

ASSOCIATION OF ETHNICITY AND CARCINOGEN WITH HEAD AND NECK CANCERS

Dr Sana Aziz^{*1}, Dr Sohail Aslam², Dr Adnan Asghar³, Shahzad Maqbool⁴,
Fatima Siddiqui⁵, Sana Sainch⁶

^{*1}ENT Resident PNS Shifa Hospital And BUHSCK

²ENT Dept HOD of ENT & Head & Neck Surgery, CMH Sialkot.

³Ent Associate Professor PNS Hafeez Hospital Islamabad

⁴ENT HOD ENT & Head and Neck Surgery PNS Shifa Hospital Karachi

⁵Ent Assistant Professor PNS Shifa Hospital Karachi

⁶ENT Resident PNS Shifa Hospital Karachi and BUHSCK

^{*1}Sanaazizkhan18@gmail.com, ²aslamsohail07@gmail.com, ³dradnanasghar@gmail.com,
⁴shahzadmaqbool73@gmail.com, ⁵Fatima.siddi7@gmail.com, ⁶sainchsana@gmail.com

ABSTRACT

Objective: The purpose of this study is to review the complex relationship between the risk of head and neck cancer, exposure to carcinogens, and ethnicity.

Study design: Cross sectional study

Place and duration of study: This study is performed at ENT department of PNS Shifa hospital Karachi from January 2023 to July 2023

Methodology: A group of 220 patients who were presented at the outdoor department were selected after analyzing their eligibility on the basis of predefined criteria of inclusion and exclusion. All the patients with age between 18 to 85 years of either gender were included in this study.

Results: The study included individuals with a mean age of 51.0 years (\pm SD 11.4), comprising 62.27% males and 37.73% females. The ethnic breakdown of biopsy-confirmed head and neck carcinoma cases revealed that 49.55% identifying as Urdu and varying percentages for Sindhi, Pashton, Balochi, Punjabi, and others. The carcinogen exposure data indicates significant usage of betel nuts (70.9%), gutka (65%), pan (79%), tobacco (83.6%), and naswar (74.09%) among participants, emphasizing the prevalence of various risk factors within the study population.

Conclusion: This study has concluded that the incidence of head and neck of carcinomas has strong association with ethnicity and the exposure to various carcinogens like betel nut, tobacco, Naswar and gutka.

Keywords: Ethnicity, head and neck carcinoma, carcinogens

INTRODUCTION

A broad category of tumors that impact different parts of the body, such as the oral cavity, pharynx, larynx, and paranasal sinuses, is referred to as head and neck cancers (HNC) ¹. A combination of hereditary and environmental variables is major contributors to the complex etiology of HNC². Two important factors that have been shown to affect the occurrence, course, and consequences of HNC are ethnicity and exposure to carcinogens. In order to better understand the relationship between ethnicity, carcinogen exposure, and the risk of head and neck cancers, a literature review has been conducted³.

The Research of Medical Science Review

The prevalence and death rates of HNC vary between ethnic groups, as research repeatedly reveals. Certain ethnic groups, such as those descended from South Asia, may be more susceptible to particular kinds of head and neck malignancies, according to studies ⁴. The observed inequalities in cancer epidemiology can be attributed to a combination of socioeconomic variables, cultural behaviors, and genetic predispositions ⁵. Therefore, it is imperative to take ethnicity into account as a major determinant.

The development and spread of head and neck cancers are significantly influenced by carcinogens, both environmental and lifestyle-related. Consumption of alcohol and tobacco products is a known risk factor, and there is a positive correlation between the two in those who use them together ⁶. Furthermore, it has come to light that a major carcinogenic factor is human papillomavirus (HPV) infection, especially in cases of oropharyngeal malignancies. The etiological landscape is made more complicated by occupational exposures, such as those in businesses that use asbestos and certain chemicals ⁷.

To comprehend the varying susceptibilities to HNC, researchers have focused on the interaction between ethnicity and carcinogen metabolism ⁸. The metabolism of tobacco-related carcinogens involves enzymes such as cytochrome P450, which contain genetic variations that have been linked to different ethnic groups ⁹. Ethnic differences in HNC risk may be influenced by these variables, which may also affect an individual's reaction to carcinogen exposure ¹⁰.

Strategic preventative and intervention measures are essential, given the complex link that exists between ethnicity, exposure to carcinogens, and HNC ¹¹. Reducing the prevalence of HNC can be facilitated by culturally sensitive public health campaigns, smoking cessation initiatives, and HPV vaccination campaigns, especially in high-risk ethnic regions ¹². The development of tailored strategies for cancer prevention can also benefit from knowledge of the genetic factors of carcinogen metabolism ¹³.

This review of the research emphasizes the complex relationship between the risk of head and neck cancer, exposure to carcinogens, and ethnicity. In order to create preventative, early detection, and intervention techniques that work, it is imperative to acknowledge these intricacies ¹⁴. To identify the precise genetic and environmental variables causing ethnic differences in HNC and to provide tailored therapies for groups that are at risk, more study is necessary.

Methodology

It is a cross-sectional study performed at the ENT department of PNS Shifa hospital Karachi from January 2023 to July 2023. The approval of the ethical review committee (ERC/2023/ENT/61) was taken before the study. A group of 220 patients who were presented at the outdoor department were selected after analyzing their eligibility on the basis of predefined criteria of inclusion and exclusion. All the patients with age between 18 to 85 years of either gender were included in this study. All the biopsy-proven cases were included in the study and any patient with co-morbidity; any other malignancy or memory issues were excluded from this study. Similarly, patients with incomplete medical records and with known genetic conditions or syndromes that increase the risk of head and neck carcinomas were not included in this study. Individuals who had recently undergone any treatment for head and neck carcinoma were also excluded to avoid any confounding effects. All the patients were briefed about the methods involved in the study and informed written consent was taken. Similarly, history was taken from each patient on the basis of a pre-designed Proforma with information about use of any carcinogen like tobacco, betel nut, gutka, pan and naswar, duration of addiction and amount of consumption. The address details were also noted for each patient along with biopsy report, staging of carcinoma and history of any treatment taken.

The data analysis was conducted using the Statistical Package for Social Sciences (SPSS) version 23.00. For quantitative variables, the calculations were mean and standard deviation; for qualitative variables, the calculations were frequencies and percentages. The p-value of less than or equal to 0.5 was considered significant.

Result

The results of this study have shown that individuals with a mean age of 51.0 years (\pm SD 11.4) made up the study population. Gender distribution showed that 62.27% of the population was male and 37.73% was

The Research of Medical Science Review

female. The ethnic composition of the population who were biopsy confirmed cases of head and neck carcinomas was varied; 49.55% identified as Urdu, while other ethnic groups included Sindhi (19.55%), Pashton (9.55%), Balochi (8.18%), Punjabi (5.45%), and others (7.73%) with a p value of 0.02 as shown in table I and figure I. This data gives a thorough overview of the demographic mix and emphasizes how different ethnic groups are represented in the research. Such data must be included in order to comprehend any differences in health outcomes across various ethnic groups and to enable a more sophisticated interpretation of the study's conclusions.

The distribution of biopsy-confirmed cases across different anatomical areas is varied, according to the statistics on head and neck carcinomas. With 44.09% of the 220 cases, the buccal mucosa was the most common location, followed by the cheek with 37.72% of cases involving carcinoma (CA). Additional locations included the floor of the mouth (1.37%), the hard palate (0.45%), the larynx (5.0%), and the tongue (11.36%) (P value 0.01). The study population's carcinomas' TNM staging provides additional insight into the kind and severity of cancer cases. Interestingly, most instances were found at stages 3 (48.64%) and 4 (35.91%) as shown in table II, indicating that many of these malignancies were advanced when they were discovered.

The tabulated data on exposure to carcinogens provide important information on the prevalence of different risk factors in the research population. A significant percentage of individuals reported using betel nuts, with 70.9% saying "Yes" and 29.1% saying "No." Of those, 65% reported using betel nuts, while 35% reported not using them. 79% of subjects reported using pan, a mixture of areca nut and betel leaf; whereas 21% said they did not. Of the participants, 83.6% acknowledged using tobacco, a known carcinogen. Only 17.4% said they did not use tobacco. Of the participants, 74.09% reported using naswar, while 25.91% said they did not use it (P value 0.03) as shown in table III and figure II.

Discussion

The data collected gives a thorough overview of the demographic mix and emphasizes how different ethnic groups are represented in the research and their relationship with incidence and frequency of head and neck carcinoma. Such data must be included in order to comprehend any differences in health outcomes across various ethnic groups and to enable a more sophisticated interpretation of the study's conclusions¹⁵. This analysis has shown that head and neck carcinomas were most common in Urdu speaking ethnic groups followed by Sindhi, Pashton, Balochi and other groups.

Comprehending the stage distribution plays a crucial role in prognostic assessments and in customizing suitable treatment strategies for individual patients¹⁶. By shedding light on the tumors' histological features, this data advances our knowledge of their biological makeup and their aggressiveness. In conclusion, the thorough information provided on biopsy-confirmed cases, TNM staging, and degree of differentiation clarifies the complex characteristics of head and neck carcinomas in the population under study, enabling researchers and clinicians to have a more sophisticated understanding of the illness.

Given that betel nut use is common in this demographic and it has been linked to cancer, public health initiatives and awareness campaigns should take this into serious consideration¹⁷. In light of the established health hazards linked to gutka, including as a heightened susceptibility to oral malignancies, our findings emphasize the significance of focused prevention efforts and intervention programs for those abusing this detrimental agent¹⁸. Comparably, the study's inclusion of Pan emphasizes the complex relationship between carcinogen exposure and the necessity for all-encompassing approaches to address the different cultural practices that raise the risk of cancer. The high prevalence highlights the ongoing difficulty of tobacco-related health problems and highlights the importance of smoking cessation programs and anti-smoking activities in this demographic¹⁹. The high frequency of Naswar use underscores the variety of tobacco exposure scenarios in this demographic and underscores the significance of comprehensive tobacco control strategies that address both smokeless and smoked tobacco products²⁰.

To sum up, this study on carcinogen exposure offers a thorough picture of the frequency of different risk variables in people who have head and neck cancers. The results highlight the critical need for focused

The Research of Medical Science Review

public health initiatives to lower the incidence of cancer, address these risk factors, and enhance general population health.

Conclusion:

This study has concluded that the incidence of head and neck of carcinomas has strong association with ethnicity and the exposure to various carcinogens like betel nut, tobacco, Naswar and gutka. The findings underscore the urgent need for targeted public health interventions to mitigate these risk factors, reduce cancer incidence, and improve overall community health.

Limitations

The study's findings may have limitations in terms of generalizability due to the specific demographic and geographic characteristics of the study population. The reliance on self-reported data for certain variables, such as carcinogen exposure, introduces the potential for recall bias. Despite efforts to control for confounding variables, there may be unmeasured factors that could influence the observed association between ethnicity, carcinogen exposure, and head and neck cancers. Variables such as genetic predispositions, occupational exposures, and lifestyle factors may not have been adequately accounted for in the study. Hence more thorough and exhaustive investigation on this issue is the need of the hour.

Conflict of interest

None

Acknowledgement

To doctors and staff of PNS al-shifa hospital

Table I: Demography and Ethnicity of population

Variables	Result
Age in years (mean \pm SD)	51. \pm 11.4
Gender	
Male	137 (62.27%)
Female	83 (37.73%)
Ethnicity	
Sindhi	43(19.55 %)
Balochi	18(8.18%)
Pashto	21 (9.55%)
Punjabi	12 (5.45%)
Urdu	109 (49.55%)
Others	17(7.73%)

The Research of Medical Science Review

Figure I: Ethnicity of study population

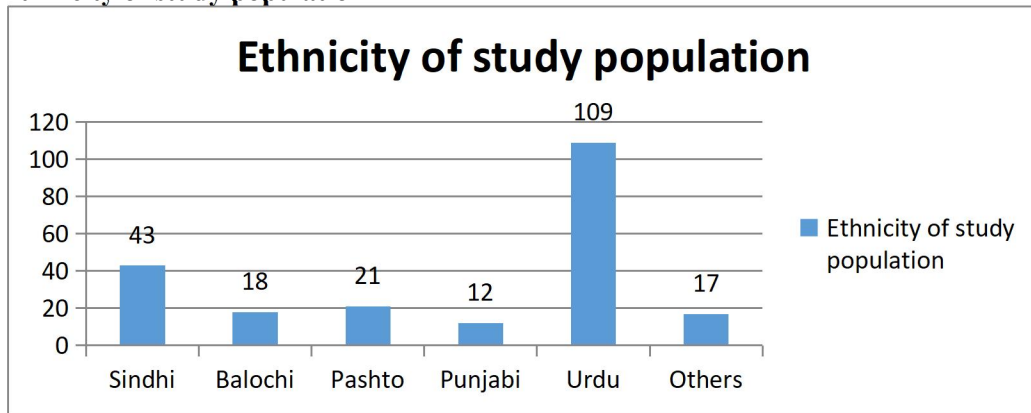


Table II: analysis of head and neck carcinomas

Variables	Frequency
Biopsy confirmed cases (n=220)	
Buccal mucosa	97 (44.09%)
CA cheek	83 (37.72 %)
CA larynx	11 (5.0%)
Floor of the mouth	3 (1.37%)
Hard palate CA	1 (0.45%)
CA tongue	25 (11.36%)
TNM Staging of the CA (n=220)	
2	34 (15.45%)
3	107(48.64%)
4	79 (35.91%)
Degree of differentiation	
Well differentiated	71 (32.27%)
Moderately differentiated	103 (46.82%)
Poorly differentiated	46 (20.91%)

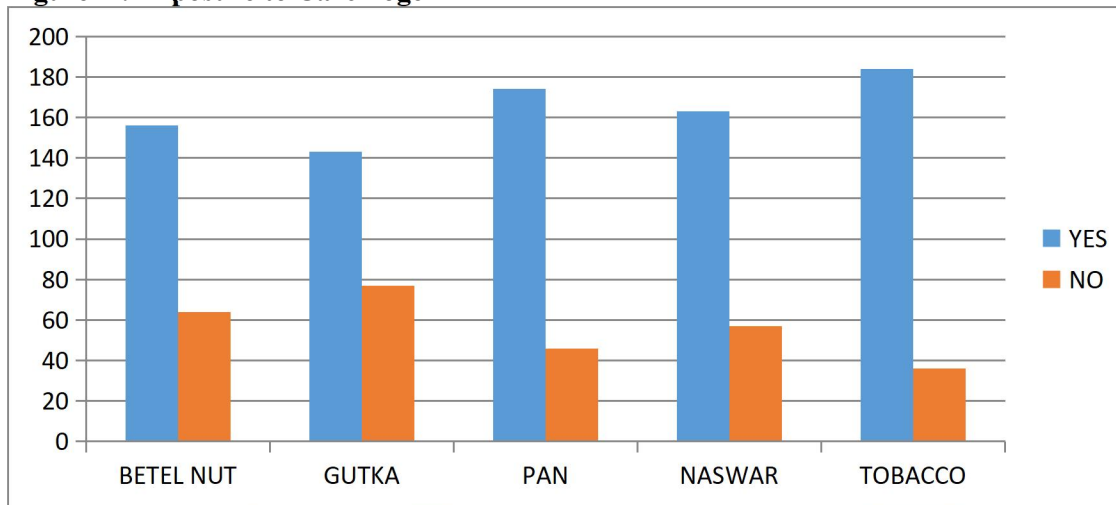
Table III: Carcinogen Exposure

Carcinogen exposure	
Variable	Frequency (n=220)
Betel nut	
• Yes	156(70.9%)
• No	64 (29.1%)
Gutka	
• Yes	143(65%)
• No	77(35%)
Pan	
• Yes	174(79%)
• No	46(21%)

The Research of Medical Science Review

• No	
Tobacco	
• Yes	184 (83.6%)
• No	36 (17.4%)
Naswar	
• Yes	163(74.09%)
• No	57(25.91%)

Figure II: Exposure to Carcinogen



REFERENCES:

- Ragin C, Liu JC, Jones G, Shoyele O, Sowunmi B, Kennett R, Gibbs D, Blackman E, Esan M, Brandwein MS, Devarajan K. Prevalence of HPV infection in racial–ethnic subgroups of head and neck cancer patients. *Carcinogenesis*. 2017 Feb 1; 38(2):218-29.
- Tataru D, Mak V, Simo R, Davies EA, Gallagher JE. Trends in the epidemiology of head and neck cancer in London. *Clinical Otolaryngology*. 2017 Feb; 42(1):104-14.
- Noel CW, Sutradhar R, Li Q, Forner D, Hallet J, Cheung M, Singh S, Coburn NG, Eskander A. Association of immigration status and Chinese and south Asian ethnicity with incidence of head and neck cancer. *JAMA Otolaryngology–Head & Neck Surgery*. 2020 Dec 1; 146(12):1125-35.
- Zhang L, Xiang Z, Hao R, Li R, Zhu Y. N-acetyltransferase 2 genetic variants confer the susceptibility to head and neck carcinoma: Evidence from 23 case–control studies. *Tumor Biology*. 2014 Apr; 35: 3585-95.
- Clarke JA, Despotis AM, Ramirez RJ, Zevallos JP, Mazul AL. Head and neck cancer survival disparities by race and rural–urban context. *Cancer Epidemiology, Biomarkers & Prevention*. 2020 Oct 1; 29(10):1955-61.
- McDermott JD, Eguchi M, Morgan R, Amini A, Goddard JA, Borrayo EA, Karam SD. Elderly black non-Hispanic patients with head and neck squamous cell cancer have the worst survival outcomes. *Journal of the National Comprehensive Cancer Network*. 2020 Sep 28; 19(1):57-67.
- Lee YC, Al-Temimi M, Ying J, Muscat J, Olshan AF, Zevallos JP, Winn DM, Li G, Sturgis EM, Morgenstern H, Zhang ZF. Risk prediction models for head and neck cancer in the US population from the INHANCE consortium. *American journal of epidemiology*. 2020 Apr 2; 189(4):330-42.
- Jia YQ, Chen XC, Chen YM, Zheng YY, Yang B. Prognostic Value of Inflammasomes in Head and Neck Carcinoma: A Meta-analysis.

The Research of Medical Science Review

- Di Credico G, Polesel J, Dal Maso L, Pauli F, Torelli N, Luce D, Radoi L, Matsuo K, Serraino D, Brennan P, Holcatova I. Alcohol drinking and head and neck cancer risk: the joint effect of intensity and duration. *British journal of cancer*. 2020 Oct 27; 123(9):1456-63.
- Mazul AL, Chidambaram S, Zevallos JP, Massa ST. Disparities in head and neck cancer incidence and trends by race/ethnicity and sex. *Head & Neck*. 2023 Jan; 45(1):75-84.
- Cho HM, Lee DW, Park JJ, Choi HM, Cho NP. Risk factors of head and neck cancer mortality compared with those of all-cause and all-cancer mortalities. *Journal of Oral Science*. 2020; 62(4):365-70.
- Rehman, M.Y.A., Taqi, M.M., Hussain, I., Nasir, J., Rizvi, S.H.H. and Syed, J.H., 2020. Elevated exposure to polycyclic aromatic hydrocarbons (PAHs) may trigger cancers in Pakistan: an environmental, occupational, and genetic perspective. *Environmental Science and Pollution Research*, 27, pp.42405-42423.
- Mohammadi H, Roochi MM, Sadeghi M, Garajei A, Heidar H, Ghaderi B, Tadakamadla J, Meybodi AA, Dallband M, Mostafavi S, Mostafavi M. Association of n-acetyltransferases 1 and 2 polymorphisms with susceptibility to head and neck cancers—a meta-analysis, meta-regression, and trial sequential analysis. *Medicina*. 2021 Oct 13; 57(10):1095.
- Day AT, Dahlstrom KR, Lee R, Karam-Hage M, Sturgis EM. Impact of a tobacco treatment program on abstinence and survival rates among current smokers with head and neck squamous cell carcinoma. *Head & neck*. 2020 Sep; 42(9):2440-52.
- Gnagnarella P, Raimondi S, Aristarco V, Johansson HA, Bellerba F, Corso F, Gandini S. Vitamin D receptor polymorphisms and cancer. *Sunlight, Vitamin D and Skin Cancer*. 2020:53-114.
- Wang X, Wang H, Zhang T, Cai L, Dai E, He J. Diabetes and its potential impact on head and neck oncogenesis. *Journal of Cancer*. 2020; 11(3):583.
- Naidoo K, Simonds H, Ebrahim AK, vAn Rensburg LJ, Merven M, Opperman J, Afrogheh A. A Descriptive Epidemiological Study of Head and Neck cancers at a Major Referral Center in Southern Africa. *Authorea Preprints*. 2020 May 11.
- Ibrahimovic M, Franzmann E, Mondul AM, Weh KM, Howard C, Hu JJ, Goodwin WJ, Kresty LA. Disparities in Head and Neck Cancer: A Case for Chemoprevention with Vitamin D. *Nutrients*. 2020 Aug 29; 12(9):2638.
- Ramezani M, Sadeghi M, Zavattaro E, Mozaffari HR. Association between CYP1A1 Ile462Val (m2, A2455G, rs1048943) polymorphism and head and neck cancer susceptibility: A meta-analysis, meta-regression, and trial sequential analysis. *Gene Reports*. 2021 Dec 1; 25: 101380.
- Kabagenyi, F., Otiti, J., Namwagala, J., Kamulegeya, A. and Kalungi, S., 2020. A descriptive study of human papilloma virus in upper aero-digestive squamous cell carcinoma at Uganda cancer institute assessed by P16 immunohistochemistry. *Cancers of the Head & Neck*, 5, pp.1-9.

CONSENT FORM

I agree to participate in the study titled:

ASSOCIATION OF ETHNICITY AND CARCINOGEN WITH HEAD AND NECK CANCERS

I have been informed about the study's possible benefits and side effects. I understand that my participation is voluntary, and I am free to withdraw from the study whenever I want to I have been told that my doctor will continue to give me all possible care and treatment even if I start the study.

I allow my doctor to take my blood sample to perform tests on it for research purposes. Furthermore, I allow my doctor or any other person authorized by my doctor to contact me at my home or at an address given by me for treatment and follow to be used for research purposes. I have been assured that the information I provided will be kept confidential and used for research purposes only.

Patient Name: _____ Witness Name _____
CNIC#: _____ CNIC#: _____
Signature: _____ Signature _____
Date: _____ Date: _____

The Research of Medical Science Review

Doctor's Signature: _____
Date: _____

CONSENT FORM

رضاء مندی نامہ برائے شمولیت تحقیق

ASSOCIATION OF ETHNICITY AND CARCINOGEN WITH HEAD AND NECK CANCERS

ڈاکٹر نے مجھے اس طبی تحقیق کے بارے میں تفصیلاً آگاہ کر دیا ہے اور میں اس تحقیق میں شامل ہونے کے لیے رضامند ہوں۔ مجھے بتا دیا گیا ہے کہ میری ذاتی اور طبی معلومات کے علاوہ مجھے اپنے خون کے نمونے فراہم کرنا ہوں گے جن پر کچھ لیبارٹری ٹیسٹ کئے جائیں گے۔
میں جانتا/جانتی ہوں کہ اس تحقیق میں شمولیت سے مجھ پر کوئی خطرناک اثرات مرتب نہیں ہوں گے۔
میں رضاکارانہ طور پر اس تحقیق کا حصہ بن رہا/ رہی ہوں۔ میں کسی بھی وقت اس تحقیق سے قطع تعلق اختیار کر سکتا/کرسکتی ہوں۔ اور میرے علاج معالجہ پر اس عمل سے کوئی اثر نہیں پڑے گا۔
مجھے بتا دیا گیا ہے کہ اس تحقیق سے مجھے کوئی مالی فائدہ حاصل نہیں ہوگا۔
علاوہ ازیں میں یہ بات اچھی طرح سمجھتا / سمجھتی ہوں کہ میری ذاتی اور طبی معلومات کو محفوظ اور صیغہ راز میں رکھا جائے گا۔

مریض کا نام: _____ گواہ کا نام: _____
شناختی کارڈ نمبر: _____ : شناختی کارڈ نمبر: _____
دستخط: _____ : دستخط: _____
تاریخ: _____ : تاریخ: _____
دستخط معالج: _____ : تاریخ: _____

ASSOCIATION OF ETHNICITY AND CARCINOGEN WITH HEAD AND NECK CANCERS

BIODATA

NAME: _____ S/O, D/O, W/O: _____
AGE: _____ GENDER: M/F PHONE: _____
COMORBIDS: _____
ADDRESS: _____
PERMANENT: _____
TEMPORARY: _____
DURATION: _____

HISTORY

CARCINOGEN:

- 1.
- 2.
- 3.

DURATION OF ADDICTION:

- 1.
- 2.
- 3.

CONSUMPTION

1. PER DAY

The Research of Medical Science Review

2. ALTERNATE DAY
3. WEEKLY

INVESTIGATION:

1. BIOPSY: _____

2. CT SCAN: _____

DIAGNOSIS: _____

STAGING TNM: _____

SURGERY:

WLE

FROZEN SECTION

REPAIR

NECK DISSECTION

DATE OF SURGERY

SURGICAL COMPLICATIONS: _____

DURATION OF COMPLICATION: _____

TREATMENT GIVEN: _____

POST-OP HISTOPATHOLOGY: _____

_____ FOLLOW UP:

CHEMO/RADIOTHERAPY STATUS: _____

NO OF CYCLES

CHEMOTHERAPY DRUGS

COMPLICATIONS OF RADIOTHERAPY: _____
