed	(2-6)		d (7)		differentiated (8-10)		
								Unknown
	N	%	Ν	%	N	%	Ν	%
40 <b>-</b> 49	0	(.0)	1	(2.2)	0	(.0)	2	(2.5)
50-59	6	(11.5	8	(17.4)	14	(14.0)	19	(23.8)
		)						
60-69	22	(42.3	2	(43.5)	37	(37.0)	24	(30.0)
		)	0					
70-79	15	(28.8	1	(21.7)	39	(39.0)	23	(28.7)
		)	0					
>80	9	(17.3	7	(15.2)	10	(10.0)	12	(15.0)
		)						
Total	52	(100.	4	(100.0)	100	(100.0)	80	(100.0)
Chi-square	=14.41, p	0) p-value =	<u>6</u> 0.275	;				
Chi-square	=14.41, <u>r</u>		-	;	•			
	=14.41, <u>p</u>		-	5	8			
	=14.41, p		-	5				
	=14.41, p		-	5	9			

1.04

Table 2: Gleason's score distribution among prostate cancer patients.

Figure 2: Gleason score distribution among prostate cancer patients by Local Government Area in Rivers State.

## Discussion

Prostate cancer is the second most common cancer in men on the planet after lung cancer and was responsible for about 4% of all male cancer deaths in 2018. [7,8] Men of African descent, especially African Americans, have the highest incidence compared to Caucasians and Asian men, with 158.3 new cases diagnosed per 100,000, and about twice the mortality in Caucasians. [1] The reasons for this disparity have been attributed to differences in social, environmental, and genetic factors. African Americans also have a higher disease burden with higher grades, elevated prostate-specific antigen (PSA) levels, and higher mortality rates. [1,8] In Africa, studies indicated that compared to their North African counterparts, blacks in sub-Saharan Africa also have a higher disease burden, higher PSA, and disease-related mortality. [9,10]

The geographical incidence of PCa has been inconsistent and sometimes undependable, and the incidence of this deadly condition in most African countries is less than what has been reported.1 This may be attributed to shorter life expectancy in Africa, suboptimal testing, poor access to health facilities.[1,8,9]

Port Harcourt is the oil-rich capital city of the coastal Rivers State in Southern Nigeria with a heterogeneous and diverse population. In our study, the mean age of the patients was 68.39year, with the 60-69years age group having the highest frequency of prostate cancer (Table 1). This is similar to a study among African American men where the average age at diagnosis was 66years.[1]

Our study indicated that the Port Harcourt Local Government (PHALGA) and ObioAkpor Local Government Areas, both in the capital city of Port Harcourt, had the highest frequency of prostate cancer compared to the rural and suburban parts of the state (Figure 1). There are several possible explanations for these observations.

The risk factors that have been linked to prostate cancer include increased consumption of saturated animal fat and red meat, lower intake of fruits, vegetables, vitamins, obesity, physical inactivity, inflammation, hyperglycemia, infections, and environmental exposure to chemicals or ionizing radiation, more peculiar with city dwellers than suburban and rural dwellers [2,11, 12].

Chu et al. [4] observed that African Americans in the United States have a 40-fold higher incidence of prostate cancer when compared to those living in Africa. A similar pattern was reported by Hsing et al. in 2000 [5] among Chinese living in the USA have a 16-fold increased incidence of prostate cancer compared to those Chinese living in China. These findings are clear indications that environmental factors play an essential role in the pathogenesis of prostate cancer.

Increased saturated fat intake in butter, cheese, cake, snacks, and fried food is generally more associated with the city and urban dwellers. It has been speculated to be a risk factor for prostate cancer. Consumption of a large amount of saturated fat from animal sources has been demonstrated to lead to the proliferation of prostate cancer cells by increasing the circulating levels of testosterone. [13,14] A study that showed that vegetarians had lower levels of circulating serum testosterone [15-17] seems to lend credence to this observation. Saturated fats are also found to cause oxidative stress in rats and increase the production of reactive oxygen species, which attack cells and cause DNA damage that is a precursor for carcinogenesis.[18]

In addition, red meat consumption has been associated with prostate cancer [19]. Rohrmann et al. [20] showed that men consuming five or more servings of processed meat per week had a higher risk of prostate cancer when compared with men who took less than one serving per week. Also important is the cooking temperature as heating food at high temperatures >125°C, such as in grilling and barbecue, could cause the formation of aromatic hydrocarbons, N-nitroso compounds, and mutagenic heterocyclic amines [21] that can cause lipid peroxidation and DNA damage by the production of free radicals.[22]

Increased intake of milk and calcium, either from supplements usage, common among city dwellers, has been largely associated with an elevated prostate cancer risk.[23] A study that followed up about 47,885 men over 24 years found an association between high calcium intake with prostate cancer.24Generally speaking, dairy products are more available and affordable in towns and cities with better supply chains and retail outlets. City dwellers have higher incomes and can generally afford milk compared to people in rural areas.

However, protective lifestyles, behaviours, and dietary habits could account for the observed lower incidence of prostate cancer in rural areas. Fresh fruits and vegetables are generally more available in rural and semi-urban areas. Consumption of vegetables and fresh tomatoes may have some benefits in preventing prostate cancer. Crucifers or Brassica vegetables such as broccoli, brussels sprouts, cauliflower, cabbage, and turnips seem to reduce prostate cancer risk.[25] This antimitotic activity is thought to be mediated by phenylmethyl isothiocyanate, sulforaphane, phytochemicals, and indole-3-carbinol [25]. Some studies in the United States on a diet rich in these vegetables have shown evidence for the protective effects against prostate cancer [26]. These findings are, however, refuted by other studies. [27,28]

Tomatoes contain high levels of lycopene which are potent antioxidants and exhibit cancer-protective activities. Intake of lycopene has been associated with a reduced risk of prostate cancer. [29,30] Another effect that may add to this anti-prostate cancer activity is that lycopene also acts on the androgen receptors and reverses the effects of dihydrotestosterone, inhibits insulin growth factor-1, stimulation through Akt and GSK3β and tyrosine phosphorylation of GSK3.[30]. Venkateswaran and Klotz [31] in their study with transgenic mice showed that lycopene produced this

observed benefit in reducing prostate cancer incidence only when selenium and vitamin E was added. Another study, The Health Professional Follow-up Study, found a decreased risk with 2-4 servings of tomatoes per week. Tomato product consumption and lycopene intake were associated with a reduced risk of prostate cancer.[32]

Finally, obesity and sedentariness are generally more common in the cities than in rural areas. Both obesity and physical inactivity have been linked to advanced and aggressive prostate cancer. [32,33] Suggested reasons include the presence in obese people alteration of circulating levels of metabolic and sex steroid hormones associated with the development of prostate cancer.[34]

The insulin resistance in the obese leads to chronic hyperinsulinemia that promotes cell growth and proliferation, potentially leading to prostate cancer initiation and progression.[35] Additionally, Keogh and McLeod found that veterans who exercised had a significantly lower risk of prostate cancer.[36] Another study found that patients with advanced prostate cancer who exercise regularly had lower PSA levels and delay in initiating androgen deprivation therapy (ADT) by two years and had a lower risk of high-grade disease than less active peers[36].

## Conclusion

Prostate cancer appears more common in the urban LGAs in Port Harcourt City compared to the semi-urban and rural LGAs in Rivers State. High-grade cancers are more frequent both in rural and urban LGA. High Gleason score prostate cancers appear more frequently among Port Harcourt City dwellers than rural LGAs. Adoption of Western dietary habits could be a factor contributing to the observed differences.

## Limitations

This is a retrospective study with relatively small sample size. Larger prospective studies and randomized control trials are required to assess the impact of regional demographics on the risk of prostate. Source of funding: None

Acknowledgements: Professor N Eke for his contribution to data acquisition and Professor ON Ekeke the Head of Surgery for his inputs during the manuscript writing.

## Conflict of interest: None

## References

- 1. Panigrahi GK, Praharaj PP, Kittaka H, Mridha AR, Black OM, Singh R, Mercer R. et al. Exosome proteomic analyses identify inflammatory phenotype and novel biomarkers in African American prostate cancer patients. Cancer Med. 2019; 8:1110-1123.
- 2. Dagnelie PC, Schuurman AG, Goldbohm RA, Van den Brandt PA. Diet, anthropometric measures, and prostate cancer risk: a review of prospective cohort and intervention studies. BJU Int. 2004; 93:1139–1150.
- 3. Bostwick DG, Burke HB, Djakiew D, Euling S, Ho SM, Landolph J, Morrison H. et al. Human prostate cancer risk factors. Cancer. 2004; 101: 2371–2490.
- 4. Chu LW, Ritchey J, Devesa SS, Quraishi SM, Zhang H, Hsing AW. Prostate cancer incidence rates in Africa. Prostate Cancer. 2011; 2011: 947870.
- 5. Hsing AW, Tsao L, Devesa SS. International trends and patterns of prostate cancer incidence and mortality. Int J Cancer. 2000; 85:60–67.
- 6. Sapira MK, Eke N, Nwofor AME. Ethnicity and Prostate Cancer in Southern Nigeria: A Preliminary Report. Niger. J. Surg. 2015; 21: 96-101.
- Bray F, Ferlay J, Soerjomataram I, Siegel RL, Torre LA, Jemal A. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. CA Cancer J Clin. 2018; 68:394–424.
- 8. Ferlay J, Colombet M, Soerjomataram I, Mathers C, Parkin DM, Piñeros M, Znaor A, Bray F. Estimating the global cancer incidence and mortality in 2018: GLOBOCAN sources and methods. Int J Cancer. 2019; 144:1941-1953.
- 9. Chan JM, Gann PH, Giovannucci EL. Role of diet in prostate cancer development and progression. J Clin Oncol. 2005; 23: 8152–8160.
- 10. Giovannucci E, Rimm EB, Colditz GA, Stampfer MJ, Ascherio A, Chute CG, Willett WC. A prospective study of dietary fat and risk of prostate cancer. J Natl Cancer Inst. 1993; 85:1571–1579.

- 11. Markozannes G, Tzoulaki I, Karli D, Evangelou E, Ntzani E, Gunter MJ, Norat T. et al. Diet, body size, physical activity and risk of prostate cancer: An umbrella review of the evidence. Eur J Cancer. 2016; 69:61-69.
- Wilson KM, Giovannucci EL, Mucci LA. Lifestyle, and dietary factors in the prevention of lethal prostate 12. cancer. Asian J Androl. 2012; 14:365-374.
- Venkateswaran V, Klotz LH. Diet and prostate cancer: mechanisms of action and implications for 13. chemoprevention. Nat Rev Urol. 2010; 7:442-453.
- 14. Pauwels EK. The protective effect of the Mediterranean diet: focus on cancer and cardiovascular risk. Med Princ Pract. 2011; 20:103–111.
- 15. Fleshner N, Zlotta AR. Prostate cancer prevention: past, present, and future. Cancer. 2007; 110:1889–1899.
- Rosenthal MB, Barnard RJ, Rose DP, Inkeles S, Hall J, Pritikin N. Effects of a high-complex-carbohydrate, 16. low-fat, low-cholesterol diet on levels of serum lipids and estradiol. Am J Med. 1985; 78: 23-27.
- Hamalainen EK, Adlercreutz H, Puska P, Pietinen P. Decrease of serum total and free testosterone during a low-17. fat high-fibre diet. J Steroid Biochem. 1983; 18:369-370.
- 18. Lloyd JC, Masko EM, Wu C, Keenan MM, Pilla DM, Aronson WJ, Chi JT. et al. Fish oil slows prostate cancer xenograft growth relative to other dietary fats and is associated with decreased mitochondrial and insulin pathway gene expression. Prostate Cancer Prostatic Dis. 2013; 16:285–291.
- Gibson TM, Ferrucci LM, Tangrea JA, Schatzkin A. Epidemiological and clinical studies of nutrition. Semin Oncol. 19. 2010: 37: 282-296.
- Rohrmann S, Platz EA, Kavanaugh CJ, Thuita L, Hoffman SC, Helzlsouer KJ. Meat and dairy consumption and 20. subsequent risk of prostate cancer in a US cohort study. Cancer Causes Control. 2007; 18:450.
- 21. Sinha R, Knize MG, Salmon CP, Brown ED, Rhodes D, Felton JS, Levander OA. et al. Heterocyclic amine content of pork products cooked by different methods and to varying degrees of doneness. Food Chem Toxicol. 1998; 36: 289-297.
- Tappel A. Heme of consumed red meat can act as a catalyst of oxidative damage and could initiate colon, breast, and 22. prostate cancers, heart disease, and other diseases. Med Hypotheses. 2007; 68:562-564.
- 23. Gao X, LaValley MP, Tucker KL. Prospective studies of dairy product and calcium intakes and prostate cancer risk: a meta-analysis. J Natl Cancer Inst. 2005; 97:1768-1777.
- Wilson KM, Shui IM, Mucci LA, Giovannucci E. Calcium and phosphorus intake and prostate cancer risk: a 24-y 24. follow-up study. Am J Clin Nutr. 2015; 101:173-183.
- Singh SV, Srivastava SK, Choi S, Lew KL, Antosiewicz J, Xiao D, Zeng Y., et al. Sulforaphane-induced cell death in 25. human prostate cancer cells is initiated by reactive oxygen species. J Biol Chem. 2005; 280: 19911–19924.
- Joseph MA, Moysich KB, Freudenheim JL, Shields PG, Bowman ED, Zhang Y, Marshall JR. et al. Cruciferous 26. vegetables, genetic polymorphisms in glutathione S-transferases M1 and T1, and prostate cancer risk. Nutr Cancer. 2004; 50:206-213.
- 27. Stram DO, Hankin JH, Wilkens LR, Park S, Henderson BE, Nomura AM, Pike MC. et al. Prostate cancer incidence and intake of fruits, vegetables, and related micronutrients: the multiethnic cohort study\* (United States) Cancer Causes Control. 2006; 17:1193-1207.
- Key TJ, Allen N, Appleby P, Overvad K, Tjonneland A, Miller A, Boeing H. et al. Fruits and vegetables and prostate 28. cancer: no association among 1104 cases in a prospective study of 130544 men in the European Prospective Investigation into Cancer and Nutrition (EPIC) Int J Cancer. 2004; 109: 119-124.
- 29. Erdman JW Jr, Ford NA, Lindshield BL. Are the health attributes of lycopene related to its antioxidant function? Arch BiochemBiophys. 2009; 483: 229-235.
- Liu X, Allen JD, Arnold JT, Blackman MR. Lycopene inhibits IGF-I signal transduction and growth in normal 30. prostate epithelial cells by decreasing DHT-modulated IGF-I production in co-cultured reactive stromal cells. Carcinogenesis. 2008; 29:816-823.
- Venkateswaran V, Klotz LH, Ramani M, Sugar LM, Jacob LE, Nam RK, Fleshner NE. A combination of 31. micronutrients is beneficial in reducing the incidence of prostate cancer and increasing survival in the Lady transgenic model. Cancer Prev Res (Phila) 2009; 2: 473-483.
- Giovannucci E, Liu Y, Platz EA, Stampfer MJ, Willett WC. Risk factors for prostate cancer incidence and 32. progression in the health professional's follow-up study. Int J Cancer. 2007;121: 1571–1578.
- Demark-Wahnefried W, Moyad MA. Dietary intervention in the management of prostate cancer. Curr Opin Urol. 33. 2007: 17:168-174.
- 34. Mcbride RB. Obesity and aggressive prostate cancer bias and biomarkers. Columbia University; 2012.
- Kaaks R, Stattin P. Obesity, endogenous hormone metabolism, and prostate cancer risk: a conundrum of "highs" and 35. "lows". Cancer Prev Res (Phila) 2010; 3:259–262.
- Keogh JW, MacLeod RD. Body composition, physical fitness, functional performance, quality of life, and fatigue 36. benefits of exercise for prostate cancer patients: a systematic review. J Pain Symptom Manage. 2012; 43: 96–110.