Rate of Occult Cervical Lymph Node Involvement in Supraglottic Squamous Cell Carcinoma

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Abstract
Introduction:
To assess the rate of cervical lymph node involvement in patients with supraglottic squamous cell carcinoma (SCC) with no lymph node in clinical assessments and radiological studies.

Materials and Methods:
Fifty-six patients who underwent elective dissection of the cervical lymph node of the second through fourth level were enrolled, and pathologic evaluation of the dissected lymph nodes was performed. Lymph node involvement and association between tumor grade, smoking and gender with lymph node involvement were assessed.

Results:
The rate of the occult neck metastasis in this series was 37.5%. There was no statistically significant association between lymph node involvement and tumor grade, smoking, or gender.

Conclusion:
Based on the results of our study, we recommend elective bilateral neck dissection in all stages of N0 supraglottic SCC patients.

Keywords:
Laryngeal cancer, Lymph nodes, Metastasis, Neck dissection, Squamous cell carcinoma.

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Introduction
Squamous cell carcinoma (SCC) is the most common cancer of the larynx and accounts for 90% of malignant laryngeal tumors. Supraglottic SCC after glottic SCC is the second most common malignant tumor of the larynx (1,2). The male-to-female (M/F) ratio is 3.8:1 for all laryngeal cancers, and more than 90% of cancers occur in patients over 40 years of age (3).

The two most important risk factors for all laryngeal cancers are smoking and alcohol consumption, and more than 97% of patients have a history of cigarette smoking (1,4). Other risk factors include laryngopharyngeal reflux, human papillomavirus (HPV), and toxins. Because of its rich network of lymphatics, supraglottic SCC has the most cervical lymph node metastasis among laryngeal cancers, and usually metastasizes to cervical lymph node levels II, III, and IV (5). The rate of pathologically involved lymph nodes in patients with supraglottic SCC who have no lymph node involvement in clinical assessments and radiological studies (occult lymph node metastasis) differs across studies and varies from 12% to 40%. On the other hand, the most important prognostic factor of supraglottic SCC is the presence of lymphatic metastasis, and 5-year survival rate decreases to 50% when there is lymphatic metastasis (6–8).

The classic approach to supraglottic SCC with no clinical or radiological cervical lymph node metastasis (N0) is surgical treatment with bilateral neck dissection of levels II, III, and IV. If the risk of occult cervical lymph node metastasis is statistically more than 20%, lymph node dissection is necessary because neck dissection in less than 20% metastasis imposes potential complications such as nerve and vessel injuries and bleeding and increases the time of surgery.

Several studies reported different rates of occult cervical lymph node metastasis (9–11), and some of these studies suggest that there was no need for neck dissection in patients with T1 N0 supraglottic SCC. The aim of this study was to assess the rate of occult cervical lymph node metastasis in N0 supraglottic SCC in the Middle East population, especially in the lower stages of laryngeal cancer, in order to potentially prevent unnecessary neck dissection in patients referred for supraglottic SCC treatment.

Materials and Methods
This was a retrospective study. Patients undergoing level II through IV elective neck dissection for supraglottic SCC between 2002 and 2014 were enrolled. The study was conducted at the tertiary center of Imam Khomeini Hospital Complex, Valiasr Hospital at Tehran University of Medical Sciences. All patients were classified as N0, meaning that they had no cervical lymph node metastasis in the clinical and radiological evaluation.

After meeting the inclusion criteria, 56 patients were enrolled. Histopathological examination of all samples was performed at the Cancer Institute of Imam Khomeini complex. Grading of tumors was based on direct laryngoscopy and biopsy, imaging and pathologic reports (T1–T4). Exclusion criteria were incomplete patient profile and the pathology reports from other centers. The associations between lymph node involvement and tumor grade, smoking, and gender were also assessed.

The IBM SPSS statistics package version 22 was used for analysis. A Chi-square test was used for the analysis of the associations between qualitative variables.

The study was approved by the Ethics Committee of the Tehran University of Medical Sciences and the National Medical Ethics Committee according to the principles of the Declaration of Helsinki.

Results
The mean age of patients was 57.6 ± 11 years. The M/F ratio was 27:1 (54:2). There was no statistically significant association between cervical lymph node metastasis and gender (chi-square test [n=56]; p=0.710).

Patients were classified into four groups based on the amount of cigarette smoking per year (less than 20 pack year [p/y], 20–40 p/y, 40–60 p/y, and more than 60 p/y). Most patients (42.9%) were in the 40–60 p/y group. There was no statistically significant association between histopathological involvement and cigarette smoking (chi-square test [n=56]; p=0.854). There was no history of alcohol consumption in our patients. Twenty-one of 56 patients had cervical lymph node involvement based on histopathologic evaluation, so the rate of occult
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cervical metastasis was 37.5% in this series. Ten of 21 patients (47.6%) with cervical lymph node metastasis had bilateral neck involvement.

Cervical lymph node metastasis based on tumor grade is summarized in Table 1. Most patients were in the T3 group (44.6%). The association between tumor grade and lymph node involvement was assessed and showed no statistically significant association between tumor grade and cervical lymph node metastasis (chi-square test [n=56]; \( P=0.389 \)).

<table>
<thead>
<tr>
<th>Tumor Grade</th>
<th>Lymph Node Involvement</th>
<th>Total</th>
<th>Percentage</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Yes (%)</td>
<td>No (%)</td>
<td></td>
</tr>
<tr>
<td>T1</td>
<td>1 (16.7)</td>
<td>5 (83.3)</td>
<td>6</td>
</tr>
<tr>
<td>T2</td>
<td>4 (28.6)</td>
<td>10 (71.4)</td>
<td>14</td>
</tr>
<tr>
<td>T3</td>
<td>10 (40)</td>
<td>15 (60)</td>
<td>25</td>
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<tr>
<td>T4</td>
<td>6 (54.5)</td>
<td>5 (45.5)</td>
<td>11</td>
</tr>
<tr>
<td>Total</td>
<td>21</td>
<td>35</td>
<td>56</td>
</tr>
</tbody>
</table>

Discussion

Our study demonstrated a significant rate of cervical lymph node metastasis in patients with supraglottic SCC. Several articles have previously reported different rates of cervical lymph node metastasis, from 12% to 40% (6–11).

Despite the classic approach to patients with N0 supraglottic SCC, which include bilateral neck dissection of levels II through IV, some authors believe that treatment of the neck in the patients with T1 N0 supraglottic SCC is not necessary (11).

Redaelli et al. showed that the risk of occult cervical metastasis in T1 patients was significantly lower than other tumor grades (11), but earlier studies showed no relation between tumor grade and cervical metastasis (12). In our study the rate of cervical nodal metastasis varied from 16.7% to 54.5% in T1 and T4 patients, respectively, but there was no statistically significant association between tumor grade and cervical lymph node metastasis. This, therefore, emphasizes the role of neck dissection for all stages.

One of the important findings in our study was that 47.6% of patients had bilateral cervical lymph node metastasis. This finding contrasts with other studies that recommended unilateral neck dissection (13).

One limitation of this study is the low sample size, which may affect the results. Further studies with more patients could provide us with more accurate conclusions.

Conclusion

This study demonstrated a significant rate of cervical lymph node metastasis in patients with supraglottic SCC. We therefore recommend elective bilateral neck dissection in all stages of N0 supraglottic SCC patients.

References

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