

Long-term Management and Outcome of Parathyroidectomy for Sporadic Primary Multiple-Gland Disease

D. Michael Rose, MD; Thomas F. Wood, MD; Andre J. Van Herle, MD; Pejman Cohan, MD; Frederick R. Singer, MD; Armando E. Giuliano, MD

Hypothesis: For a specific subset of patients with sporadic primary multiple-gland parathyroid disease, subtotal parathyroidectomy results in long-term normocalcemia in the majority of patients, with a minimal complication rate.

Design: Retrospective analysis of outcomes in patients undergoing parathyroidectomy performed by a single surgeon (A.E.G.) between 1984 and 1999.

Setting: A multidisciplinary endocrine service based at a tertiary referral center.

Patients: Patients undergoing subtotal parathyroidectomy for primary hyperparathyroidism due to sporadic multiple-gland disease identified from a single surgeon's operative records (A.E.G.).

Main Outcome Measures: Data analyzed included demographic factors, operative and pathologic findings, and postoperative and long-term clinical and laboratory results, including calcium and intact parathyroid hormone levels.

Results: Of 379 patients undergoing parathyroidectomy for hyperparathyroidism between 1984 and 1999,

49 (13%) had sporadic multiple-gland disease. Median preoperative calcium and intact parathyroid hormone (iPTH) levels were 2.7 mmol/L (10.8 mg/dL) and 11.79 pmol/L, respectively. Postoperative calcium and iPTH levels were available in 39 patients, and median values were 2.28 mmol/L (9.1 mg/dL) and 2.84 pmol/L, respectively. Long-term follow-up was available for 36 patients (73%), and duration ranged from 6 to 180 months (median, 44 months). Median calcium and iPTH levels at follow-up were 2.3 mmol/L (9.2 mg/dL) and 3.26 pmol/L, respectively, with 3 (8%) of 36 patients having evidence of persistent or recurrent hyperparathyroidism. No patient had biochemical evidence of hypoparathyroidism at long-term follow-up. Five patients (14%) had persistent elevated iPTH levels (range, 8.11-10.95 pmol/L) and normal calcium levels.

Conclusions: Subtotal parathyroidectomy for sporadic primary multiple-gland disease resulted in a long-term normocalcemia rate of 92%, with minimal complications. Selective subtotal parathyroidectomy can yield excellent long-term results in patients with multiple-gland disease.

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From the John Wayne Cancer Institute at Saint John's Health Center, Santa Monica, Calif (Drs Rose, Wood, Singer, and Giuliano), and the UCLA School of Medicine, Division of Endocrinology, University of California, Los Angeles (Drs Van Herle and Cohan).

PARATHYROIDECTOMY for primary hyperparathyroidism (HPT) is generally regarded as one of the most consistently successful surgical procedures, curing approximately 95% of all patients undergoing resections performed by experienced surgeons.¹ Patients undergoing resection of a single parathyroid adenoma, however, represent approximately 85% of these cases in most series. Fewer outcome data are available following operative intervention for the subset of patients with sporadic primary multiple-gland disease. The reported incidence of multiglandular disease has ranged from 6.8% to 39% of all patients with primary HPT, and it in-

cludes patients with sporadic or familial hyperplasia, multiple adenomas, and multiple endocrine neoplasia (MEN) syndromes.²⁻⁴

The appropriate management of multiple-gland disease in primary HPT continues to be debated because there is little information on outcome following treatment. In addition, the available data are largely a combination of outcomes for patients with both sporadic and familial disease; primary and secondary, or tertiary HPT; and both 4-gland hyperplasia and multiple parathyroid adenomas. The intraoperative finding of multiple-gland disease at the time of parathyroidectomy is common in large series of patients with HPT; however, the appropriate operative

PATIENTS AND METHODS

A single surgeon's (A.E.G.) operative records were examined to identify patients undergoing operation for HPT. A formalized protocol was approved by the institutional review board of the John Wayne Cancer Institute and the Saint John's Health Center, Santa Monica, Calif, and patients participated in the study after providing informed consent. The protocol included completion of a questionnaire requesting information on number of operations for HPT, complications following operation, and current care by an endocrinologist. Patients were requested to have blood drawn for an evaluation of current calcium and intact parathyroid hormone (iPTH) levels.

Data obtained included demographic factors, presenting symptoms, preoperative clinical data, operative and pathologic factors, immediate postoperative clinical and laboratory findings, and long-term clinical and laboratory findings. Patients with secondary or tertiary HPT, evidence of MEN syndrome, or familial HPT were excluded from analysis. Operative and pathologic reports were reviewed, and patients who had both intraoperative evidence of multiple-gland enlargement and who had undergone complete excision of at least 2 histopathologically hypercellular glands were included in the study.

All patients in this study underwent formal bilateral neck exploration without preoperative localization. At operation, all 4 glands were identified and inspected for enlargement. Failure to identify 4 glands prompted exploration of the carotid sheath and retroesophageal space, transcervical thymectomy, and ipsilateral thyroid lobectomy if the fourth gland remained unidentified. In cases of 4-gland enlargement, a subtotal parathyroidectomy was performed, leaving approximately 100 mg of parathyroid tissue in an anatomically accessible position should re-exploration be required. This remnant was marked with a metallic clip and permanent suture to aid identification during subsequent reexploration. Parathyroid autotransplantation was performed only if the remaining glands were possibly compromised. In patients with asymmetric hyperplasia or multiple adenomas, biopsies were performed on grossly normal glands, which were left in place, and all grossly enlarged glands were excised.

intervention remains uncertain. To determine the results of a standardized operative approach in a homogeneous subset of patients with sporadic multiple-gland disease, we evaluated immediate postoperative and long-term clinical and biochemical outcomes of a single surgeon's experience throughout the last 15 years.

RESULTS

PATIENT DEMOGRAPHICS

Between 1984 and 1999, 379 patients underwent parathyroidectomy for HPT. Three hundred seventeen patients (84%) had a single adenoma, 49 (13%) had spo-

Table 1. Demographics and Presenting Symptoms of 49 Patients With Primary Multiple-Gland Disease

Symptom	Patients, No. (%)
Neuropsychiatric	10 (20)
Bony disease	9 (18)
Gastrointestinal	8 (16)
Musculoskeletal	6 (12)
Nephrolithiasis	3 (6)
Asymptomatic	22 (45)

Table 2. Operative Procedures in 49 Patients With Primary Multiple-Gland Disease

	Patients, No. (%)
No. of operations	
1	39 (80)
2	8 (16)
3	2 (4)
No. of glands resected	
4½	1 (2)
3½	30 (62)
3	3 (6)
2½	5 (10)
2	10 (20)

radic multiple-gland disease, 9 (2%) had secondary or tertiary HPT, and 4 (1%) had MEN syndromes. The 49 patients with multiple-gland disease had a median age of 64 years (range, 26-77 years) and a male-female ratio of 15:34. Their symptoms at initial visit are presented in **Table 1**. Of note, 55% of the patients were symptomatic, usually with multiple symptoms. Median preoperative calcium and iPTH levels were 2.7 mmol/L (10.8 mg/dL) (range, 2.28-3.35 mmol/L [9.1-13.4 mg/dL]) and 11.8 pmol/L (range, 5.37-141.31 pmol/L), respectively.

OPERATIVE PROCEDURES AND FINDINGS

The majority of the 49 patients with sporadic multiple-gland disease underwent a single operation (**Table 2**). Of the 10 patients (20%) who underwent multiple surgical procedures, 8 had recurrent HPT after undergoing a previous parathyroidectomy elsewhere (resection of a single abnormal gland in 7 patients, and 2 abnormal glands in 1 patient). These patients visited the service 2 to 180 months (median, 72 months) following their initial surgical procedure. Two other patients failed to achieve normocalcemia on immediate postoperative analysis and underwent a second exploration within 72 hours. One patient underwent conversion to a subtotal 3½-gland parathyroidectomy, and the second patient was found to have a fifth hypercellular gland and underwent excision of 4½ parathyroids.

Four glands were identified in 43 (88%) of the 49 cases. For purposes of analysis, a properly identified gland was any gland removed by an outside surgeon during prior exploration, or any gland identified by our group during subsequent operation. The total number of resected glands is presented in Table 2; the majority of patients

underwent a 3½-gland resection for 4-gland hyperplasia. Patients undergoing resection of fewer than 3½ glands had grossly normal parathyroid glands identified intraoperatively; biopsies were performed on these normal-appearing glands for frozen-section examination. All resected glands had histopathologic evidence of increased cellularity and decreased fat content consistent with hyperplasia. The median weight of the removed glands was 200 mg (range, 21-10400 mg). Two patients underwent autotransplantation of parathyroid tissue because of concern about the viability of the remaining gland.

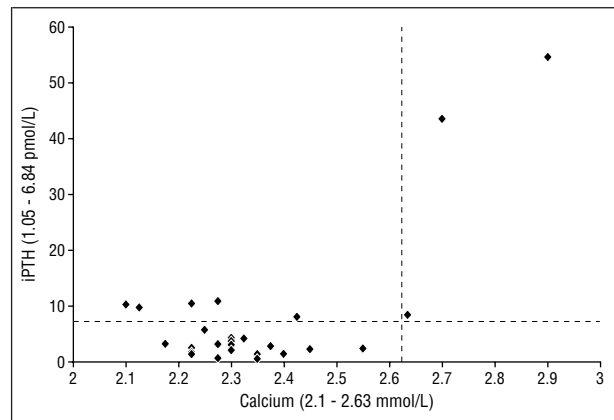
POSTOPERATIVE AND LONG-TERM RESULTS

Postoperative calcium and iPTH levels were available in 39 patients, and median values were 2.28 mmol/L (9.1 mg/dL) (range, 1.95-2.75 mmol/L [7.8-11.0 mg/dL]) and 2.84 pmol/L (range, 0.21-15.58 pmol/L), respectively. Only 1 patient had elevated calcium and iPTH levels in the immediate postoperative period; however, an additional 5 patients had elevated iPTH levels and normal calcium levels, suggestive of secondary HPT. Long-term data were available for 36 patients (73%), who were followed up for a range of 6 to 180 months (median, 44 months). The 13 patients unavailable for follow-up had either died or moved with no forwarding information. A single patient suffered a temporary recurrent laryngeal nerve injury following parathyroidectomy. Of note, this patient concurrently underwent a total thyroidectomy for incidental malignant material found at operation; however, she subsequently had normal vocal cord mobility, which was documented endoscopically, and she is currently asymptomatic with a normal voice. Median calcium and iPTH levels at follow-up were 2.3 mmol/L (9.2 mg/dL) (range, 2.1-2.9 mmol/L [8.4-11.6 mg/dL]) and 3.26 pmol/L (range, 0.63-54.86 pmol/L), respectively (**Figure**). No patient had biochemical evidence of permanent hypoparathyroidism. Three (8%) of the 36 patients had evidence of recurrent or persistent HPT, and an additional 5 patients (14%) had elevated iPTH levels (range, 8.10-10.95 pmol/L), with normal calcium levels.

Of the 3 patients who failed to achieve normocalcemia, 1 underwent a subtotal 3½-gland parathyroidectomy in 2 operative procedures with transplantation and was noted to have marginal recurrent HPT 24 months postoperatively (calcium=2.63 mmol/L [10.5 mg/dL]; iPTH=8.53 pmol/L). A second patient underwent 3 operations throughout 7 years, with the removal of 3 glands, including a 10.4-g mediastinal parathyroid; the patient has persistent HPT. The last patient underwent removal of 2 enlarged glands; however, the right inferior parathyroid was not identified at operation. She has refused further operative intervention.

COMMENT

Parathyroidectomy is the only curative option, and the treatment of choice in patients with primary HPT. The benefit of parathyroidectomy as treatment of this condition has been well documented in multiple studies.⁵⁻⁷ In an attempt to provide adequate controls, Chan et al⁵ compared symptomatic and clinical outcomes in patients un-



Calcium and intact parathyroid hormone (iPTH) levels at long-term follow-up in patients undergoing subtotal parathyroidectomy. The dotted lines represent the upper limits of the normal level for calcium (2.63 mmol/L [10.5 mg/dL]) and iPTH (6.84 pmol/L). To convert calcium values to conventional measurements (milligrams per deciliter), divide the given values by 0.25.

dergoing parathyroidectomy, with patients undergoing thyroidectomy used as controls. At a mean follow-up of 20 months, the 121 parathyroid patients had fewer symptoms and lower rates of hematuria, bone fracture, joint swelling, weight loss, duodenal and gastric ulcerations, and hypertension. Burney et al⁶ also demonstrated significant improvement in functional status and patient perception of health status following parathyroidectomy for primary HPT. A recent nonrandomized prospective study of patients with primary HPT evaluated outcomes following parathyroidectomy vs close follow-up.⁷ Recommendations for resection in this study were based on guidelines from the National Institutes of Health Consensus Conference.⁸ Of the 61 patients undergoing parathyroidectomy in that study, 49 were asymptomatic and 12 were symptomatic with nephrolithiasis preoperatively. At 10 years' follow-up, biochemical parameters were normal, bone mineral density was significantly increased, and all 61 patients remained free of symptoms.

The debate regarding the appropriate operative intervention for sporadic primary multiple-gland disease has continued unabated for decades. The operative approaches include a routine 3½-gland resection (leaving approximately 50-100 mg of the most "normal-appearing" gland),⁹⁻¹¹ excising only those glands that are grossly enlarged at operative exploration,¹²⁻¹⁵ and a total 4-gland parathyroidectomy with subsequent autotransplantation.¹⁶⁻¹⁸ All 3 techniques have yielded excellent results in multiple series. Unfortunately, many of these reports are based on heterogeneous groups that include patients with solitary adenomas and familial disease. This heterogeneity substantially complicates analysis of postoperative outcomes in patients with sporadic multiple-gland disease. The paucity of information on this specific subset of patients led to the current study. In addition, there are very few long-term data on iPTH levels in patients undergoing parathyroidectomy for sporadic multiple-gland disease.

Results of studies reporting long-term outcomes for subtotal parathyroidectomy and including patients with all causes of multiple-gland disease are summarized in the first portion of **Table 3**. Of note, these study groups

Table 3. Long-term Outcomes of Patients Undergoing Subtotal Parathyroidectomy

Published Study (Year)	Study Years	No. of Patients	Mean Follow-up, mo	Postoperative Normocalcemia, %
All Causes of Multiple-Gland Disease				
Block et al ⁹ (1978)	1960-1975	29	...	79
Scholz et al ¹⁰ (1978)	1968-1972	49	...	73
Edis et al ¹¹ (1979)	1959-1976	55	...	82
Harness et al ¹⁹ (1979)	1971-1978	58	...	88
Proye et al ²⁰ (1998)	1971-1992	190	89	90
Sporadic Primary 4-Gland Hyperplasia				
Bondeson et al ²¹ (1985)	1960-1980	39	66	90
Wallfelt et al ²² (1990)	1956-1984	93	79	80
Tezelman et al ² (1993)	1982-1992	58	...	97
Sporadic Primary Multiple Adenomas				
Attie et al ²³ (1990)	1953-1990	31	75	93
Tezelman et al ² (1993)	1982-1992	49	70	100

*Ellipses indicate data are not reported.

include heterogeneous populations of patients with diagnoses including familial HPT, MEN syndromes, and secondary HPT.^{9-11,19,20} All of these authors followed a policy of selective subtotal parathyroidectomy, in which only grossly enlarged glands were excised, leaving grossly normal parathyroids in place. The percentage of patients who were rendered normocalcemic postoperatively ranged from 73% to 90%, with a trend toward improvement in the more recent series.^{19,20} Proye et al²⁰ reported the largest experience in this heterogeneous group to date, with excellent long-term outcomes. They concluded that “conservative surgery is an acceptable treatment for primary HPT due to multiglandular disease discovered at first cervicotomy.”

Other studies have examined outcomes following subtotal parathyroidectomy in more specific subgroups of patients with multiple-gland disease, including those with sporadic primary 4-gland hyperplasia^{2,21,22} and sporadic primary multiple adenomas^{2,23} (Table 3). The 80% to 100% postoperative rates of normocalcemia found in these patients were somewhat better than those for patients with all causes of multiple-gland disease (although these reports are more recent than the previous, more inclusive studies). Once again, a recurrent theme of these studies is operative conservatism (ie, removing only those glands noted to be grossly enlarged). Wallfelt et al²² found that persistent or recurrent hypercalcemia was more common following resection of 3 or 3½ glands than of 2 or 2½ glands, although the determination of extent of resection was not clearly defined. In addition, the incidence of postoperative normocalcemia was

significantly lower in patients with diffuse hyperplasia (80%) than in those with a single adenoma (91%).

The definition and presence of sporadic primary multiple adenomas have been widely debated topics in the literature. The intraoperative finding of 2 or 3 enlarged glands may represent multiple parathyroid adenomas or asymmetric 4-gland hyperplasia, with microscopic hyperplasia within the normal-appearing glands. Unfortunately, no histopathologic features can distinguish between a parathyroid adenoma and hyperplasia. Therefore, the diagnosis remains a clinical decision based on the findings of the operating surgeon. The distinction between these 2 diagnoses, however, may have little clinical import. Attie et al²³ and Tezelman et al² have reported long-term outcomes following resection of multiple adenomas with excellent normocalcemia rates of 93% to 100% at more than 70 months of follow-up. Both groups conclude that multiple adenomas do exist as an independent pathologic entity and recommend selective parathyroidectomy for cure. Because the clinical management remains identical for any degree of multiple-gland disease found at operation, all of these patients can be successfully treated with selective subtotal parathyroidectomy. Whether the remaining glands are normal or hyperplastic on a microscopic level seems to have no clinical ramifications.

Twenty percent of the patients in the current series underwent multiple operative procedures, primarily owing to recurrent hypercalcemia after previous parathyroidectomy elsewhere. This is not unexpected in this patient population because multiple-gland disease has been previously recognized as a risk for reoperative parathyroidectomy. Shen et al²⁴ found that 37% of failed parathyroidectomies reflected incomplete resection of multiple abnormal glands. In another review of reoperative cases, Weber et al²⁵ concluded that the most common causes of persistent or recurrent HPT were either inadequate neck exploration or inadequate resection of hyperplastic tissue. Similar to our own series, Doherty et al²⁶ noted that 7 of 8 patients who had undergone previous operation elsewhere had abnormal glands that were not resected at the initial operation. Clearly, patients with primary HPT secondary to multiple-gland disease are at risk for inadequate resection, and should undergo complete exploration by an experienced endocrine surgeon.

Of note, 14% of the patients for whom long-term follow-up was available had evidence of elevated iPTH levels and normal serum calcium levels. This phenomenon has been noted by other authors, including Mandal and Udelsman,²⁷ who reported secondary HPT in 12% of their patients following parathyroidectomy. These findings were noted in the immediate postoperative period and had largely resolved within 6 months, in contrast to the findings in our series. This secondary hyperparathyroid profile may be the result of remaining microscopic parathyroid hyperplasia or an alteration in the set point required to maintain a normal calcium level. Regardless, in no normocalcemic patient have we found this to be of clinical concern, and we do not recommend further intervention.

In conclusion, selective subtotal parathyroidectomy for patients with sporadic primary multiple-gland disease can result in excellent long-term normocalce-

mia with minimal complications. Based on analysis of a homogenous group of patients with primary sporadic multiple-gland disease found during cervical exploration, we recommend conservative resection of all grossly abnormal glands with biopsy of normal-appearing glands at operative exploration, or a 3½-gland resection in the case of 4-gland disease. In addition, a complete neck exploration and identification of all glands by an experienced surgeon is of paramount importance for optimal treatment of this group of patients.

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Reprints: Armando E. Giuliano, MD, John Wayne Cancer Institute, 2200 Santa Monica Blvd, Santa Monica, CA 90404 (e-mail: giulianoa@jwci.org).

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DISCUSSION

Theodore X. O'Connell, MD, Los Angeles, Calif: This paper makes an effort to separate the select subgroup from all other patients with parathyroid disease and tries to make specific comments and recommendations regarding them. I feel this is a very worthwhile undertaking, and expands our insight into the treatment of patients with this condition. I have several comments and questions for the authors. First, in the recent years there have been 2 schools of thought in doing parathyroid surgery. The first school is that every patient needs to have full neck exploration with identification and biopsy of all 4 glands. Proponents of this theory find multiple-gland disease in a significant number of patients, justifying this approach. The other school of thought is to use preoperative or intraoperative localization procedures and do a more directed minimal surgery just to remove the single adenoma, or perhaps explore the side where the adenoma is found to substantiate that there is also a normal gland on that side. Supporters of this approach state that while the full neck explorers may find more "multifocal adenomas," many of these may be clinically insignificant. This is based on the excellent short-term results with this more limited approach. Obviously, the authors are proponents of the first school of thought of complete neck exploration. I would like to ask them whether they still pursue this approach or are using the less invasive and more localized procedures, either with preoperative or intraoperative localization with radio nuclear medicine techniques. Since there are many surgeons who follow the more limited localization approach, I would ask the authors if there is anything in the preoperative workup of these patients with multiple-gland disease, such as types or degrees of symptoms, calcium levels, PTH levels, etc, which would indicate that they have multigland disease, and therefore, would perhaps not be a candidate for the limited localized procedures.

Second, although the failure rate in this series of 8% is quite low and comparable with other series, I would like to ask the authors what factors accounted for these failures, and what steps would have been taken in retrospect to eliminate them.

Third, I was particularly interested in the paper to see that at least some patients preoperatively had a normal calcium value since the range of preoperative calcium was from 9.1 to 13.4 mg/dL. I would like to ask the authors how many of these patients had normocalcemic hyperparathyroidism. It was also noted in the paper that approximately 14% of the patients postoperatively had normocalcemic hyperparathyroidism, that is, normal calciums with elevated PTH levels. Are these the same patients who had the condition preoperatively?

The final question regards this group of patients. Normocalcemic hyperparathyroidism with normal renal function usually is due to calcium-losing nephropathy. As calcium is lost in the urine, the serum calcium falls, and then as the normal homeostatic mechanism, increased PTH is released to correct to a normal serum calcium. After many years, the patients may actually develop parathyroid hyperplasia as a homeostatic re-

sponse. These patients can be identified as having normal serum calcium, elevated PTH, and elevated urinary calcium levels. They can be treated nonoperatively by the use of thiazide diuretics, which decreases the calcium loss by the kidney.

I would like to ask the authors if any of these patients with normocalcemic hyperparathyroidism either preoperatively or postoperatively were assessed for calcium-losing nephropathy, and, if this was found, how many were treated with thiazide diuretics and, if so, what was the response?

Eberhard Mack, MD, Madison, Wis: Could you tell us a little bit more about the history of those patients? Were these patients with multiple-gland disease exposed to high levels of radiation, or were some of these patients treated with lithium?

William Turner, MD, Jackson, Miss: Would the authors comment on any value for intraoperative parathormone assay now that they have completed this study?

Jon van Heerden, MD, Rochester, Minn: We have become quite enthusiastic about the intraoperative use of parathyroid hormone assays, but we have yet to find what the absolute indications are. Perhaps multiglandular disease is the indication for intraoperative PTH, and I wonder if the authors would comment.

As a cautionary note, any operation that has a 20% reoperation rate is not a good operation. Multiglandular disease continues to be problematic.

Any study that has a loss of 27% in regards to long-term follow-up has to be viewed with a little caution.

Claude H. Organ, Jr, MD, Oakland, Calif: Dr van Heerden has already expressed my concern about the 20% reoperation rate. Were those patients who were eucalcemic postoperatively also asymptomatic?

Norman W. Thompson, MD, Ann Arbor, Mich: I have the same philosophy about subtotal parathyroidectomy as the authors. However, their premise is that they were dealing with sporadic multiple-gland disease, and I would like to ask them how they have proven that. The follow-up isn't that long that MEN(I) could absolutely be ruled out, and that is the group most likely to recur. Finally, the most important figure in their paper is the zero incidence of permanent hypoparathyroidism. For that they are to be congratulated.

Richard A. Prinz, MD, Chicago, Ill: Was there any difference in the outcome in patients treated with true subtotal parathyroidectomy (in other words, 3½- or 4½-gland removal) compared with selective removal of enlarged glands? In my mind, those may be very different entities. There were 3 recurrences in your series. What were the initial operations in those 3 patients? Were they subtotal parathyroidectomies or selective removal of enlarged glands?

Dr Giuliano: I would like to thank the discussants, especially Dr O'Connell, for their excellent comments. First, let me address our approach to minimally invasive parathyroidectomy. We have recently started to perform that procedure. In patients who have a clearly defined hot spot on preoperative sestamibi scan, we find a very high rate of success using intraoperative gamma probe localization. You can predict that it will fail in patients who have poor localization or questionable findings on sestamibi scan. With a good hot spot, you are likely to have a successful minimally invasive procedure.

On our failures and on our reoperation rate, I think there may have been some misunderstanding. We had 10 patients who had reoperation. Eight of those were referred to us after prior exploration. So in our hands we had only 2 patients who were re-explored. One patient had 4 abnormal glands identified at operation. We did a 3½-gland resection only to find that the calcium did not fall. We went back and found a fifth gland. So our own reoperation rate is not as high as it would appear. You can often predict failures among patients with sporadic multigland disease. Those tend to occur in patients who have had prior operation, and patients in whom you do not personally identify all 4 glands.

Dr O'Connell, it is not clear to me what the persistent elevated PTH with normal calcium years after the operation signifies. Secondary hyperparathyroidism is reported following parathyroidectomy for primary hyperparathyroidism. It usually resolves in 6 to 12 months. These patients had longer follow-up and still had some persistently elevated hormone levels and normal calcium. Whether that represents some low-grade persistent hyperparathyroidism, some change in the set point for parathyroid hormone, or some secondary hyperparathyroidism is unknown.

Dr Mack, none of these patients had radiation. I am not sure about lithium. These are relatively straightforward patients, who were explored for hyperparathyroidism.

Intraoperative PTH may be of value in these patients. We have not started using this technique, and this may be a group of patients who would benefit from it.

Dr van Heerden, I think I commented on the fact that only 2 of these patients in our hands required a second operation.

Dr Organ, these patients were asymptomatic when their calcium and parathyroid levels were corrected. Most were asymptomatic preoperatively.

We eliminated patients with family history. I think some could have MEN syndrome, although we doubt that many could. We have had excellent long-term results, Dr Thompson. Thank you for your kind comments. Our 1 transient nerve injury was in a patient who had an incidental carcinoma of the thyroid who had a total thyroidectomy at the same time. Fortunately, that complication resolved and we are very pleased to have no hypoparathyroidism. We noted no difference in outcome, Dr Prinz, for patients who had selective operation and those who had a 3½-gland operation. However, we do not routinely do 3½-gland resections. We undertook this study because while rotating on my service, Dr Rose pointed out to me that my approach appeared to be a very subjective approach to the management of this often difficult problem and because one of our endocrinologists said point blank that hyperplasia is not a surgical disease because of poor published results. So we sought to find out our long-term results, especially with respect to parathyroid hormone levels, which has not been well studied previously. While there may be subjectivity in the determination of what is a normal-sized gland, I prefer, Dr Rose, the term "experience" rather than subjectivity. We believe our results show that the overwhelming majority of patients with sporadic multiglandular disease treated with selective parathyroidectomy have favorable long-term outcomes with normalization of both calcium and parathyroid hormone.