Mediastinal 4L lymph node dissection in left lung cancer

Katsuhiro Okuda, Hiroshi Haneda, Keisuke Yokota, Tsutomu Tatematsu, Ryoichi Nakanishi

Department of Oncology, Immunology and Surgery, Nagoya City University Graduate School of Medical Sciences, Nagoya, Japan *Correspondence to:* Katsuhiro Okuda, MD, PhD. Department of Oncology, Immunology and Surgery, Nagoya City University Graduate School of Medical Science, 1 Kawasumi, Mizuho-cho, Mizuho-ku, Nagoya 467-8601, Japan. Email: kokuda@med.nagoya-cu.ac.jp. *Comment on:* Wang YN, Yao S, Wang CL, *et al.* Clinical Significance of 4L Lymph Node Dissection in Left Lung Cancer. J Clin Oncol 2018;36:2935-42.

Received: 25 February 2019; Accepted: 04 June 2019; published: 17 June 2019. doi: 10.21037/shc.2019.06.01 View this article at: http://dx.doi.org/10.21037/shc.2019.06.01

Lymph node dissection is an important surgical treatment for the accurate staging and selection of patients who require postoperative adjuvant therapy, including chemotherapy and radiotherapy, for non-small cell lung cancer (NSCLC). Furthermore, the complete removal of unsuspected or microscopic cancer by lymph node dissection can reduce the risk of local recurrence and metastasis and increase the possibility of a cure. Lymph node dissection can be roughly divided into three procedures: systematic lymph node dissection, wherein the lymph nodes are removed along with the surrounding adipose tissue as a cluster; selective lymph node dissection, which involves dissection to omit the range of dissection depending on the tumor location; and sampling, where only an arbitrary lymph node is extracted. No significant difference in the treatment outcome was noted between systematic lymph node dissection and sampling in the Z0030 trial, a randomized controlled trial of the American College of Surgery Oncology Group (1). In addition, the operation time of systematic lymph node dissection was longer than that of sampling lymph node dissection, but no marked difference in the postoperative complication rate or surgical related mortality rate was noted (2). Although many studies and reviews have explored the usefulness of mediastinal lymph node dissection, the findings are controversial, with some stating that it is effective (3,4) while others found that it does not contribute to the prognosis (5-7).

Mediastinal lymph node resection is necessary for the accurate staging and resection of possible metastatic lesions. However, the omission of lymph node dissection or sampling dissection used to be performed for the patients who are not suspected of mediastinal lymph node metastasis by preoperative imaging. Dissecting mediastinal lymph nodes carries a risk of a prolonged operation time and postoperative complications. However, no data has unequivocally shown that lymph node dissection has a better prognosis than sampling. Therefore, the utility of systematic mediastinal lymph node dissection has not yet been established. Furthermore, the indications for patients requiring systematic lymph node dissection differ among institutions and surgeons, and the precision differs depending on the surgical technique used, such as thoracotomy versus video-assisted thoracoscopic surgery (VATS). This makes it difficult to clarify the utility of mediastinal systematic lymph node dissection.

There are several reports concerning the validity of systematic lymph node dissection (1-7), but relatively few have described 4L lymph node dissection. Wang et al. (8) investigated the validity of 4L lymph node dissection in left lung cancer and reported the possibility of the prognosis being improved by adding 4L lymph node dissection. However, because that study was a retrospective study with no mention of how patients were selected for 4L lymph node dissection versus non-dissection, it was impossible to conclude that 4L lymph node dissection was effective in improving the prognosis. Furthermore, the study included all patients with left lung cancer, regardless of upper or lower lobe; the method of lymph node dissection should be divided at least by the location of the tumor. The results of the progression-free survival and overall survival is important and interesting, but readers want to know the recurrence form, especially the 4L lymph node recurrence or not. Because the patients underwent mediastinal 4 L lymph node dissection will be selected by each institution and surgeon based on the general condition of the patient or tumor progression, it may be difficult to clarify whether the 4L lymph node dissection improve prognosis or not. Darling et al. (9) reported that 21 patients (4%) had occult N2 disease identified by mediastinal lymph node dissection that was not found by rigorous pre-randomization

Page 2 of 3

systematic sampling, and non-4L metastasis was found in 207 cases of left-sided NSCLC. It is also necessary to pay close attention to the risk of serious complications, such as bleeding requiring blood transfusion, recurrent nerve paralysis or chylothorax, caused by 4L lymph node dissection.

In the study of mediastinal lymph node dissection, we have to evaluate the short-term survival, including perioperative complications, as well as the long-term survival. Given recent improvements in the accuracy of preoperative clinical staging, especially for lymph node metastasis, thanks to advances in preoperative imaging, such as positron emission tomography, the role of mediastinal lymph node dissection for staging is expected to change. The technique and range of mediastinal lymph node dissection differs among surgeons and surgical operation type, including thoracotomy, VATS or robotic-assisted thoracoscopic surgery (10). In order to clarify the usefulness of mediastinal lymph node dissection, a study must consider such detailed background factors.

Mediastinal lymph node dissection including 4L lymph nodes would be necessary for truly accurate staging and decision of postoperative adjuvant therapy, at least for the patients underwent left upper lobectomy. However, because there is a risk of serious perioperative complications, joint prospective studies of other institutions with as little bias as possible due to differences in case selection and surgical method will be necessary to prove the usefulness of 4L lymph node dissection.

Acknowledgments

Funding: None.

Footnote

Provenance and Peer Review: This article was commissioned by the editorial office, *Shanghai Chest.* The article did not undergo external peer review.

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at http://dx.doi. org/10.21037/shc.2019.06.01). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

Open Access Statement: This is an Open Access article distributed in accordance with the Creative Commons Attribution-NonCommercial-NoDerivs 4.0 International License (CC BY-NC-ND 4.0), which permits the non-commercial replication and distribution of the article with the strict proviso that no changes or edits are made and the original work is properly cited (including links to both the formal publication through the relevant DOI and the license). See: https://creativecommons.org/licenses/by-nc-nd/4.0/.

References

- Izbicki JR, Passlick B, Pantel K, et al. Effectiveness of radical systematic mediastinal lymphadenectomy in patients with resectable non-small cell lung cancer: results of a prospective randomized trial. Ann Surg 1998;227:138-44.
- Allen MS, Darling GE, Pechet TT, et al. Morbidity and mortality of major pulmonary resections in patients with early-stage lung cancer: initial results of the randomized, prospective ACOSOG Z0030 trial. Ann Thorac Surg 2006;81:1013-9; discussion 1019-20.
- Wright G, Manser RL, Byrnes G, et al. Surgery for nonsmall cell lung cancer: systematic review and meta-analysis of randomised controlled trials. Thorax 2006;61:597-603.
- Mokhles S, Macbeth F, Treasure T, et al. Systematic lymphadenectomy versus sampling of ipsilateral mediastinal lymph-nodes during lobectomy for nonsmall-cell lung cancer: a systematic review of randomized trials and a meta-analysis. Eur J Cardiothorac Surg 2017;51:1149-56.
- Meng D, Zhou Z, Wang Y, et al. Lymphadenectomy for clinical early-stage non-small-cell lung cancer: a systematic review and meta-analysis. Eur J Cardiothorac Surg 2016;50:597-604.
- Huang X, Wang J, Chen Q, et al. Mediastinal lymph node dissection versus mediastinal lymph node sampling for early stage non-small cell lung cancer: a systematic review and meta-analysis. PLoS One 2014;9:e109979.
- Darling GE, Allen MS, Decker PA, et al. Randomized trial of mediastinal lymph node sampling versus complete lymphadenectomy during pulmonary resection in the patient with N0 or N1 (less than hilar) non-small cell carcinoma: results of the American College of Surgery Oncology Group Z0030 Trial. J Thorac Cardiovasc Surg 2011;141:662-70.
- Wang YN, Yao S, Wang CL, et al. Clinical Significance of 4L Lymph Node Dissection in Left Lung Cancer. J Clin Oncol 2018;36:2935-42.

Shanghai Chest, 2019

 Darling GE, Allen MS, Decker PA, et al. Number of lymph nodes harvested from a mediastinal lymphadenectomy: results of the randomized, prospective American College of Surgeons Oncology Group Z0030 trial. Chest 2011;139:1124-9.

doi: 10.21037/shc.2019.06.01

Cite this article as: Okuda K, Haneda H, Yokota K, Tatematsu T, Nakanishi R. Mediastinal 4L lymph node dissection in left lung cancer. Shanghai Chest 2019;3:31.

 Shimada Y, Saji H, Kakihana M, et al. Retrospective analysis of nodal spread patterns according to tumor location in pathological N2 non-small cell lung cancer. World J Surg 2012;36:2865-71.