

Locoregional recurrence after VATS surgery for NSCLC

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Comment on: Haruki T, Miwa K, Araki K, *et al.* Distribution and Prevalence of Locoregional Recurrence after Video-Assisted Thoracoscopic Surgery for Primary Lung Cancer. *Thorac Cardiovasc Surg* 2016;64:526-32.

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In the present article, Doctor Haruki and associates (1) seek to address the very invigorating question of the distribution and prevalence of locoregional recurrence after video-assisted thoracoscopic surgery (VATS). This remarkable work not only contributes to a deeper understanding of the patterns of recurrence, but also questions some of the traditional and prominent surgical dogmas in the area of VATS lung resection for primary lung cancer.

For this purpose, the authors retrospectively reviewed 248 patients with primary lung cancer submitted to scheduled VATS lobectomy or segmentectomy (with mediastinal lymph node dissection) in their department, over a 7 years period (January 2005–December 2011). The results showed that there were 47 cases of postoperative recurrences among the 248 included patients. They were classified as follows: 26 distant, 6 locoregional and distant, and 15 locoregional recurrences. The rate of locoregional recurrences was 6.0%. Of these 15 cases, 2 concerned the bronchial stump and lung parenchyma transection line (0.4%), five the ipsilateral pleura (2.0%), and 8 recurred within ipsilateral hilar and mediastinal lymph nodes (3.2%). Univariate analysis showed 3 variables to be significant associated factors for locoregional recurrence: pleural invasion ($P=0.02$), lymphovascular invasion ($P=0.04$), and pathological stage I *vs.* \geq II ($P<0.01$). Multivariate analysis revealed that advanced stages remained the only significant associated factors for locoregional recurrence ($P<0.01$, OR: 3.3). Finally, the authors concluded that although their observed locoregional recurrence rate appears as acceptable—and within the range of data reported in the

available international medical literature—they should explore more effective treatment modalities against histologically proven locally advanced lung cancer to prevent not only distant metastases, but also locoregional recurrences.

Locoregional recurrences are frequently used as endpoint (primary or more often secondary) for outcomes analysis and are currently considered as indirect indicators for quality of care in breast cancer surgery: not only quality of surgical treatment given to the patient, but also subsequent quality of life following surgery (2,3). However, in the specific framework of thoracic surgery, a limited number of studies have addressed the interesting, but seldom debated question of patterns of recurrence during the past decade. Most of the latter have been single institutions' series (4-8). This innovative article is most timely and helps to clarify some of the controversies about oncologic efficacy and technical quality of operation. Its main findings might have a relevant impact on patients' management in the near future.

Concerning factors of recurrence, we currently know that the issue is not whether to perform a VATS or a thoracotomy—we have enough evidence and this debate seems outdated! (9-11)—but how to decrease postoperative recurrence rate and which armamentarium we could use for this purpose. The authors nicely indicate paths for reflection and action. To summarize, the cornerstone is strict compliance to some well-established fundamental surgical rules (12). For example, the operative specimen should be manipulated with caution in order to lower the risk of dissemination (avoid use of lung grasping forceps;

retrieve specimen with a protection bag); perform routine frozen section analysis of the bronchial resection margin; dissect lymph nodes very carefully, without severing the capsula and strive to an en-bloc removal. No compromise towards basic oncological principles should be tolerated. In each situation where they might be questionable, a conversion to open thoracotomy should be undertaken.

The current contribution shows an essential way to explore, the prognostic impact of the extent of lymph node dissection in early stage lung cancer. Although Ichinose *et al.* and Maniwa *et al.* in their retrospective studies (9,13), recently followed by Adachi in a propensity score matching analysis (14), have shown that lobe-specific lymphadenectomy for clinical stage I lung cancer was safe and resulted in acceptable locoregional control, the extent of lymph node dissection is still a matter of controversy. The manuscript of Mordant *et al.* might be helpful in this context (15). It sheds light on the key prognostic role of the location of pN1 lymph nodes in case of multi-station disease, and of the number of metastatic stations. Survival of multi-station N1 disease was comparable to the survival of skip N2 (single station N0–N2) disease, and significantly worse than single-station N1 disease. We should also remind that the N1 category *per se* is mixing up 2 subcategories with sharply differing prognosis: survival of intra-lobar N1 mirrors the lower range of survival for stage I, while prognosis of extralobar N1 is in the higher range of stage IIIA (16,17). Future works focusing on the extent of lymph node dissection should also address definition of guidelines in order to reduce the part of locoregional recurrences which are in the reach of optimized surgical care.

Another critical aspect is multidisciplinary interaction. We know that radiation therapy may improve local control, but is subjected to some toxicity and increased complications in case of redo surgery. The effect of adjuvant chemotherapy is unfortunately marginal only, and unlikely to compensate for incomplete resection. Targeted therapies and anti-angiogenic medications are not recommended, and there is no information available about immunotherapy (18,19). The relatively low contribution of adjuvant therapies to long-term survival places a strong accent onto quality requirements of surgical care.

For the time being, Doctor Haruki and associates are to be congratulated on their investigations in this area. Their results will certainly prove to be most beneficial to the thoracic surgery community from the standpoint of medical care and affect future patient management.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

References

1. Haruki T, Miwa K, Araki K, et al. Distribution and Prevalence of Locoregional Recurrence after Video-Assisted Thoracoscopic Surgery for Primary Lung Cancer. *Thorac Cardiovasc Surg* 2016;64:526-32.
2. Ernst MF, Voogd AC, Coebergh JW, et al. Using locoregional recurrence as an indicator of the quality of breast cancer treatment. *Eur J Cancer* 2004;40:487-93.
3. Morgan DA, Kurtz JM. Is local control necessarily an indicator of quality? *Eur J Cancer* 2004;40:472-3.
4. Koike T, Koike T, Yoshiya K, et al. Risk factor analysis of locoregional recurrence after sublobar resection in patients with clinical stage IA non-small cell lung cancer. *J Thorac Cardiovasc Surg* 2013;146:372-8.
5. Sawabata N. Locoregional recurrence after pulmonary sublobar resection of non-small cell lung cancer: can it be reduced by considering cancer cells at the surgical margin? *Gen Thorac Cardiovasc Surg* 2013;61:9-16.
6. El-Sherif A, Fernando HC, Santos R, et al. Margin and local recurrence after sublobar resection of non-small cell lung cancer. *Ann Surg Oncol* 2007;14:2400-5.
7. Fan C, Gao S, Hui Z, et al. Risk factors for locoregional recurrence in patients with resected N1 non-small cell lung cancer: a retrospective study to identify patterns of failure and implications for adjuvant radiotherapy. *Radiat Oncol* 2013;8:286.
8. Zhang Z, Zhang Y, Feng H, et al. Is video-assisted thoracic surgery lobectomy better than thoracotomy for early-stage non-small-cell lung cancer? A systematic review and meta-analysis. *Eur J Cardiothorac Surg* 2013;44:407-14.
9. Ichinose J, Kohno T, Fujimori S, et al. Locoregional control of thoracoscopic lobectomy with selective lymphadenectomy for lung cancer. *Ann Thorac Surg* 2010;90:235-9.
10. Flores RM, Ihekweazu UN, Rizk N, et al. Patterns of recurrence and incidence of second primary tumors after lobectomy by means of video-assisted thoracoscopic

- surgery (VATS) versus thoracotomy for lung cancer. *J Thorac Cardiovasc Surg* 2011;141:59-64.
11. Lee PC, Nasar A, Port JL, et al. Long-term survival after lobectomy for non-small cell lung cancer by video-assisted thoracic surgery versus thoracotomy. *Ann Thorac Surg* 2013;96:951-60; discussion 960-1.
 12. Thomas P, Dahan M, Riquet M, et al. Practical issues in the surgical treatment of non-small cell lung cancer. Recommendations from the French Society of Thoracic and Cardiovascular Surgery. *Rev Mal Respir* 2008;25:1031-6.
 13. Maniwa T, Okumura T, Isaka M, et al. Recurrence of mediastinal node cancer after lobe-specific systematic nodal dissection for non-small-cell lung cancer. *Eur J Cardiothorac Surg* 2013;44:e59-64.
 14. Adachi H, Sakamaki K, Nishii T, et al. Lobe-Specific Lymph Node Dissection as a Standard Procedure in Surgery for Non-Small Cell Lung Cancer: A Propensity Score Matching Study. *J Thorac Oncol* 2017;12:85-93.
 15. Mordant P, Pricopi C, Legras A, et al. Prognostic factors after surgical resection of N1 non-small cell lung cancer. *Eur J Surg Oncol* 2015;41:696-701.
 16. van Velzen E, de la Rivière AB, Elbers HJ, et al. Type of lymph node involvement and survival in pathologic N1 stage III non-small cell lung carcinoma. *Ann Thorac Surg* 1999;67:903-7.
 17. Riquet M, Pricopi C, Arame A, et al. From anatomy to lung cancer: questioning lobe-specific mediastinal lymphadenectomy reliability. *J Thorac Dis* 2016;8:2387-2390.
 18. Kelly K, Altorki NK, Eberhardt WE, et al. Adjuvant Erlotinib Versus Placebo in Patients With Stage IB-III A Non-Small-Cell Lung Cancer (RADIANT): A Randomized, Double-Blind, Phase III Trial. *J Clin Oncol* 2015;33:4007-14.
 19. Goss GD, O'Callaghan C, Lorimer I, et al. Gefitinib versus placebo in completely resected non-small-cell lung cancer: results of the NCIC CTG BR19 study. *J Clin Oncol* 2013;31:3320-6.

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