The Effect of Standardized Order Set for Intravenous Calcium Treatment on the Length of Hospital Stay in Secondary Hyperparathyroidism following Parathyroidectomy. Retrospective Pre-Post Study

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Abstract

Objective: The objective of this study is to assess if administering a post-operative standardized order set for intravenous calcium treatment for dialysis patients with secondary hyperparathyroidism who have undergone a parathyroidectomy will reduce the length of the hospital stay.

Methods: The medical records of dialysis patients who underwent subtotal PTX for symptomatic secondary hyperparathyroidism were reviewed prior to and after the introduction of a standard postoperative order set. The genders, duration of hospital stay as well as readmission rates of all patients were analyzed.

Results: The study population included 50 patients, with a mean age of 49 years. 25 patients were administered the standardized order set of intravenous calcium treatment following a subtotal parathyroidectomy. There was no statistical difference (p=0.7063) between the length of hospital stay in the group of the patients that received the post-operative standardized order set of intravenous calcium and the group that did not. Only 1 patient was readmitted during the study period.

Conclusion: There was no statistical difference in the length of hospital stay between the dialysis patients with secondary hyperparathyroidism who received the standardized order set of intravenous calcium and the patients who did not following a PTX.

Keywords: Secondary hyperparathyroidism; Parathyroidectomy; Intravenous calcium; Length of hospital stay

Abbreviations: SHPT: Secondary Hyperparathyroidism; PTH: Parathyroid Hormone Levels; KDOQI: Kidney Disease Outcomes Quality Initiative; PTX: Parathyroidectomy; Alkp: Alkaline Phosphatase

Introduction

Secondary hyperparathyroidism (SHPT) commonly develops in patients with chronic kidney disease [1-17]. SHPT is characterized by parathyroid hyperplasia and overproduction of the parathyroid hormone, resulting in hypercalcemia and hyperphosphatemia [4,6-8,11,15-17]. As a result of the calcium and phosphate imbalance, SHPT is commonly associated with other co morbidities, including skeletal and cardiovascular diseases, leading to increased mortality [2,4,7,8,10,16,18].

The majority of patients with hyperparathyroidism are medically managed by phosphate binders, vitamin D analogues and/or calcimimetics in order to reduce the phosphate and parathyroid hormone (PTH) serum levels. However, according to the Kidney disease Outcomes Quality Initiative (KDOQI) guidelines, in cases refractory to medical treatment, parathyroidectomy (PTX) is recommended [1-7,11-17,19]. Three surgical parathyroidectomy approaches have been developed: subtotal parathyroidectomy, total parathyroidectomy with parathyroid autotransplantation, and total parathyroidectomy without autografting [11,12]. Subtotal PTX or total PTX with autotransplantation are the preferred approaches [3,5,11,12].

Post-operative hypocalcaemia, a result of reduced PTH serum levels, is a complication that is frequently encountered following PTX [15,18]. Rapid, severe and prolonged hypocalcaemia, known as hungry bone syndrome, commonly develops in patients with SHPT and severe bone disease [1,2,7,19-21]. The duration of hospital stay is typically prolonged these patients [1,5]. To alleviate this complication, intravenous calcium is given following the PTX [1-3,6,15,20]. The purpose of this study is to assess if administering a post-operative standardized order set of intravenous calcium treatment for dialysis patients with secondary hyperparathyroidism who have undergone a PTX will reduce the length of the hospital stay.

Methods

Patients

In this retrospective pre-post study, 50 patients' charts with chronic kidney disease that underwent subtotal PTX by a single general surgeon in Saskatoon, SK between April 2010 and May 2015 for symptomatic secondary hyperparathyroidism were reviewed. Patients with primary and tertiary hyperparathyroidism were excluded from the study. Research Ethics Board approval was obtained for this study. The information abstracted in this study included, gender and age of the patients, the number of patients who received standardized orders of intravenous calcium, length of hospital stay and readmission rates. The length of hospital stay was defined as the number of days in the hospital from the date of surgery to the date of discharge.

The patients were divided into two study groups: one that received post-operative standardized order set for intravenous calcium treatment...
(Table 1), and one that did not. Patients’ gender, age, and length of hospital stay between the two groups were compared.

**Statistical analysis**

We checked all variables regarding assumptions underlying the use of parametric and nonparametric statistics, and analyzed them accordingly. We performed Student’s t-test, for the normally distributed variable (age), and Wilcoxon test for the non-normally distributed variable (length of stay). Chi-square test was used to examine the association between categorical variables. Data were analyzed using SAS software (v 9.4; SAS Cary, NC, USA). The level of significance was set at 0.05 (two-tailed).

**Results**

The study population included 18 females (36%) and 32 males (64%), with a mean age of 49 years, a median of 51.5 years, and a range of 20-75 years. The mean length of hospital stay was 5.3 days with a median of 4.0 days. Of the study population, only 1 (2%) patient was readmitted. 25 patients (50%) were treated with a standardized order set of intravenous calcium post-operatively. The patients’ gender, age, length of stay and readmission rates and whether or not they received a standardized order set are outlined in Table 2.

Our study revealed no statistical difference (p>0.05) in the length of hospital stay (p=0.7063) between the group of patients that received the post-operative standardized order set of intravenous calcium (mean of 4.88 days) and the group that did not (mean of 5.72 days) (Table 3).

There were no significant differences in regards to age (p=0.1897) and sex (p=0.5557) when comparing the group of patients that received the standardized order set of intravenous calcium post-operatively and the group that did not. Outlined in Table 3.

<table>
<thead>
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<th>Parameter</th>
<th>Given</th>
<th>Not Given</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male/Female, %</td>
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<td>46.88/55.56</td>
<td>0.5557</td>
</tr>
<tr>
<td>Age</td>
<td>46.48±11.45</td>
<td>51.60±15.46</td>
<td>0.1897</td>
</tr>
<tr>
<td>Length of hospital stay</td>
<td>5.72±5.56</td>
<td>4.88±2.82</td>
<td>0.7063</td>
</tr>
</tbody>
</table>

Table 3: Comparison of the parameters in patients who were administrated the post-operative standardized order set of intravenous calcium and the group that did not.

**Discussion**

In dialysis patients, secondary hyperparathyroidism is a common complication requiring surgical PTX if refractory to medical treatment. PTX drastically improves SHPT [14]. PTX has been shown to improve bone mineral density and reduce the incidence of fractures by 30% in patients receiving dialysis for chronic kidney disease [2,20]. Moreover, PTX is cost-effective, increases survival rates and improves quality of life [14]. In this study, all the patients received a subtotal PTX.

Despite of the benefits of the surgical procedure, postoperative hypocalcaemia is a common and serious complication. It results from calcium metabolism imbalance due increased absorption of calcium from the circulation into the bones, and cessation of osteoclastic activity and continued osteoblastic activity [1,7,18-21]. Postoperative hypocalcaemia is seldom symptomatic unless serum calcium levels fall below 2 mmol/L [18]. When it becomes symptomatic, hypocalcaemia can present as seizures, neuromuscular symptoms, such as numbness, tingling and muscle cramps, and may predispose to ventricular arrhythmias and sudden cardiac death. Postoperative hypocalcaemia and the associated clinical manifestations can be prevented through strict monitoring of the calcium serum levels, and administration of prophylactic intravenous calcium infusion and vitamin D analogues [1,3,18,19,21].

There is inconsistency regarding the management of postoperative hypocalcaemia. Cozzolino et al. has proposed initiating calcium infusion when a rapid decline in serum calcium is noted. Tominaga et al. suggested intravenous calcium treatment when Alkaline phosphatase (AlkP) levels are >500 IU/L. Cheng et al. recommended a preventive approach to start calcium infusion immediately following surgery in any patients who have a preoperative AlkP level of >200 IU/L. The KDOQI guidelines suggest that calcium infusion should begin when the blood levels of corrected total calcium fall below 7.2 mg/dl. However, administrating intravenous calcium immediately postoperatively may prevent hypocalcaemia altogether [1].

Hypocalcaemia following PTX can prolong the length of hospital stay. A prolonged length of stay can increase the risk of morbidity and mortality, and lead to increased financial burden on the healthcare system [2]. Recent studies have concluded that the mean length of hospital stay following a PTX for secondary hyperparathyroidism is 3.0 to 9.2 days, which is similar to our mean stay of 5.3 days [1]. This study investigated if administrating a standardized order of intravenous calcium following a PTX would reduce the length of hospital stay. Our data showed that there was no statistical significance between the group of patients that received the post-operative standardized order set of intravenous calcium and the group of patients that did not, even when variables including age and gender were accounted for. This could have been largely due to limitations of the study, including a small sample size and a non-randomized study design. Further studies are warranted to assess postoperative

hypocalcaemia in the setting of PTX, including studying a larger sample size, utilizing various order sets, and taking into account patients' co-morbidities.

Other studies have formulated various guidelines for treatment and prevention of postoperative hypocalcaemia following a PTX [1,18,19,21]. However, analysis of the effects of a standardized order set of intravenous calcium immediately following parathyroidectomy in dialysis patients with secondary hyperparathyroidism has not been previously done. Thus, we wanted to look at a standardized order set that would reduce the patient's length of hospital stay. Although our study did not find a significant difference between the group of patients who received the standardized order set and the group that did not, we have elucidated areas and topics for further research.

Conclusions

The management of postoperative hypocalcaemia following a parathyroidectomy in dialysis patients remains poorly defined [3,19]. Our study did not reveal that dialysis patients with secondary hyperparathyroidism receiving a postoperative standardized order set of intravenous calcium following a parathyroidectomy would have a shortened the hospital stay. This could be due to the inadequate sample size and other metrics that were not analyzed, such as number of inpatient episodes of significant hypocalcaemia, nursing time, patient co-morbidities, and perioperative and other postoperative complications.

Conflict of Interest

No conflict of interest.

References