Original Article

Incidence of Hypocalcaemia in

Hypocalcaemia in Total Thyroidectomy

Patients Undergoing Total Thyroidectomy

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ABSTRACT

Objective: To determine the incidence of hypocalcaemia in patients undergoing total thyroidectomy.

Study Design: Cross-sectional study.

Place and Duration of Study: This study was conducted at the Department of General Surgery, Central Park Teaching Hospital, and Lahore from 1st March 2016 to 31st March 2019.

Materials and Methods: Two hundred and seventy eight patients who underwent total thyroidectomy for benign multinodular goiter and thyroid carcinoma after ethical committee approval. Euthyroid patients between the ages of 20 to 60 years without any gender bias having ASA class I, II and III were included in the study. Patients having concurrent parathyroid pathologies, redo surgery, coagulation disorders, renal failure and vitamin D deficiency were excluded. Post-operative hypocalcaemia was measure in immediate follow-up period and after 6 months.

Results: The mean age of patients was 42.22 ± 9.25 years. The frequency of transient hypocalcaemia was found to be 11.15% and permanent hypocalcaemia was 1.8%. Transient hypocalcaemia varied by statistically significant frequency amongst different age groups (p=0.000) while permanent hypocalcaemia did not vary significantly amongst different age groups (p=0.116). There was no association of development of transient or permanent hypocalcaemia with gender and diagnosis of the patients (p>0.05).

Conclusion: The frequency of transient hypocalcaemia and permanent hypocalcaemia following total thyroidectomy was found to be 11.15% and 1.8% respectively.

Key Words: Hypocalcaemia, Multinodular goiter, Thyroid carcinoma total thyroidectomy.

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INTRODUCTION

Total thyroidectomy has gradually cemented it place as the standard procedure of choice nowadays for the management of most patients presenting with multinodular goitre and thyroid carcinoma because of the increased recurrence rates with more conservative surgical options.¹

The parathyroid glands lie in close proximity to the thyroid gland and thus they can be inadvertently damaged during surgery of thyroid gland leading to hyperparathyroidism which principally manifests as Postoperative hypocalcaemia which is the commonest complication of total thyroidectomy these days.^{2,3}

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Received: June, 2019 Accepted: August, 2019 Printed: November, 2019 The incidence of transient hypocalcaemia has been reported to range from 1.6% to 71% while the incidence of permanent hypocalcaemia has been reported between 0.2% to 17%. ⁴⁻⁷ Hypocalcaemia results from damage to parathyroid glands which may result from parathyroid gland devascularisation, direct damage to the parathyroid glands during dissection, thermal damage, venous congestion, incidental Para thyroidectomy, local edema and non-visualization of parathyroid glands during surgery.^{8,9}

Hypocalcaemia can manifest clinically as carp pedal spasm, circumoral numbness, parenthesis, muscle cramps, stridor, lethargy, Trousseau sign, Chvostek sign, convulsions and cardiac arrhythmias. ¹⁰ Serum calcium levels and intact parathyroid hormone (iPTH) are the most common tests used for assessment of postoperative hypocalcaemia and hyperparathyroidism but due to the increased cost and decreased availability, measurement of iPTH has not been employed as a standard test. Moreover, serum calcium levels are a reliable predictor of determining hypocalcaemia and hyperparathyroidism after total thyroidectomy. ¹¹.

MATERIALS AND METHODS

We conducted this cross-sectional study on a total of 278 patients who were admitted to the Department of General Surgery, Central Park Teaching Hospital Lahore from 1st March 2016 to 31st March 2019 and

underwent total thyroidectomy operation. The inclusion criteria consisted of patients of either sex between the ages of 20 to 60 years; ASA Grade I to III; and diagnosed cases of carcinoma thyroid andmultinodular goitre having normal thyroid functions test undergoing Total Thyroidectomy. Patients having concurrent parathyroid abnormalities; chronic renal failure; vitamin D deficiency; hypertension; ASA Grade IV and V; deranged thyroid function tests and previous history of neck surgery including thyroid and parathyroid surgery were excluded from the study. Patients were operated on the elective operation list. The workup started with history, detailed clinical assessment and relevant laboratory investigations including thyroid profile and radiological investigations like ultrasound of neck. Fine needle aspiration cytology (FNAC) was requested from histopathology department of the hospital. The baseline serum calcium level was also recorded and only patients with normal calcium levels were included in the study. All the patients underwent total thyroidectomy by the conventional knot tie. The surgery was led by a professor or an associate professor who were assisted by a senior registrar or post graduate trainees.

Serum calcium levels were assessed in all patients daily till discharge from the hospital. Most patients were discharged on the third post-operative day. Serum calcium level less than 8 mg/dL at any instance was taken as hypocalcaemia. Patients will be given calcium and vitamin D3 supplement (calcium carbonate 1250mg; cholecalciferol 125IU, Qalsan D) one tablet twice daily. Patients were then followed for serum calcium levels at 3 and 6 months of surgery. The calcium levels were checked on follow up visits such that calcium supplements were not taken for 24 hours. Patients having serum calcium levels below 8 mg/dL after 6 months of surgery were labeled as cases of permanent hypocalcaemia while those in which hypocalcaemia settled were labeled as cases of transient hypocalcaemia. Follow up was ensured by taking contact numbers of all patients. The data was analyzed through SPSS-25.

RESULTS

The mean age of patients included in the study was 42.22±9.25 years.

Table No. 1: Stratification of transient and Permanent hypocalcaemia amongst different age groups

ge groups (years)		Total	P value			
20-40	41-60	Total	r value			
Transient hypocalcaemia						
2	29	31	0.000			
150	97	247				
Permanent hypocalcaemia						
1	4	5	0.116			
151	122	273				
1	20-40 pocalcae 2 150 ypocalcae	20-40 41-60 pocalcaemia 2 29 150 97 ypocalcaemia 1 4	20-40 41-60 10tal			

There was a female predominance with 212 out of 278 patients (76.26%) while 68 patients (23.74%) were male giving the female to male ratio of 3.2:1. Out of 278 total patients, 152 patients (54.68%) were between the ages of 20-40 years while 126 patients (45.32%) had ages between 41-60 years. Ninety one patients (32.73%) were classified as ASA Grade I, 142 patients (51.08%) were ASA Grade II, while the remaining 45 patients (16.19%) were classified as ASA Grade III respectively.

Table No. 2: Stratification of transient and permanent hypocalcaemia according to gender

Variable	Gender		Total	P value		
	Male	Female	Total	r value		
Transient hypocalcaemia						
	8	23	31	0.774		
	58	189	247	0.774		
Permanent hypocalcaemia						
	2	3	5	0.389		
	64	209	273	0.369		

Table No. 3: Stratification of transient and permanent hypocalcaemia according to diagnosis

Variable	Diagnosis		Total	Danalara			
	MNG	Carcinoma	Total	P value			
Transient hypocalcaemia							
	25	6	31	0.774			
	205	42	247	0.774			
Permanent hypocalcaemia							
	3	2	5	0.175			
	227	46	273	0.175			

As regards the diagnosis, 230 patients (82.73%) were cases of benign multinodular goiter while 48 patients (17.27%) cases undergoing total thyroidectomy were cases of thyroid carcinoma. The mean preoperative serum calcium level of all the patients included in the study was 9.25±0.54 mg/dL. The mean serum calcium level on the third post-operative day (day of discharge) was 8.67±0.67 mg/dL. The frequency of transient hypocalcaemia was calculated to be 11.15% (31 out of 278 patients), while the frequency of permanent hypocalcaemia came out to be 1.8% (5 out of 278 patients) respectively (Table Figs. 1-2). There is statistically significant difference in frequency of transient hypocalcaemia (p=0.000), however the frequency of permanent hypocalcaemia between different age groups was statistically non-significant (p=0.116) [Table 1]. There was no statistically significant difference in the frequency of transient hypocalcaemia (p=0.774) as well as permanent There was hypocalcaemia (p=0.389) [Table 2]. no statistically significant difference in the frequency of transient hypocalcaemia (p=0.744) as well as permanent hypocalcaemia (p=0.175) [Table 3].

DISCUSSION

Total thyroidectomy has become the surgical procedure of choice for both multinodular goiter and most cases of thyroid carcinoma and hypocalcaemia are one of the most common complications. ¹² Studies have reported a disparate results regarding the frequency of hypocalcaemia after total thyroidectomy.

In this study, the mean age of patients was found to be 42.22±9.25 years. A study from Karachi by Baloch et al¹³ reported a comparable mean age of 42.1 years. Edafo et al¹⁴ reported a mean age of 46.7 years, while a higher mean age of 50.7±15.9 was reported in a study by Baldassarre et al.¹⁵ Our study comprised of 76.26% female patients. Baloch et al¹³ reported a comparable frequency of female patients equal to 78.45%, while studies by Edafo et al¹⁴, Baldassarre et al¹⁵ and Basim et al¹⁶ reported the frequency of female gender to be 77.3%, 81.1% and 83.5% respectively.

In our study, 82.73% were diagnosed cases of benign multinodular goiter while the 17.27% patients had diagnosis of thyroid carcinoma. However in the study by Baloch et al¹³, thyroid carcinoma was the predominant diagnosis in 87.58% cases and only 12.41% patients were cases of benign diseases.

The frequency of transient hypocalcaemia was found to be 11.15% while the frequency of permanent hypocalcaemia i-e. Hypocalcaemia persisting for more than 6 months after surgery was observed in 1.8% patients. The frequency of transient hypocalcaemia was found to be significantly associated (p=0.000) with the age of the patients at the time of surgery as shown in Table 1. The odds of development of permanent hypocalcaemia were also greater in older patients. However there was no gender predisposition of developing postoperative hypocalcaemia after total thyroidectomy. Similarly the preoperative diagnosis had no bearing on the development of hypocalcaemia.

A study by Nair et al⁹ published in 2013 reported the frequency of transient hypocalcaemia following total thyroidectomy to be 23.6% which was higher than our result while the frequency of permanent hypocalcaemia was reported as 1.61% which was comparable to our study. Contrary to our results, the study reported that there was no difference in the frequency of transient hypocalcaemia (p=0.732) or permanent hypocalcaemia (p=0.332) between different age groups. Total thyroidectomy was also found to be significantly associated with the diagnosis of multinodular goiter for transient hypocalcaemia (p=0.02).

A similar study from Karachi by Baloch et al¹³ reported that the frequency of transient and permanent hypocalcaemia was 7% and 0.11% respectively. Another study by Basim et al¹⁶ from India in 2017 reported the incidence of hypocalcaemia to be 24.27%. The study also reported that the frequency of hypocalcaemia was greater in patients more than 50 years of age but the frequency didn't vary by a statistically significant proportion (p=0.35).

Baldassarre et al¹⁵ reported the incidence of hypocalcaemia in total thyroidectomy to be 9% which

was comparable to our study. However this study reported that postoperative hypocalcaemia was significantly associated with female gender (p<0.001, and malignancy (p<0.001).

Edafe et al¹⁷ reported the frequency of post-thyroidectomy transient and permanent hypocalcaemia as 29% and 5.5% respectively. ¹⁴Iqbal et al in 2010 reported the frequency of transient hypocalcaemia to be 21.6% with none of the patients developing permanent hypocalcaemia. Esimontas et al in 2018 from Lithuania reported a much higher frequency of post-operative hypocalcaemia in 64.2% patients. ¹⁸ While Ritter et al¹⁹ reported the frequency of transient and permanent hypocalcaemia to be 18% and 1.9% respectively.

The above mentioned studies highlight the wide disparity in the reported incidence of postoperative hypocalcaemia. Hypocalcaemia is an important postoperative complication following total thyroidectomy that develops due to hyperparathyroidism resulting from direct or indirect damage to the parathyroid glands. It not only increases the hospital stay with studies reporting an increase in stay up to 7 days but also increases the financial burden.²⁰

Various techniques have been reported by different studies for prevention of postoperative hypocalcaemia such as prophylactic postoperative calcium supplementation, identification and preservation of parathyroid glands and their vasculature, auto transplantation of parathyroid glands and use of magnification loupes for better visualization amongst others. ²¹⁻²⁴ Moreover there is an increased tendency of overprescribing calcium and vitamin D supplements in these patients which predisposes the patients to development of complications like renal stones etc. ²⁵

Thus we recommended that meticulous dissection technique aimed at preserving parathyroid glands should be employed to prevent hypocalcaemia after total thyroidectomy. Prophylactic administration of calcium and vitamin D supplements can prevent postoperative hypocalcaemia, however their prolonged use should only be prescribed in the light of serum levels of calcium since the overall frequency of permanent hypocalcaemia after total thyroidectomy is very less. Owing to the increased cost of iPTH assay, it was not done in our patients. Further research is recommended to find out frequency of hypocalcaemia following total thyroidectomy with the newer dissection techniques like LigaSure small jaw, Focus harmonic scalpel and laparoscopic thyroidectomy.

CONCLUSION

The frequency of transient hypocalcaemia and permanent hypocalcaemia following total thyroidectomy was found to be 11.15% and 1.8% respectively. We recommend the use of calcium and vitamin D supplements in the postoperative period to prevent the development of hypocalcaemia; however prolonged use of these supplements should be based on serum calcium levels.

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Conflict of Interest: The study has no conflict of interest to declare by any author.

REFERENCES

Revisiting Critically:

- Gharib H, Papini E, Garber JR, Duick DS, Harrell RM, Hegedüs L, et al. American Association of Clinical Endocrinologists, American College of Endocrinology, and Associazione Medici Endocrinologi Medical Guidelines forClinical Practice for the diagnosis and management of thyroid nodules - 2016 update. Endocr Pract 2016;22(5):622-39.
- Chang YK, Lang BHH. To identify or not to identify parathyroid glands during total thyroidectomy. Gland Surg 2017;6(Suppl 1):S20–9
- 3. Sitges-Serra A, Gómez J, Barczynski M, Lorente-Poch L, Iacobone M, Sancho J. A nomogram to predict the likelihood of permanent hypoparathyroidism after total thyroidectomy based on delayed serum calcium and iPTH measurements. Gland Surg 2017;6(Suppl 1):S11–9
- Chahardahmasumi E, Salehidoost R, Amini M, Aminorroaya A, Rezvanian H, Kachooei A, et al. Assessment of the early and late complication after thyroidectomy. Adv Biomed Res 2019;8:14
- Seo ST, Chang JW, Jin J, Lim YC, Rha KS, et al. Transient and permanent hypocalcemia after total thyroidectomy: Early predictive factors and longterm follow-up results. Surg 2015; 158(6):1492–9.
- 6. Wu J, Harrison B. Hypocalcemia after thyroidectomy: the need for improved definitions. World J End Surg 2010;2:17-20.
- Langner E, Tincani AJ, Negro AD. Use of prophylactic oral calcium after total thyroidectomy: a prospective study. Arch Endocrinol Metab 2017;61(5):447-54
- Malik MZ, Mirza AA, Farooqi SA, Chaudhary NA, Waqar M, Bhatti HW. Role of preoperative administration of vitamin D and calcium in postoperative transient hypocalcemia after total thyroidectomy. Cureus 2019;11(4):e4579
- Nair CG, Babu MJ, Menon R, Jacob P. Hypocalcaemia following total thyroidectomy: An analysis of 806 patients. Indian J Endocrinol Metab 2013;17(2):298–303
- Hadker N, Egan J, Sanders J, Lagast H, Clarke BL. Understanding the burden of illness associated with

- hypoparathyroidism reported among patients in the PARADOX study. Endocr Pract 2014;20(7):671-9.
- 11. Noureldine SI, Genther DJ, Lopez M, Agrawal N, Tufano RP. Early predictors of hypocalcemia after total thyroidectomy: an analysis of 304 patients using a short-stay monitoring protocol. JAMA Otolaryngol Head Neck Surg 2014;140(11):1006–13
- 12. Makay Ö. Less than total thyroidectomy for goiter: when &how? Gland Surg 2017;6(Suppl 1):S49–58
- 13. Baloch N, Taj S, Anwer M, Naseem M. Frequency of hypocalcaemia following total thyroidectomy. Pak J Med Sci 2019;35(1):262–5
- 14. Edafe O, Prasad P, Harrison BJ, Balasubramanian SP. Incidence and predictors of post-thyroidectomy hypocalcaemia in a tertiary endocrine surgical unit. Ann R Coll Surg Engl 2014;96(3):219–23
- 15. Baldassarre RL, Chang DC, Brumund KT, Bouvet M. Predictors of hypocalcemia after thyroidectomy: results from the nationwide inpatient sample. ISRN Surg 2012;2012:838614
- Basim M, Abdul Siyad AK. Incidence of hypocalcemia following total thyroidectomy and its relation with age. IOSR J Dental Med Sci 2017;16(6):19-23.
- 17. Iqbal M, Subhan A, Baig SM, Shah SM. Frequency of hypocalcaemia in total thyroidectomy. J Surg Pak (Int) 2010;15(2):87–9.
- Eismontas V, Slepavicius A, Janusonis V, Zeromskas P, Beisa V, Strupas K, et al. Predictors of postoperative hypocalcemia occurring after a total thyroidectomy: results of prospective multicenter study. BMC Surg 2018;18(1):55.
 Ritter K, Elfenbein D, Schneider DF, Chen H,
- 19. Ritter K, Elfenbein D, Schneider DF, Chen H, Sippel RS. Hypoparathyroidism after total thyroidectomy: incidence and resolution. J Surg Res 2015; 197(2): 348–53.
- 20. Grainger J, Ahmed M, Gama R, Liew L, Buch H, Cullen RJ. Post-thyroidectomy hypocalcemia: Impact on length of stay. Ear Nose Throat J 2015;94(7):276-81.
- 21. Xing T, Hu Y, Wang B, Zhu J. Role of oral calcium supplementation alone or with vitamin D in preventing post-thyroidectomy hypocalcaemia: a meta-analysis. Medicine (Baltimore) 2019;98(8): e14455.
- 22. Maxwell AK, Shonka DC Jr, Robinson DJ, Levine PA. Association of preoperative calcium and calcitriol therapy with postoperative hypocalcemia after total thyroidectomy. JAMA Otolaryngol Head Neck Surg 2017;143(7):679–84.
- 23. Park I, Rhu J, Woo JW, Choi JH, Kim JS, Kim JH. Preserving parathyroid gland vasculature to reduce post-thyroidectomy hypocalcemia. World J Surg 2016;40(6):1382-9.
- 24. Antakia R, Edafe O, Uttley L, Balasubramanian SP. Effectiveness of preventative and other surgical measures on hypocalcemia following bilateral thyroid surgery: a systematic review and meta-analysis. Thyroid 2015;25(1):95-106.
- 25. Al-Dhahri SF, Mubasher M, Mufarji K, Allam OS, et al. Factors predicting post-thyroidectomy hypoparathyroidism recovery. World J Surg 2014;38(9):2304-10