

## Conference Paper

# Neck Dissection in Patients with Oral Squamous Cell Carcinoma in Dharmais Hospital, Jakarta

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## Abstract

Oral squamous cell carcinoma (OSCC) is one of the rarest cancers worldwide. In Indonesia, the incidence is less than five thousands per year, and the mortality rate is almost 50%. More than 50% patients with OSCC have lymph node metastasis; the proportion of occult metastasis is 24-42%. Those with lymph node metastasis have the worst possibility of survival. This study aimed to estimate the survival of OSCC patients with neck dissections. We conducted a retrospective cohort study of 78 patients with OSCC who were treated in Dharmais National Cancer Hospital between 1 January 2003 and 31 January 2013. The three years survival rate post diagnosis, post neck dissection was calculated using Kaplan –Meier survival curves and statistically tested using a log-rank test. Cox proportional hazard models were applied to assess the prognostic significance of neck dissections. Of the total patients in this study (n=78), 53.8% of patients had surgery. Of patients who underwent surgery, 71.4% had a neck dissection surgery. These patients were in either early or advanced stages of cancer. Overall survival showed that patients who received neck dissections had better survival rates (58.2%) than patients who did not receive neck dissection (32.2%). Stratification at every stage of cancer (I, II, III, and IV) showed better survival in patients with neck dissections. The risk of patients without neck dissections is higher than patient with neck dissection to die, Hazard Ratio(HR)=2.19 (CI95% 1.04-4.62, p=0.028). Adequate neck dissection surgery increases chances of survival in patients with OSCC.

**Keywords:** Neck Dissection, Squamous Cell Carcinoma, Oral Cancer, Survival

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## 1. INTRODUCTION

Therapies can improve survival in those with oral squamous cell carcinoma (OSCC). Surgery is a successful cancer therapy with neck dissection often yielding satisfactory results. However, therapy fails for primary tumors in almost half of cases despite 90% of patients having a clear margin [2]. Other therapies are adjuvant radiotherapy and chemotherapy. Patients with metastatic lymph nodes have a recurrence risk of 62% and a survival rate of 27% if they are not given adjuvant radiotherapy after surgery [6].

Removing lymph nodes is an important factor to achieve a positive therapy result in OSCC. However, the benefit of a neck dissection is still unknown, because neck dissections mostly happen for advanced stages [3]. This study aimed to estimate the survival of OSCC patients with neck dissections.

## 2. METHODS

### 2.1. Study Design

This paper was part of a larger study on HPV, P16, EGFR and VEGF expression and margin status and neck dissection as prognostic factors of three years survival patients post diagnosis with OSCC in Dharmais Hospital. After ethical approval by the Committee of Medical Research Ethics of the Dharmais National Cancer Hospital (No. KEPK/041/X/2015) and the Committee of Ethical Research Public Health (No. 250/UN2.F10/PPM.00.02/2015), we performed a retrospective cohort study. A total 78 patients were assessed. Patients who were eligible must have had oral cancer and squamous carcinoma cell confirmed by a histology examination. Patients whose cancer stage could not be determined and who did not have formalin-fixed, paraffin-embedded (FFPE) tissues available were excluded from the study.

### 2.2. Statistical analysis

The study design was to determine the three year overall survival post diagnosis of OSCC patients in Dharmais National Cancer Hospital. Independent variables included age, sex, stage, differentiation, surgery, neck dissection, margin status, chemotherapy

and radiotherapy. The comparison of OSCC patient survival was described by a Kaplan–Meier curve and significance was determined using a log rank test. A hazard ratio (HR) was calculated using a Cox semiparametric proportional hazards model.

### 3. RESULTS

From 2003 to 2013, we had analyzed 78 patients from 266 patients. The distribution of demographic data is shown in Table 1. The average age was 47 years old (range 19–78 years old). Patients over 40 years old made up 64.1% of the sample and were mostly male. The proportion of advance stage OSCC was 73.1% with well differentiation of 59.0%. There were 42 patients who received surgery, most of whom (30/42) received a neck dissection. The remaining patients received chemotherapy (47.4%) or radiotherapy (50%).

Kaplan–Meier estimated the three years overall survival post diagnosis was 42.5% (Fig.1). Of the 78 patients, three year survival patients post diagnosis with neck dissection was 58.2% and survival with no neck dissection was 32.2%. The survival rate of those with neck dissections was significantly higher ( $p=0.030$ ). In this group, the HR was 2.197 (95% CI 1.046-4.615). When stratified by surgery status ( $n=42$ ), the group that received neck dissections survived longer than the group that did not receive neck dissections, with 58.2% and 34.4% survival rate respectively, but it's not statistically significant ( $p=0.276$ ). The HR was 1.760 (95% CI 0.636-4.870). After adjusting the variables age, sex, differentiation, chemotherapy and radiotherapy, the multivariate analysis (Table 3) showed that the neck dissection group had better survival rates than the group with no neck dissections (HR = 1.795) (table 2).

### 4. DISCUSSION

Surgery is the main therapy to treat solid tumors. According to the Indonesia Society of Surgical Oncologist [4], every surgery should reach radical neck dissection and free margin. Over the years, there have been controversies about whether radical neck dissection should or should not be given to patients. A study by Wang et al. (2013) showed that patients with lymph node dissection, adjuvant chemotherapy or local excision, had a better prognosis compared with patients with no lymph node dissection [5]. Yuen et al. (1997) found that patients with elective neck dissections had an increased survival rate compared to those without [7]. D'Cruz et al. (2015) found that patients with early stage cancer who elected for radical neck dissection had a 12.5%

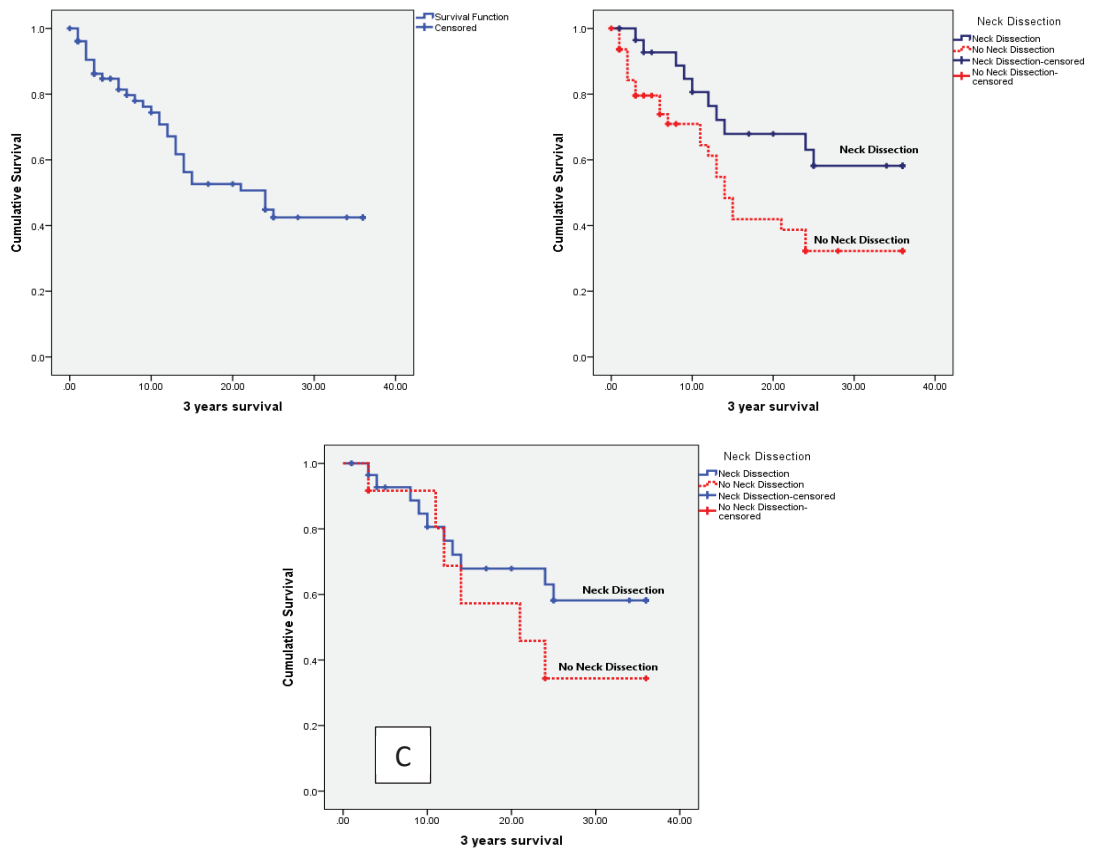
TABLE 1: Characteristics of patient with OSCC in Dharmais Hospital.

Variable	F	%
Age		
≤ 40 years	28	35.6
> 40 years	50	64.1
Sex		
Male	47	60.3
Female	31	39.7
Stage		
I-II	21	26.9
III-IV	57	73.1
Differentiation		
Good	46	59.0
Poor	31	39.7
Unknown	1	1.3
Surgery		
Yes	42	53.8
No	36	46.2
Neck Dissection		
Yes	30	38.5
No	48	61.5
Margin Status		
Free Margin	37	47.4
Close Margin	41	52.6
Chemotherapy		
Yes	37	47.4
No	41	52.6
Radiotherapy		
Yes	39	50
No	39	50

TABLE 2: Univariate Cox semiparametric proportional hazard model.

Variable	Hazard ratio (95% CI)	p value
Neck Dissection (n=78)		
Yes	1.00	
No	2.197(1.046-4.615)	0.038
Neck Dissection (n=42)		
Yes	1.00	
No	1.760 (0.636-4.870)	0.276

better chance of survival, a reduced risk of death (by 36%), and a reduced recurrence rate (by 55%) compared with patients with therapeutic neck dissections [1]. Our study



**Figure 1:** (A) Overall survival curve. (B) Survival curve for OSCC patients (n=78). (C) Survival curve for OSCC patients after stratified by surgery (n=42).

had the same results as these studies. Patients with neck dissections had significantly higher survival rates. However, patients with early stages of cancer (both I and II) had the same results. The group that got neck dissection surgery had better survival.

TABLE 3: Multivariate Cox regression overall survival.

Variables	Hazard ratio CI95%	p value
Sex (Male vs Female)	0.530 (0.242-1.159)	0.112
Neck Dissection (Yes vs No)	0.557 (0.249-1.246)	0.154
Age (<=40 vs >40)	2.122 (0.975-4.622)	0.058
Differentiation (poor vs good)	1.672 (0.837-3.340)	0.145
Radiotherapy (yes vs no)	0.445 (0.215-0.920)	0.029
Chemotherapy (yes vs no)	1.925 (0.891-4.157)	0.096

Other studies have found that more than 50% patients with OSCC have metastatic lymph nodes positively confirmed by histology. Metastatic lymph nodes are poor prognostic factors in OSCC. In many cases also found occult metastatic lymph node, the numbers vary from 24-42%. The therapy for this condition was radical neck dissection with lymph node level I-IV [2].

Due to the controversies regarding the efficacy of neck dissection, more studies are needed to assess the proper therapy in oral cancer develop. Many researchers suggest a sentinel node biopsy to be used in oral cavity cancer. While this method has a less clear interpretation compared to applied melanoma, the technique is promising. The advantage of sentinel node biopsy is a reduction of morbidity in the absence of a neck dissection, and the technique has proven worthwhile and accurate [3].

## 5. CONCLUSIONS

We suggested that OSCC patients should be treated with neck dissections to improve likelihood of survival. In the future, the combination of neck dissection and sentinel node biopsy will lead to a better survival and reduce morbidity.

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