

## Completion Thyroidectomy: Revisited a Quarter of a Century Later

Benzon M. Dy, MD<sup>1</sup>, Veljko Strajina, MD<sup>2</sup>, Michael Tuttle, MD<sup>3</sup>, and Ashok R. Shaha, MD<sup>3</sup>

<sup>1</sup>Mayo Clinic, Breast, Endocrine, Metabolic and GI Surgery, Rochester; <sup>2</sup>Mayo Clinic, Surgery, Rochester; <sup>3</sup>Department of Head and Neck Surgery, Memorial Sloan-Kettering Cancer Center, New York

The excellent outcome for patients with thyroid cancer is increasingly recognized, with Japan and the United States considerably interested in observing selected microcarcinomas.<sup>1,2</sup> In the last decade, our experience has shown that most thyroid microcarcinomas can be observed safely.<sup>3</sup> It therefore is understandable why the potential overtreatment of thyroid cancer has been debated, not only among professionals, but also in the media and general public.

The complications of thyroidectomy may be worse than the thyroid cancer itself, and the complications are directly related to the extent of surgery. Even experienced surgeons report a 1% rate of permanent hypoparathyroidism when both thyroid lobes are removed.<sup>4–7</sup> Conversely, this complication is virtually unknown after thyroid lobectomy. Some patients may require thyroid hormone supplementation after thyroid lobectomy, but it appears to be tolerated much better than full replacement, which is needed after total thyroidectomy. Certain patients, particularly young women, appear never to reach their prior quality of life despite achieving a normal level of thyroid-stimulating hormone.

The philosophy of “let the punishment fit the crime” or “let the treatment not be worse than the disease” is critical in the management of thyroid cancer. In 1992, we published a paper, “Completion Thyroidectomy: A Critical Appraisal” in *Surgery* defining the role of completion thyroidectomy for selected patients.<sup>8</sup> In this report, we reviewed the evolution of the thought process regarding

restrictive use of contralateral thyroid lobectomy as a secondary surgical procedure for differentiated thyroid cancer.

No level 1 evidence exists to provide a clear answer regarding the adequate extent of surgery for thyroid cancer. Both national database studies and single-institution retrospective studies give somewhat mixed results,<sup>9–12</sup> with more recent studies demonstrating no effect of surgery extent on survival.

Conceptually, how would the removal of the contralateral lobe have a positive effect on either patient survival or recurrence of differentiated thyroid cancer? The argument can be summarized into three assumptions: (1) total thyroidectomy enables use of serum thyroglobulin (Tg) levels in surveillance; (2) total thyroidectomy prevents development of contralateral malignancy; and (3) total thyroidectomy allows for the administration of radioactive iodine (RAI).

The correlation between the Tg level and the risk of persistent or recurrent disease has been documented,<sup>13</sup> but it is unknown whether Tg level alone would substantially change management strategy or result in improved patient outcome. Subjecting a patient to the additional morbidity of a total thyroidectomy to achieve more sensitive biochemical surveillance has questionable scientific foundations. As our understanding of malignancy risk in thyroid nodules improves, so does our ability to stratify risk noninvasively (or at least with minimal invasiveness). With that ability and routine preoperative ultrasound, the same approach to stratifying malignancy risk can be applied to both thyroid lobes, independently of contralateral findings. Ongoing surveillance can be advised if it is supported by history (no family history of thyroid cancer or neck radiation) or sonographic, and/or fine-needle aspiration biopsy

findings within the ipsilateral thyroid lobe. The concept of risk-reducing contralateral thyroid lobectomy appears to be obsolete in this era of modern imaging.

During the last 50 years in the United States, the entire philosophy of managing thyroid cancer has revolved around routine use of RAI.<sup>14</sup> More recently, it has been recognized that a subset of patients may not benefit from RAI, and some large tertiary referral centers have started to advocate for selective use of RAI.<sup>15,16</sup> The decision to administer RAI is based on the anticipated aggressive biology of the tumor rather than direct evidence of the benefit RAI offers for a particular subgroup of patients.

Our ability to predict the biologic behavior of the tumor has evolved with the development of risk stratification models at Mayo Clinic (MACIS), Lahey Clinic (AIMS), and Memorial Sloan Kettering Cancer Center (GAMES).<sup>17–20</sup> In general, markers of more aggressive tumor biology include older patient age, greater tumor size, gross extrathyroidal extension, bulky nodal metastases, and higher grade of tumor or aggressive histologic subtype. Four of these factors often can be diagnosed or strongly suspected preoperatively.

Certain features previously popularized as an indication for completion thyroidectomy have been recognized as poor predictors of aggressive tumor biology: (1) central compartment micrometastases have a questionable effect on prognosis;<sup>21</sup> (2) the presence of multifocality of micropapillary carcinoma has very little implication in the overall management of thyroid cancer;<sup>22</sup> and (3) minimally invasive follicular carcinoma with few foci of vascular invasion or capsular invasion, as noted by van Heerden,<sup>18</sup> can be considered a “nonthreatening malignancy”. Moreover, individuals with these features do not appear to exhibit a strong indication for completion thyroidectomy.

In essentially two clinical scenarios a surgeon faces a dilemma whether to perform completion thyroidectomy: (1) a patient with known malignancy but new pathologic findings that could indicate an RAI; and (2) a patient with a diagnosis of malignancy after a diagnostic lobectomy. Considering the abilities for preoperative risk stratification offered by current imaging technology and the tendency of a more aggressive cancer to present such a situation early on,<sup>23</sup> the first scenario should be infrequent. Intraoperative, frozen-section pathology can be particularly helpful in the second scenario, not only by establishing the diagnosis of malignancy, but also by providing the information needed to decide between lobectomy and total thyroidectomy (e.g., presence of extensive vascular invasion in follicular thyroid cancer or a more aggressive histologic variant of papillary thyroid carcinoma). However, the use of frozen-section pathology appears to be highly institution dependent, with conflicting reports regarding reliability and cost effectiveness.<sup>24,25</sup> In either situation, as nicely summarized

by the American Thyroid Association almost 10 years ago, this question can be answered simply: completion thyroidectomy should be offered to patients for whom a near-total or total thyroidectomy would have been recommended had the diagnosis been available before the initial surgery,<sup>26</sup> with some relatively minor caveats. Completion thyroidectomy is not simply a total thyroidectomy in stages, and a valid question can be raised if completion thyroidectomy, performed as a “redo operation,” carries a higher risk. For all practical purposes, the risks of technical complications are likely equivalent because most of the operation is performed in essentially virgin planes, particularly the critical portions of the operation.

Finally, it should be remembered that RAI rather than removal of the contralateral thyroid lobe may provide the benefit for the patient. Total thyroidectomy only paves the way for it. Therefore, the main question for a surgeon considering completion thyroidectomy is will the patient benefit from RAI? However, answering this question may not be straightforward. Although different risk stratification models exist, these systems have limitations, including the relatively low quality of data supporting their role in decision-making. Therefore, institutions that treat patients with thyroid cancer use multidisciplinary input from surgeons, endocrinologists, nuclear medicine specialists, pathologists, radiation oncologists, and medical oncologists. Although the decision regarding completion thyroidectomy may be guided by risk stratification models, the choice should be made in cooperation with the multidisciplinary team and the patient.

## REFERENCES

1. Ito Y, Miyauchi A, Inoue H, et al. An observational trial for papillary thyroid microcarcinoma in Japanese patients. *World J Surg.* 2010;34:28–35.
2. Ito Y, Miyauchi A, Kihara M, Higashiyama T, Kobayashi K, Miya A. Patient age is significantly related to the progression of papillary microcarcinoma of the thyroid under observation. *Thyroid.* 2014;24:27–34.
3. Tuttle RM, Fagin JA, Minkowitz G, et al. Natural history and tumor volume kinetics of papillary thyroid cancers during active surveillance. *JAMA Otolaryngol Head Neck Surg.* 2017;143:1015–20.
4. Ritter K, Elfenbein D, Schneider DF, Chen H, Sippel RS. Hypoparathyroidism after total thyroidectomy: incidence and resolution. *J Surg Res.* 2015;197:348–53.
5. Asari R, Passler C, Kaczirek K, Scheuba C, Niederle B. Hypoparathyroidism after total thyroidectomy: a prospective study. *Arch Surg.* 2008;143:132–7; discussion 138.
6. Su A, Wang B, Gong Y, Gong R, Li Z, Zhu J. Risk factors of hypoparathyroidism following total thyroidectomy with central lymph node dissection. *Med Baltimore.* 2017;96:e8162.
7. Kim SM, Kim HK, Kim KJ, et al. Recovery from permanent hypoparathyroidism after total thyroidectomy. *Thyroid.* 2015;25:830–33.

8. Shaha AR, Jaffe BM. Completion thyroidectomy: a critical appraisal. *Surgery*. 1992;112:1148–52; discussion 1152–43.
9. Bilimoria KY, Bentrem DJ, Ko CY, et al. Extent of surgery affects survival for papillary thyroid cancer. *Ann Surg*. 2007;246:375–81; discussion 381–74.
10. Hay ID, Grant CS, Bergstralh EJ, Thompson GB, van Heerden JA, Goellner JR. Unilateral total lobectomy: is it sufficient surgical treatment for patients with AMES low-risk papillary thyroid carcinoma? *Surgery*. 1998;124:958–64; discussion 964–56.
11. Adam MA, Pura J, Goffredo P, et al. Impact of extent of surgery on survival for papillary thyroid cancer patients younger than 45 years. *J Clin Endocrinol Metab*. 2015;100:115–21.
12. Nixon IJ, Ganly I, Patel SG, et al. Thyroid lobectomy for treatment of well-differentiated intrathyroid malignancy. *Surgery*. 2012;151:571–79.
13. Duren M, Siperstein AE, Shen W, Duh QY, Morita E, Clark OH. Value of stimulated serum thyroglobulin levels for detecting persistent or recurrent differentiated thyroid cancer in high- and low-risk patients. *Surgery*. 1999;126:13–9.
14. Mazzaferri EL. An overview of the management of papillary and follicular thyroid carcinoma. *Thyroid*. 1999;9:421–27.
15. Nixon IJ, Patel SG, Palmer FL, et al. Selective use of radioactive iodine in intermediate-risk papillary thyroid cancer. *Arch Otolaryngol Head Neck Surg*. 2012;138:1141–46.
16. Hay ID, Hutchinson ME, Gonzalez-Losada T, et al. Papillary thyroid microcarcinoma: a study of 900 cases observed in a 60-year period. *Surgery*. 2008;144:980–7; discussion 987–8.
17. Hay ID, Thompson GB, Grant CS, et al. Papillary thyroid carcinoma managed at the Mayo Clinic during six decades (1940–1999): temporal trends in initial therapy and long-term outcome in 2444 consecutively treated patients. *World J Surg*. 2002;26:879–85.
18. Hay ID, Bergstralh EJ, Goellner JR, et al. Predicting outcome in papillary thyroid carcinoma: development of a reliable prognostic scoring system in a cohort of 1779 patients surgically treated at one institution during 1940 through 1989. *Surgery*. 1993;114:1050–57; discussion 1057–58.
19. Shaha AR, Loree TR, Shah JP. Intermediate-risk group for differentiated carcinoma of thyroid. *Surgery*. 1994;116:1036–40; discussion 1040–31.
20. Cady B, Rossi R. An expanded view of risk-group definition in differentiated thyroid carcinoma. *Surgery*. 1988;104:947–53.
21. Cranshaw IM, Carnaille B. Micrometastases in thyroid cancer: an important finding? *Surg Oncol*. 2008;17:253–58.
22. Chow SM, Law SC, Chan JK, Au SK, Yau S, Lau WH. Papillary microcarcinoma of the thyroid: prognostic significance of lymph node metastasis and multifocality. *Cancer*. 2003;98:31–40.
23. Lang BH, Lo CY, Chan WF, Lam KY, Wan KY. Staging systems for papillary thyroid carcinoma: a review and comparison. *Ann Surg*. 2007;245:366–78.
24. Paphavasit A, Thompson GB, Hay ID, et al. Follicular and Hurthle cell thyroid neoplasms: is frozen-section evaluation worthwhile? *Arch Surg*. 1997;132:674–78; discussion 678–80.
25. Udelsman R, Westra WH, Donovan PI, Sohn TA, Cameron JL. Randomized prospective evaluation of frozen-section analysis for follicular neoplasms of the thyroid. *Ann Surg*. 2001;233:716–22.
26. American Thyroid Association Guidelines Taskforce on Thyroid N, Differentiated Thyroid C, Cooper DS, et al. Revised American Thyroid Association management guidelines for patients with thyroid nodules and differentiated thyroid cancer. *Thyroid*. 2009;19:1167–214.