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Assessing the Role of Antibiotic Therapy in Transoral Endoscopic Thyroidectomy Vestibular Approach

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Abstract

Background: The transoral endoscopic thyroidectomy vestibular approach (TOETVA) is a remote access endoscopic approach that results in the disruption of the mucosa in the oral cavity. Despite its growing popularity, consensus has not been reached regarding the optimal type and duration of perioperative antibiotics.

Methods: We conducted a literature review examining the use of perioperative antibiotics during TOETVA. We compared the infection rate in studies that used only prophylactic antibiotic regimens (< 24 hours), with extended antibiotic regimens (> 24 hours).

Results: A PubMed literature review yielded 268 reports after searching for TOETVA or Transoral Thyroidectomy. Twenty-seven of those mentioned the antibiotic regimen of their series and were included in this analysis. In total, 1887 TOETVA cases were included. There was no significant difference in rate of infections between the two groups.

Conclusion: Based on our review, we do not see an increased risk of infection when only prophylactic antibiotics are used.

Keywords

TOETVA, Thyroidectomy, Antibiotics, Prophylaxis, Transoral thyroidectomy

Introduction

The traditional open cervical thyroidectomy is considered a clean procedure due to the absence of microbiologic contamination of the surgical field, especially as there is no disturbance of mucosa of the respiratory or gastrointestinal tracts [1]. These procedures have a very low rate of surgical site infection (SSI), estimated in the literature to be around 0.3% and 0.6% [2]. For this reason, antibiotic prophylaxis (pABX) is not suggested for the trans-cervical approach to endocrine surgery by most national and international guidelines [1,2]. Despite this, many surgeons worldwide continue to routinely use pABX. The debate as to whether the use of pABX is required is ongoing. While a large number of surgical societies recommend against antibiotic use in general (the caveat being that patients are not immunocompromised and/or are free of certain co-morbidities), its use is significantly influenced by social and geographic factors. For example, in England and Ireland 9% of patients receive routine antibiotic prophylaxis, whereas in Italy and China 38.7% and 57.4-100% receive pABX, respectively [2-5]. Without evidence-based oversight, the overuse of antibiotics in certain regions will invariably lead to negative repercussions such as an increase in antibiotic resistance and unnecessary costs [6].

The transoral endoscopic thyroidectomy vestibular approach (TOETVA) involves the removal of whole or part of

the thyroid gland from three small incisions in the mucosa of the mouth. The transoral thyroidectomy was first approached through a sublingual route but later this technique was abandoned due to complications related to the violation of the floor of the mouth [7-9]. The transoral technique has evolved to using a vestibular approach and modifications of this technique gained popularity for the treatment of thyroid disease. This route entails the placement of three endoscopic ports in the vestibular area of the mouth with dissection carried out beneath the platysma muscle, first over the chin, then down between the two strap muscles [10-12].

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The mucosa of the oral cavity is colonized with a diverse bacterial flora. Because of the violation of the oral mucosa, these procedures are classified as clean contaminated cases (having controlled contamination of the operative field) assuming that there is no concomitant infectious process at the time of surgery (e.g. tooth abscess, etc.) [1,13,14]. In clean contaminated cases as well as in the transoral vestibular approach, unless otherwise indicated, only antibiotic prophylaxis is recommended.

A clarification between antibiotic prophylaxis and treatment is necessary in order to clearly understand the difference between the two. Antibiotic prophylaxis is when administration is within 1 hour of incision time and up to 24 hours after the operation, whereas antibiotic treatment is the administration after an infection has already begun [14]. Often, the administration of pABX requires a single dose within 60 minutes of incision time to provide adequate tissue concentration throughout the operation. Additional doses during the procedure are advisable if surgery is prolonged (i.e. greater than 4 hours), if there is major blood loss, or an antimicrobial with a short half-life is used [14]. The antibiotics should be discontinued no later than 24 hours after surgery, as the prolonged use of prophylaxis leads to bacterial resistance and increased hospital costs [14].

While the surgical technique of TOETVA has been refined throughout the years since its inception, the use of perioperative antibiotics is widely variable throughout institutions. The purpose of this study is to review the current state of antibiotic use during TOETVA in the available literature to see if there is a difference in the rate of infection compared with length of antibiotic uses. The hope is that this can contribute to appropriate antibiotic recommendations in the future.

Methods

A comprehensive literature review was carried out on PubMed, using the search terms "TOETVA" or "Transoral Thyroidectomy". Only articles written in English were included. A total of 268 results were reviewed. Inclusion criteria included greater than five cases in a series, and those

studies which described IV or oral antibiotics duration and infection rate. Exclusion criteria included less than 5 cases in a study, cadaver or animal studies, studies without mention of antibiotics or infections. Full exclusion process can be seen in Table 1. Type of antibiotics used was recorded when available, as well as specific infectious complications and how they were managed. Results were examined to determine differences outcome's between patients receiving prophylactic antibiotics versus extended antibiotics. Prophylactic antibiotics were defined as < 24 hours of antibiotics, whereas extended antibiotics (eABX) were defined as > 24 hours of antibiotics. Statistical analysis was performed using SAS. P < 0.05 was considered statistically significant.

Results

Out of the 268 studies reviewed, 27 studies were included in the analysis [15-39]. One of the studies was a prospective randomized trial examining prophylactic vs extended antibiotic use in TOETVA [38]. One study compared four different groups of patients receiving different antibiotic regimens [37]. A total of 29 different groups were examined which translates to a total of 1887 TOETVA cases. Five hundred and four cases (41%) received pABX only. One thousand three hundred and eighty-three cases received eABX (59%). In the extended group, the average time of IV antibiotics was 2 days. The average length of oral antibiotics was 3.2 days.

A total of 15 infections were noted, 6 from the pABX group (1.1%) and 9 from the extended antibiotic group (0.6%). There was no statistically significant different between the rate of infections in the two groups (Table 2). Specified infections included chin cellulitis, pharyngitis, abscess and infected seroma. Two abscesses required surgical drainage in the operating room. One seroma was treated with needle aspiration and antibiotics. The management of the other infections was not specified.

The most common IV antibiotic regimens were amoxicillin/clavulanic acid; cefazolin and metronidazole; and clindamycin. The most common oral antibiotic regimen, if specified, was amoxicillin/clavulanic acid. A more complete list of antibiotic regimens can be seen in Table 3.

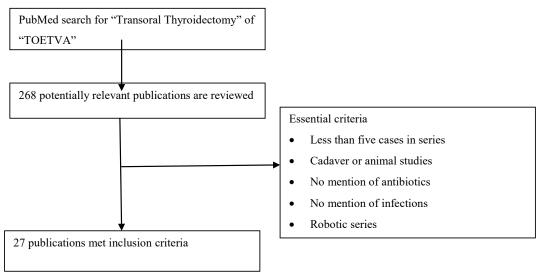


Table 1: Exclusion criteria.

Table 2: Differences in Antibiotic Duration and Rates of Infection Between the Prophylactic Only and Extended Antibiotic Groups.

	Prophylactic Antibiotics Only	Extended Antibiotics	Total	P-value
Number of Studies	11	16	27	
Number of Patients	504 (26.7%)	1383 (73.3%)	1887	
Average IV antibiotic time (days, standard deviation)	1	2 (1.6)	2	
Average PO antibiotic time (days, standard deviation)	0	5.1 (2.6)	3.1	
Number of Infections	6 (1.2%)	9 (0.7%)	15 (0.8%)	0.28

IV: intravenous; PO: per os

Table 3: Most Commonly Listed IV and PO Antibiotics.

Most Common IV Antibiotics	
Amoxicillin/Clavulanic Acid	
Ampicillin/Sulbactam	
Ancef and Metronidazole	
Cefazedone	
Cefotetan	
Clindamycin	
Oxacephem	
Most Common PO antibiotics	
Amoxicillin/Clavulanic Acid	

There was one prospective trial comparing prophylactic antibiotics only to extended antibiotics in TOETVA [38]. Twenty-five patients in each group participated. Both groups received a second-generation cephalosporin pre-operatively. The extended antibiotic group received amoxicillin/clavulanic acid orally for 7 days post-operatively. There were no surgical site infections in either group.

Discussion

In the 12 years since its origin in 2008, TOETVA and its development have been of great interest to the endocrine surgery community, with many favorable reports of its use throughout the world. Well received by patients and surgeons alike, the technique has been refined and improved. One area that requires further development is the topic of perioperative antibiotics in TOETVA. Prolonged antibiotic use when not indicated will lead to an increase in antibiotic resistance and unnecessary costs [6]. While many surgeons have moved to only using prophylactic antibiotics (less than 24 hours), the original methodology described multiples days of IV and oral antibiotics [10]. In our review, we found that 73% of patients who underwent TOETVA were given extended duration of antibiotics. This is despite the fact that there was no increase in the rate of infections when the length of antibiotic therapy is decreased to prophylactic antibiotics only.

The difference in duration of antibiotic use is partially socio-cultural as well. Two series compared the length of stay of TOETVA patients in the US and Italy, and found to have no increased complications when sending patients home on day of surgery [25,39]. In Asia it can be more common to have patients stay in the hospital longer before transitioning to home care. Here we see longer IV and oral antibiotic use [17].

The oral flora is composed of 750 different types of bacteria, ranging from aerobic gram-positive cocci to gram-negative anaerobes [1]. Therefore, antibiotic choice for both prophylaxis and therapy should strive to cover a broad range of gram positive and negative aerobes and anaerobes. In our study, we saw that all institutions used broad spectrum antibiotics, most commonly second generation cephalosporins with or without metronidazole, amoxicillin/clavulanic acid, or ampicillin/sulbactam. If oral antibiotics were used, only amoxicillin/clavulanic acid was listed.

In our literature review, only one prospective randomized trial was found to address antibiotic use in TOETVA [38]. Fifty patients were split into two equal groups. The control group which would receive IV antibiotics at the time or surgery as well as seven days of amoxicillin/clavulanic acid, and the experimental group which would receive only IV antibiotics at the time of surgery. Primary endpoint was signs of infection on the first post-operative day. They did not find a difference in the rate of infection between the two groups. They did find that patients in the antibiotic treatment group had an incidence of gastrointestinal upset not seen in the prophylactic only group, which they conclude is from the antibiotics.

One area of further study would be to examine the use of post-operative drains in TOETVA. The most cited technique for TOETVA calls for the use of post-operative drains, but some centers no longer use them, or no longer use them for all cases. Drain use is intended to decrease fluid collection which can become infected, but their impact on surgical site infection and antibiotic duration ("covering the drain") is not entirely clear [40]. Their use and effect on seroma formation or infection rate was not specifically examined in this study.

Conclusion

Based on current guidelines, no pABX are necessary for the traditional cervical approach to a thyroidectomy. Similar guidelines have not been set for TOETVA. Based on its clean-contaminated wound class, it would be reasonable to only use pABX for TOETVA. In our literature review we see up to 73% of patients undergoing TOETVA will receive an extended duration of antibiotics. Based on our review, we do not see an increased incidence of infection when only pABX are used. As the popularity of TOETVA continues to grow around the world, it is necessary to have a better understanding of what length of antibiotic therapy is necessary to help decrease the risk of antibiotic resistance and decrease costs.

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