Surgical management of Squamous Cell Carcinoma of the lower lip: An experience of 109 cases

Wenhao Rena 1, Yin Lia 1, Changyang Liu a , Cui Qiang a , Linmei Zhang a , Ling Gao a , Zhi Wang b , Keqian Zhia 1

1 MD, DDS, MD, MD Department of Oral and Maxillofacial Surgery, Stomatological Hospital of Xi’an Jiaotong University College of Medicine, Xi’an, Shaanxi, People’s Republic of China.710004
2 MD, Department of Anesthesiology, Stomatological Hospital of Xi’an Jiaotong University College of Medicine, Xi’an, Shaanxi, People’s Republic of China.710004

Correspondence:
Department of Oral and Maxillofacial Surgery
Stomatological Hospital of Xi’an Jiaotong University
College of Medicine
Number 98, West Fifth Road
Xi’an, Shaanxi
People’s Republic of China. 710004
zhikeqian@sina.com

Received: 06/10/2013
Accepted: 25/12/2013

Abstract
Objectives: We are presenting our experience collected from a series of 109 cases with SCC of the lower lip focusing on clinical features of patients and surgical approach.
Study Design: We retrospectively analyzed medical records of patients diagnosed with Squamous Cell Carcinoma (SCC) of the lower lip at the Oral and Maxillofacial surgery at Xi’an Jiaotong University during a period between 1999 and 2008.
Results: A total of 109 patients with lip cancer were included in the study. When no frozen-section test was performed, the neoplasia was removed with a margin of at least 6 mm. Different surgical techniques were used for lip reconstruction after tumor excision. Neck dissection was performed in all patients with clinically palpable lymph nodes. Median follow-up was 38 months. During follow-up, recurrence occurred in 5 patients, 3 patients developed neck metastases, distant metastases developed in 1 patient. Five patients died during observation period.
Conclusions: The patient-related and defect-related issues must be taken into consideration during reconstruction for surgical defect. For N0 patients, we recommend wait-and-see policy. Early detection, careful follow-up and prompt neck is essential for the successful treatment.

Key words: Lip cancer, surgical management, reconstruction.
Introduction
Lip cancer is a common malignancy of the oral cavity and accounts for 23.6% to 30% of malignant tumors of the oral cavity (1-5). The most frequent malignancy related to the lips is squamous cell carcinoma (SCC), while adenocarcinomas and melanoma occur rarely (3,4). The lower lip was the most frequently affected site, which accounts for more than 90% of the cases (2,6). SCC of the lip is thought to be related to sun exposure. It also explains the greater frequency of the pathologic findings on the lower than on the upper lip, because the upper lip is far less exposed to the sun than the lower (7). However, the etiology of lip cancer is multifactorial, including exposure to sunlight, tobacco, genetic predisposition, immunosuppression and immunodeficiency (7,8). The definitive pathogenic pathway remains unclear.

Surgery is the treatment of choice for most of the tumors of the lip. Surgical resection requires a full-thickness resection of the skin, muscle and underlying mucosa to allow a safe surgical margin. Numerous reconstruction methods after tumor removal have been reported, however, the reconstruction of lip defect remains a challenge (9). Neck dissection should be performed patients with N+ disease, although there is some debate about the measures that should be adopted in N0 cases (1,8,10,11). In this work we are presenting our experience collected from a series of 109 cases with SCC of the lower lip from January 1999 to April 2008 focusing on clinical features of patients and surgical approach.

Material and Methods
A retrospective chart review was performed to analyze patients with a diagnosis of SCC of the lower lip treated in the Department of oral and maxillofacial surgery, Xi’an JiaoTong University stomatological hospital, from January 1999 through April 2008. One hundred and nine patients with preoperative diagnosis of SCC of the lower lip were identified in the medical records within this period. The charts were reviewed, and the following information was recorded: Age and sex, smoking habits and sun exposure, clinical stage, macroscopic features of the primary lesions, type of surgical treatment, follow-up, and outcome. Analysis of the data was performed using the statistical package software SPSS 13.0 (SPSS Inc., Chicago, IL, USA).

Results
A total of 109 patients with lip cancer were included in the study. There were 87 men (79.8%) and 22 women (20.2%). The overall male-to-female ratio was 4 to 1. Mean patient age was 62.5 years (range, 32 to 83 years). Distribution of patients according to age is presented in

| Age | No. | %  |
|-----|-----|----|
| ≤50 | 14  | 12.8 |
| 51-60 | 27  | 24.8 |
| 61-70 | 45  | 41.3 |
| 71-80 | 20  | 18.3 |
| >80  | 3   | 2.8  |

| Gender | No. | %  |
|--------|-----|----|
| male    | 87  | 79.8 |
| female  | 22  | 20.2 |

| TNM stage | No. | % |
|-----------|-----|----|
| T stage   |     |    |
| 1         | 39  | 35.8 |
| 2         | 52  | 47.7 |
| 3         | 16  | 14.7 |
| 4         | 2   | 1.8  |
| N stage   |     |    |
| 0         | 87  | 79.8 |
| 1         | 19  | 17.4 |
| 2         | 3   | 2.8  |
| 3         | 1   | 0.9  |
| M stage   |     |    |
| 0         | 109 | 100 |
| 1         | 0   | 0    |

| Site     | No. | %  |
|----------|-----|----|
| right side | 34  | 31.2 |
| left side  | 27  | 24.8 |
| both sides | 48  | 44  |

Table 1. Patient and tumor characteristics.

52 (47.7%) in T2, and 16 patients (14.7%) were in T3 stage. In 2 patients (1.8%) the lesion was in stage T4; 14 patients (12.8%) presented with nodal disease at diagnosis. While in 95 patients (87.2%) there were no signs of clinical regional lymphadenopathy. All patients were in M0. We observed a close relationship between the tumor size at presentation and the neck stage. Two patients with T1-2 disease (2/91) developed lymph node metastases during the follow-up period. In contrast, 6 of 18 patients with T3-4 disease developed lymph node metastases during the study period.

Neck dissection was performed in all patients with clinically palpable lymph nodes (14/109, 12.8%): 3 patients (21.4%) underwent radical neck dissection(RND), 11 patients (78.6%) underwent radical neck dissection with conservation of the external jugular vein, deep branches of the cervical plexus, and sternocleido-
Surgical management of lower lip cancer

mastoid muscle (12). For the 8 patients who were N0 at diagnosis and who developed lymph node metastasis during the follow-up period, RND were performed in 5, bilateral RND were performed in 2, and radical neck dissection (RND) and contralateral supraomohyoid neck dissection were performed in 1. All the N+ patients were given postoperative radiotherapy. Different surgical techniques were used for lip reconstruction after tumor excision. For 36 patients (33%) with defects less than one third the length of the lower lip, we used a U or V, W-shape excision. For 56 patients (51.4%) with defects between one third and two thirds lip length of the lower lip, we performed Abbe flap, Bernard flap, Karapandzic flap. For 17 patients (15.6%) with defects more than two thirds lip length of the lower lip, Bernard technique and cheek advancement flaps was used for reconstruction (Figs. 1,2).

Median follow-up was 38 months (range, 12 to 76 months). During follow-up, local recurrence occurred in 5 patients, 3 patients developed neck metastases, 6 of whom were salvaged with surgery and/or radiotherapy. Distant metastases developed in 1 patient. Five patients died during the observation period, 3 of causes unrelated to the disease treated, 2 of the patients died of lip cancer.

Discussion

Lip cancer is common in malignant neoplasm of the oral cavity. It also has been reported the most common malignant neoplasm of the oral cavity or the second frequency skin cancer in the head and neck region in some studies (8,13). The lower lip was the most frequently affected site. The predilection for this subsite has been attributed to sun exposure, because the lower lip receives considerably more direct sunlight than the upper lip. Sunlight and smoking are substantiated as risk factors for lip cancer development by epidemiological data (14,15). The sex distribution revealed only a slight tendency toward men, which is in contrast to previous studies that reported a male-to-female ratio of 5–8:1 (1,8,10,16).

Several methods for treatment of SCC of the lip are available, including surgery, radiotherapy, chemotherapy, and combinations of these. We preferred surgical removal of the tumor followed by the reconstruction of lip defects for most patients with SCC of the lip. All the patients underwent the surgical treatment in this series. The main goal of the surgery is to completely remove the tumor and the surgical treatment involves full-thickness of the skin, muscle and underlying mucosa to allow a safe surgical margin. At present, there is no consensus on the definition of an adequate margin of resection. It has been reported that a 3 mm margin with excision of early SCC of lower lip (stage I/II) seems to be appropriate, when a frozen-section test of the margins is performed (17). If it is not performed, a 6 mm margin is necessary (17). Brodland et al. (18) recommended at least a 6 mm margin for high-risk tumors of primary cutaneous squamous cell carcinoma. Some surgeons consider a minimum of 10 mm additional normal tissue at all margins of the resection (16). In this series, when no frozen-section test was performed, we kept a margin of 6-10 mm.

The lips play a key role in facial expression, speech, and eating. The goal of lip reconstruction for the surgical defect should be to maintain oral competence and maximize cosmesis. There are many techniques have been reported for lip reconstruction. However, as a result of the unique anatomy of the lips, reconstruction of the lip can be challenging, especially for the large defect. Based our experience, a wedge-shaped or V-shaped excision for smaller tumors can be perform for less than one third the length in lower lip defects. If necessary to allow for adequate resection margins, we prefer the U-shaped excisions. They are simple to perform and provide good results.

It may be a challenge for the surgeon on making decision for the defects that are between one third and two thirds the length in low lip. The abbe flap (9), Karapandzic flap (19), Staircase technique (20,21), Gillies fan flap (9,22) and Bernard technique and its modifications (23,24) has been reported to be used for reconstructions involving up to two thirds of the lower lip. According to the size and location of the defect, local tissue characteristics, individual needs of the patient and patient’s condition, we performed local sliding flap, Abbe flap, Bernard technique, Karapandzic flap for 56 patients with defects between one third and two thirds lip length of the lower lip. In our experience, the local sliding flap is easy to
performed and suitable to the small tumors; the Abbe flap is used for defects medial to the commissure and could results in a proportional reduction in the size of both the upper and the lower lip, however it need for a two-stage procedure; The Karapandzic flap preserves the nerve and bloody supply but may result in microstomia and distortion of the commissure when used for large defect; Bernard Technique and its several modifications could be used for large defect and the functional and aesthetic result outcome was acceptable. Although not commonly used, the staircase technique has been reported extremely practical and offers good aesthetic and functional results for the up to two thirds defect of the lower lip in several study (1,10,20,21). For the more than two thirds of the lower lip defect, we used Bernard technique and cheek advancement flaps. Generally, immediate reconstruction after resecting more than two thirds of the lower lip gives poor results. We consider most of these patients need secondary surgery and the result is acceptable. Generally, we don’t recommend the free flap for reconstruction of lower lip, because of the poor appearance and function. There is no standard or most acceptable surgical technique in lip reconstruction and there was little detailed assessment on postoperative functional and aesthetic outcome for patients following a lip reconstruction in current literature. However, the patient-related and defect-related issues must be taken into consideration during treatment planning. The reported rate of lymph node metastases at diagnosis is 3% to 29% (1,8,10). In this study, 12.8% of all patients had lymph node involvement at diagnosis. There were 8 N0 cases that became positive during the follow-up period, the percentage with lymph node metastases increased to 20.2%. Many studies have indicated a remarkable worsening of the survival rate for cases that become N+ during follow-up (25). Therefore, how to treat patients with clinical stage N0 disease is in controversy. Some surgeons have adopted an aggressive approach, they suggested that a prophylactic neck dissection should be performed in all N0 cases (11,26,27). Some other authors recommend the “wait-and-see” approach, proceeding to therapeutic RND and radiotherapy, if necessary (8). We consider it is important to identify the patients who will definitely benefit from an aggressive approach to the lymph nodes. The high risk of lymph node metastases such as tumor size, high histologic grade, locally recurrent, perineural invasion in N0 cases has been reported (4,10,28-30). However, it was found that the tumor size does not correlate closely with regional lymph node metastases in other study (8). In this study, We also observed a close relationship between the tumor size and the neck stage at presentation. Two patients with T1-2 disease (2/91) developed neck metastases during the follow-up period. In contrast, 6 of 18 patients with T3-4 disease developed neck metastases during the study period. JPM Vanderlei et al observed high incidence of neck metastases (more than 20%) in tumors measuring 3 to 4 cm. They suggest that prophylactic neck dissection should be indicated in lip SCCs larger than 3 cm (31)(Vanderlei, 2013 #528).We recommend the ultrasound-guided wait-and-see policy for T1-2 patients with a clinical N0 neck. Certainly compliance of the patient is required. Early detection and treatment could lead to decreased morbidity and mortality. Therefore, the role of dentists is very important in the early diagnosis of SCC of the lip. Because most of the tumors were diagnosed early (83.5%), SCC of the lip generally has a good outcome. Careful follow-up and prompt neck treatment remain important steps in the surgical management of patients with SCC of the lower lip.

Conclusions

Lower lip cancer is a frequent disease of the oral cavity. General agreement has not been reached concerning in lower lip reconstruction and unresolved questions remain with regard to N0 treatment. We report a series of 109 patients and our experiences on surgical management of SCC of the lower lip. The patient-related and defect-related issues must be taken into consideration during lip reconstruction. For N0 patients, we recommend wait-and-see policy. Careful follow-up and prompt neck treatment remain important steps. Early detection is essential for the successful treatment of SCC of the lip.

References

1. Salgarelli AC, Sartorelli F, Cangiano A, Collini M. Treatment of lower lip cancer: An experience of 48 cases. International Journal of Oral and Maxillofacial Surgery. 2005;34:27-32.
2. Chen J, Katz RV, Krutchkoff DJ, Eisenberg E. Lip cancer. Incidence trends in Connecticut, 1935-1985. Cancer. 1992;70:2025-30.
3. Hoffman HT, Karmell LH, Funk GF, Robinson RA, Mencik HR. The National Cancer Data Base report on cancer of the head and neck. Arch Otolaryngol Head Neck Surg. 1998;124:951-62.
4. Zitsch RP, Park CW, Renner GJ, Rea JL. Outcome analysis for lip carcinoma. Otolaryngology-Head and Neck Surgery. 1995;113:589-96.
5. Hasson O. Squamous cell carcinoma of the lower lip. Journal of Oral and Maxillofacial Surgery. 2008;66:1259-62.
6. de Visscher JG, Schapweld M, Otter R, Visser O, van der Waal I. Epidemiology of cancer of the lip in The Netherlands. Oral Oncol. 1998;34:421-6.
7. de Visscher JG, van der Waal I. Etiology of cancer of the lip. A review. Int J Oral Maxillofac Surg. 1998;27:199-203.
8. Vukanovic M, Jezdic Z, Petrovic M, Medenica LM, Lens M. Surgical management of squamous cell carcinoma of the lip. Analysis of a 10-year experience in 223 patients. Journal of Oral and Maxillofacial Surgery. 2007;65:675-9.
9. Coppel GL, Lin DT, Burkey BB. Current concepts in lip reconstruction. Curr Opin Otolaryngol Head Neck Surg. 2004;12:281-7.
10. Salgarelli AC, Sartorelli F, Cangiano A, Pagani R, Collini M. Surgical treatment of lip cancer: Our experience with 106 cases. Journal of Oral and Maxillofacial Surgery. 2009;67:840-5.
11. Yilmaz S, Ercocen AR. Is elective neck dissection in t1-2, n0
patients with lower lip cancer necessary? Annals of Plastic Surgery. 2009;62:381-3.
12. Longjiang L, Yuming W, Changmei W, Lijuan W. Rational radical neck dissection for oral cancer. Chin Med J (Engl). 2003;116:1123-6.
13. Casal D, Carmo L, Melancia T, Zagalo C, Cid O, Rosa-Santos J. Lip cancer: A 5-year review in a tertiary referral centre. Journal of Plastic Reconstructive and Aesthetic Surgery. 2010;63:2040-5.
14. Lindqvist C, Teppo L. Epidemiological evaluation of sunlight as a risk factor of lip cancer. Br J Cancer. 1978;37:983-9.
15. Moore SR, Johnson NW, Pierce AM, Wilson DF. The epidemiology of lip cancer: A review of global incidence and aetiology. Oral Diseases. 1999;5:185-95.
16. Lydiatt W. Management of lower lip cancer: A retrospective analysis of 118 patients and review of the literature. Archives of facial plastic surgery. 2003;5:533.
17. de Visscher J, Gooris RJJ, Vermey A, Roodenburg JLN. Surgical margins for resection of squamous cell carcinoma of the lower lip. International Journal of Oral and Maxillofacial Surgery. 2002;31:154-7.
18. Brodland DG, Zitelli JA. Surgical margins for excision of primary cutaneous squamous cell carcinoma. J Am Acad Dermatol. 1992;27:241-8.
19. Karapandzic M. Reconstruction of lip defects by local arterial flaps. Br J Plast Surg. 1974;27:93-7.
20. Salgarelli AC, Persia M, Ciancio P, Pagani R. The staircase technique for treatment of cancer of the lower lip: A report of 36 cases. Journal of Oral and Maxillofacial Surgery. 2001;59:399-402.
21. Roldan JC, Teschke M, Fritzer E, Dunsche A, Haerle F, Wiltfang J, et al. Reconstruction of the lower lip: Rationale to preserve the aesthetic units of the face. Plastic and Reconstructive Surgery. 2007;120:1231-9.
22. Gillies HD, Millard DR. The Principles and Art of Plastic Surgery. 1st Ed. Boston: Little, Brown & Co.; 1957. Back to cited text no. p.4.
23. Herrera E, Bosch RJ, Barrera MV. Reconstruction of the lower lip: Bernard technique and its variants. Dermatologic Surgery. 2008;34:648-55.
24. Freeman BS. Myoplastic modification of the bernard cheiloplasty. Plastic and reconstructive surgery and the transplantation bulletin. 1958;21:453-60.
25. Khalil HH, Elaffandi AH, Afifi A, Alsayed Y, Mahboub T, El-Refaie KM. Sentinel lymph node biopsy (slnb) in management of n0 stage t-1-t-2 tip cancer as a “same day” procedure. Oral Oncology. 2008;44:608-12.
26. Bucur A, Stefanescu L. Management of patients with squamous cell carcinoma of the lower lip and N0-neck. J Craniomaxillofac Surg. 2004;32:16-8.
27. Akpinar H, Bulut H, Berberoglu U. Is suprahyoid dissection a diagnostic operation in lower lip carcinoma? Journal of Experimental & Clinical Cancer Research. 2002;21:29-30.
28. Zitsch RP, Lee BW, Smith RB. Cervical lymph node metastases and squamous cell carcinoma of the lip. Head Neck. 1999;21:447-53.
29. Vartanian JG, Carvalho AL, de Arajo MJ, Hattori M, Magrin J, Kowalski LP. Predictive factors and distribution of lymph node metastasis in lip cancer patients and their implications on the treatment of the neck. Oral Oncology. 2004;40:223-7.
30. Luna-Ortiz K, Guemes-Meza A, Villavicencio-Valencia V, Mosqueda-Taylor A. Lip cancer experience in mexico. An 11-year retrospective study. Oral Oncology. 2004;40:992-9.
31. Vanderlei JPD, Pereira-Filho FJJ, da Cruz FA, de Mello FLV, Kruschewsky LD, de Freitas LCC,et al. Management of neck metastases in t2n0 lip squamous cell carcinoma. American Journal of Otolaryngology. 2013;34:103-6.

**Funding**
This study was supported by National Natural Science Foundation of China (81272957), Specialized Research Fund for the Doctoral Program of Higher Education (20120201110063, 20130201120054), The Fundamental Research Funds for the Central Universities(XJJ2013057).

**Conflict of Interest Statement**
The authors declare that they have no conflict of interest.