Endoscopic surgical treatment of epistaxis in hereditary haemorrhagic telangiectasia: our experience

Il trattamento endoscopico dell’epistassi nella teleangectasia emorragica ereditaria: la nostra esperienza

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SUMMARY
Objectives. Hereditary haemorrhagic telangiectasia (HHT) is a rare autosomal dominant disease characterised by epistaxis. Surgical procedures for epistaxis vary from diathermo-coagulation to nasal closure. The aim of this paper is to report our experience in endoscopic surgical management of epistaxis in HHT patients.

Methods. This is a descriptive, longitudinal study carried out at the Otorhinolaryngology Department of IRCCS Policlinico San Matteo in Pavia, a reference centre for the treatment and diagnosis of HHT. We retrospectively evaluated HHT patients who underwent surgery for epistaxis from 1996 to 2015, including only those treated with endoscopic surgery.

Results. Among the 591 patients hospitalised and screened for HHT, 323 (54.7%) underwent endoscopic surgery for epistaxis, for a total of 679 procedures. General anaesthesia was used in 77.2% of procedures; argon plasma coagulation was the instrument of choice in the majority of patients, followed by lasers and quantum molecular resonance technology.

Conclusions. We report one of the largest cohorts undergoing endoscopic treatment of epistaxis in HHT patients. This mini-invasive surgical treatment allowed us to control epistaxis without major complications and nasal packing and can be repeated over time. For these reasons, we recommend it as first choice in case of epistaxis in HHT patients.

KEY WORDS: hereditary haemorrhagic telangiectasia, Rendu Osler Weber disease, epistaxis, nosebleeds, endoscopy, argon plasma coagulation

RIASSUNTO
Obiettivi. L’obiettivo di questo studio è riportare la nostra esperienza nel trattamento endoscopico dell’epistassi nei pazienti con Teleangectasia Emorragica Ereditaria (HHT).

Metodi. Si tratta di uno studio longitudinale retrospettivo svolto presso l’UOC di Otorinolaringoiatria della Fondazione IRCCS Policlinico San Matteo di Pavia, centro di riferimento per la diagnosi e la cura dell’HHT. Sono stati valutati i pazienti sottoposti dal 1996 al 2015 a trattamento chirurgico delle epistassi ricorrenti, includendo solo quelli trattati con tecnica endoscopica.

Risultati. Dei 591 pazienti ricoverati e sottoposti a screening per HHT, 323 (54.7%) sono stati sottoposti a trattamento chirurgico endoscopico per epistassi, per un totale di 679 procedure. Il 77,2% delle procedure è stato eseguito in anestesia generale; il sistema ad Argon Plasma è stato lo strumento di scelta nella maggior parte dei pazienti, seguito dai laser e dalla quantum molecular resonance.

Conclusioni. Riportiamo una delle più ampie casistiche nel trattamento endoscopico dell’epistassi nei pazienti con HHT. Tale approccio consente di gestire i pazienti senza necessità di tamponamento nasale, è scelto da complicanze maggiori, ripetibile nel tempo e riteniamo che debba essere preso in considerazione come trattamento di prima scelta in caso di epistassi in pazienti con HHT.

PAROLE CHIAVE: teleangiectasia emorragica ereditaria, malattia di Rendu Osler Weber, epistassi, endoscopia, argon plasma
Introduction

Hereditary haemorrhagic telangiectasia (HHT) is an autosomal dominant disorder characterised by multiorgan systemic vascular dysplasia. The prevalence of the disease is estimated to be between 1:5000 and 1:8000 in the European population. HHT is characterised by a highly variable phenotype and complete penetrance by the age of 40 years; vascular dysplasia leads to visceral arteriovenous malformations (AVMs) and mucosal and skin telangiectasias. Recurrent epistaxis from telangiectasias of the nasal mucosa is the most frequent symptom, which is present in more than 95% of HHT patients. AVMs are frequently found in the lungs (15-50% of patients), liver (32-78%) and brain (23%) and should be recognised in these patients since they may cause severe life-threatening complications. The clinical diagnosis of HHT is confirmed if three of the four diagnostic Curacao criteria (positive family history, recurrent epistaxis, mucocutaneous telangiectasias, visceral AVMs) are met. Two major genes are involved in HHT: ENG (OMIM #187300) and ACVRL1 (OMIM #600376), coding for proteins involved in the TGF-β/BMP pathway.

In HHT patients, the clinical manifestations of epistaxis are variable, from rare blood spots to patients in whom nosebleed severity can require frequent blood transfusions, with a substantial worsening in quality of life. Over the years, different grading scale systems have been proposed to quantify the severity of epistaxis related to HHT. Moreover, in recent years, to enhance the visualisation of nasal vascular structures, our group developed new methods of intraoperative endoscopy based on narrow-band imaging and fluorescein-guided endoscopy. However, epistaxis in HHT remains a clinically unsolved problem; current therapies do not lead to a definitive resolution but only to temporary control of the condition. Multiple therapeutic approaches have been proposed in the literature and can be schematically classified into the destruction of telangiectasia, reduction of blood flow, reduction of trauma, improvement of protection and control of fibrinolytic activity.

Regarding medical therapy, different approaches have been proposed and described with variable results. From a surgical perspective, the management of epistaxis depends on clinical severity. In patients with mild nosebleeds, various forms of mini-invasive surgical techniques including bipolar cautery, argon plasma coagulation (APC), lasers and Diego-PK Shaver treatment can be indicated, while more invasive procedures, such as modified Young’s procedure, are reserved for cases of severe epistaxis. Various therapeutic approaches for epistaxis in HHT have been described, but there still is no general agreement on the best treatment. The purpose of this study was to describe our experience in the endoscopic management of epistaxis in HHT patients, focusing on mini-invasive surgical procedures.

Materials and methods

The Department of Otorhinolaryngology of I.R.C.C.S. Policlinico San Matteo in Pavia is a national reference centre for diagnosis and treatment of HHT. After Local Ethics Committee approval (Comitato Etiico di Pavia, reference number 1-29/1/14), we retrospectively reviewed data of HHT patients treated from 1996 until 2015 for nosebleeds, including demographics, surgical technique, number of procedures carried out on each patient, time between each procedure, characteristics of nasal telangiectasias, presence of septal perforation and intraoperative bleeding. Patient records were stored in a Filemaker Pro Advance database (version 14.0.1, Filemaker Inc.) and extracted for this study. Molecular analyses on peripheral blood DNA were performed as described in a previous paper: 100 ng of genomic DNA was amplified in the coding regions of both ENG and ACVRL1.

The endoscopic endonasal mini-invasive surgical procedure was performed in the operating theatre under local or general anaesthesia. During surgery, patients were placed in a reverse Trendelenburg position. Careful preparation of the nasal cavities was undertaken: cottons soaked with xylometazoline chlorohydrate 0.1% and oxybuprocaine chlorohydrate 0.01% were placed in the nasal fossa and left in place for 5-10 min to decongest and clean the nasal mucosa. Surgery was performed under endoscopic endonasal control with a 0° rigid 4 mm optic (Karl Storz and Co., Germany) to selectively treat involved mucosa and telangiectasias located along the nasal fossae. After surgery, an antibiotic ointment was placed in the nasal cavities and no nasal packing was necessary. In some patients whose intraoperative bleeding was more severe than expected and not managed elsewhere, we applied selective bipolar cauterisation or an intranasal haemostatic matrix such as Floseal (Baxter, U.S.), Surgiflo (Ethicon, U.S.), Perclot (Cryolife, U.S.), or Haemocer (BioCer GmbH, Germany) as an adjunct to local coagulation control of haemostasis. In the postoperative period, patients were periodically evaluated (every 6–12 months) for epistaxis with medical evaluation and/or follow-up phone call. In cases in which good control of epistaxis was obtained, follow-up was continued as described; in cases of worsening of symptoms, a new surgical procedure was performed. Non-responders (31 patients) were given medical therapy with oral thalidomide and excluded from the present analysis.
**Statistical analysis**

Descriptive statistics were used for demographic characteristics for this sample of patients. The Shapiro-Wilk test was used to test the normal distribution of quantitative variables. When quantitative variables were normally distributed, the results were expressed as mean value and standard deviation (SD), otherwise median and interquartile range (IQR; 25th - 75th percentile) were reported. Qualitative variables were summarised as counts and percentages. The comparisons between gender or patients with one procedure versus patients with more than one procedure were performed with chi-square test for categorical variables and Student's t test (or Kruskall-Wallis test if data are skewed) for continuous variables. All tests were two-sided and p values < 0.05 were considered statistically significant. The data analysis was performed with the STATA statistical package (release 14.0, 2015, Stata Corporation, College Station, Texas, USA).

**Results**

Five hundred and ninety-one HHT patients, diagnosed according to the Curacao criteria, were hospitalised and screened at our department. Of these, 323 patients (54.7%) underwent endoscopic surgery for epistaxis, for a total of 679 procedures. Concerning mutational screening, analyses were completed on 207 patients: 152 patients (73.4%) carried a mutation in \textit{ACVRL1}, while 55 (26.6%) carried a mutation in \textit{ENG}.

The group consisted of 182 men (56.3%) and 141 women (43.6%), with a mean age of 56 years at first surgery (range 5–86 years); interestingly, we found a significant difference between males (mean 53 years, SD 14 years) and females (mean 57 years, SD 15 years) regarding mean age at first surgery (p = 0.0158).

General anaesthesia was used in 524 procedures (77.2%), while local anaesthesia was used in 155 (22.8%). In 175 patients (54.2%), a single surgical procedure was effective to obtain acceptable control of epistaxis (median follow-up in this group was 48 months), while 148 patients (45.8%) required more than one procedure over time, for a total number of 679 procedures (Tab. I). During the follow-up period, 25 patients (7.7%) were lost and 31 patients (9.6%) died. The mean number of surgical procedures in our group was 1.9 in females (SD 1.4), 2.3 in males (SD 1.8) and 2.1 (SD 1.6) overall. No significant difference was noted between \textit{ACVRL1} and \textit{ENG} patients regarding the number of surgical procedures (one surgery vs two or more; p = 0.1389). In patients with more than one surgery, the time between procedures was analysed (Tab. II; Fig. 1). Preoperative epistaxis severity was scored according to Pagella et al. and was moderate or severe in all patients 6. Characteristics of telangiectasias were analysed before the first surgery and this data was available for 160/323 patients (49.5%): according to the description performed by Pagella et al. in 2009 6, the most common pattern was “mixed” (90 patients, 56.3%), followed by the “large” (56 patients, 35%) and “punctate” pattern (14 patients, 8.7%) (Fig. 2). The nasal septum and lateral nasal walls were involved, respectively, in 91.9% and 91.3% of patients, and the nasal floor in 43.7% and nasal valve in 31%. Interestingly, a significant difference (p = 0.033) between males and females was found in the pattern of telangiectasias: the punctate pattern was more common in females (11 vs 3), while mixed and large patterns were more common in males (respectively 51 vs 39 and 33 vs 23).

In our sample of HHT patients treated with endoscopic endonasal mini-invasive technique, APC (Fig. 3) was the

![Figure 1. Plot showing the time between surgical procedures (in days) in HHT patients with more than one surgery. Columns: medians; lines: interquartile ranges. Females represented in white and males in black.](image-url)
instrument of choice (661 procedures, 97.3%), while lasers (diode, thulium) and quantum molecular resonance technology (Figs. 4, 5) were used in 15 procedures (2.2%). Since 2004, intranasal haemostatic matrices such as Floseal (Baxter, U.S.), Surgiflo (Ethicon, U.S.), Perclot (Cryolife, U.S.), or Haemocer (BioCer GmbH, Germany) were introduced in our clinical practice in order to permit adequate management of severe intraoperative bleeding and better visualisation of the intraoperative field. In our experience, we successfully used these devices during surgery in 83 procedures (12.2%).

Septal perforation was observed in 75 patients (23.2%). Among these, 31 patients (41.3%) had a septal perforation before our first surgery, so in these patients the complication was not related to our treatment. On the other hand, 44 patients (58.7%) developed a septal perforation during
our treatment: 18 patients (40.9%) after one procedure, 8 patients (18.2%) after two procedures, 12 patients (27.3%) after three procedures, and 6 patients (13.6%) after four or more procedures. In most of these patients, previous surgery in another hospital was performed, which could explain the relatively high rate of septal perforations in our group.

Discussion

Epistaxis in HHT remains a problem with regards to definitive treatment: current therapies permit control of nosebleeds, but not complete resolution. As stated by Silva et al., first-line treatment of HHT-related nosebleeds includes hygiene of the nasal cavities and avoidance of triggering factors. Different therapies, both medical and surgical, have been proposed in the literature. In our opinion, endoscopic endonasal mini-invasive techniques have several advantages over other techniques, and these are represented by less morbidity, reduced trauma of nasal mucosa, low risk of septal perforation, treatment repeatability, no need for post-surgical nasal packing, the possibility of local anaesthesia and short time of hospitalisation. A review of the literature suggests that the choice of treatment should be based on the severity of nosebleeds: in cases of bleeding of mild entity, a mini-invasive technique can be proposed; on the other hand, in patients with severe epistaxis more invasive surgeries should be chosen (e.g. septodermoplasty, septectomy, closure of the nostrils). However, in 2013, our group demonstrated the efficacy and long-term benefit of a mini-invasive approach (APC) in a group of HHT patients affected by severe epistaxis with a history of blood transfusion. In this paper, we report our 20-year experience in the surgical management of epistaxis in HHT, focusing on endoscopic mini-invasive techniques. The Department of Otorhinolaryngology at I.R.C.C.S. Policlinico San Matteo in Pavia is a reference centre for diagnosis and treatment of
HHT in Italy, and, over the period analysed (1996-2015), 591 patients with a diagnosis of HHT (according to Cura-cao criteria) were referred to our centre. The first concept that should be stressed is that only 54.7% of these patients required surgery for control of epistaxis. This is, to the best of our knowledge, one of the most extensive reported case series on epistaxis surgical treatment in HHT. For the remaining 268 patients, although epistaxis was still present (in fact, it is usually the most common symptom that leads to diagnostic suspicion), quality of life-related to this problem was deemed acceptable and did not require further interventions. Another interesting piece of data is that in 175 patients (54.2%), a single surgical procedure was effective to obtain acceptable control of epistaxis (median follow-up: 48 months), while 148 patients (45.8%) required more than one procedure over time; even if not objective, this is a good outcome measure of patient satisfaction with epistaxis control. In our opinion, in fact, it is essential to take into consideration the personal satisfaction of each patient, given that each treatment used for epistaxis in HHT is symptomatic. According to this management strategy, Sautter and Smith also stated in 2016 that, regardless of treatment modality, HHT-related epistaxis would continue.

The mean number of surgical procedures in our group was 2.1 (1.9 in females and 2.3 in males, no significant difference). No significant difference was reported between ACVRL1 and ENG patients regarding the number of procedures. Repetition of a mini-invasive surgical procedure, given its low risk of short-term and long-term complications (44/323 patients developed a septal perforation during treatment, most underwent previous repeated cauterisations in other hospitals) is possible when nasal bleeding returns and has a negative impact on the quality of life. A “staged” procedure should be performed when telangiectasias are present on opposing sides of the anterior nasal septum in order to avoid septal perforation. Moreover, some authors report that submucosal injections of bevacizumab associated with surgical treatment may increase the risk of perforation.

Regarding the characteristics of nasal telangiectasias, data was available for 160 of 323 patients (49.5%): the most common pattern was mixed (90 patients, 56.3%), followed by the large (56 patients, 35%), and punctate patterns (14 patients, 8.7%). The nasal septum and lateral nasal walls were involved, respectively, in 91.9% and 91.3% of patients, nasal floor in 43.7% and nasal valve in 31%. Wide variability can be found not only in the nasal pattern of telangiectasias, but also in the number of nasal subsites involved, even in patients with the same pattern. Furthermore, we confirmed that different subsites in nasal fossae, and even the same subsite, can present similar or different patterns. We also observed a higher incidence of mixed and large patterns in surgical candidates; this is in accordance with a previous paper in which we observed a significantly higher frequency of nosebleeds in patients presenting the large pattern.

Regarding surgical technique applied, APC was the instrument of choice in most of our patients (661 procedures, 97.3%), while lasers and quantum molecular resonance technology were used only in 2.7% of cases. Indeed, APC is our instrument of choice in the treatment of epistaxis in HHT patients independently of preoperative epistaxis severity, which was moderate or severe before surgery in all patients.

Bergler first described APC treatment of nosebleeds in 1999 on 12 HHT patients. From a technical point of view, APC is based on high-frequency electric energy transmitted through ionised argon gas. The generator is set at 40–60 W and the argon gas flow at 1.2 L/min. Endoscopic endonasal visualisation permits selective treatment in a noncontact mode of telangiectasias located along all the nasal fossae (also in their posterior portion) and in the nasopharynx. Coagulation of tissue is limited to a 1-2 mm depth. This permits low risk of local tissue damage because the temperature is rarely over 100°C. Finally, the applicator tips can be sterilised, and the cost of argon gas is very low.

In the literature, multiple techniques have been described for endoscopic surgical treatment of nasal telangiectasias (Tab. III). Ablative therapies, such as sclerotherapy, electro-surgery and laser treatment, show good outcomes in reducing epistaxis in HHT patients. Sclerotherapy with foamed sodium tetradecyl sulfate has been shown to be effective and safe in four studies, including one randomised clinical trial. The potential side effects are tissue necrosis, cellulitis at the site of injection and, more rarely, anaphylaxis, pulmonary embolus and permanent blindness due to central retinal artery or ophthalmic artery occlusion. Bipolar electrosurgery is preferable to monopolar electrosurgery due to the lower risk of septal perforation, and Ghaheri et al. reported on the efficacy of bipolar cautery for epistaxis treatment in HHT.

Laser photocoagulation is also a viable option, and many different types of laser exist. Lasers with higher tissue penetration that use haemoglobin as a chromophore, such as the Nd:YAG laser, are preferred over lasers with less penetration. The problem with laser use is that not all devices can be used with flexible fibres, and the manoeuvrability inside nasal cavities can be limited. Moreover, increased thermal damage to surrounding tissue is a concern when using lasers.

Multiple uncontrolled series demonstrated that laser treatments are efficient in reducing epistaxis duration, frequency and severity in HHT patients.
### Table III. Surgical techniques for epistaxis treatment in HHT patients.

| Study                        | Study design       | No. of patients | Treatment                      | Outcome of interest                               | Results                                      |
|------------------------------|--------------------|-----------------|--------------------------------|---------------------------------------------------|----------------------------------------------|
| Boyer H. et al., 2011        | Retrospective      | 7               | Sclerotherapy                  | Epistaxis frequency and severity                  | 100% improved                               |
| Morais D. et al., 2012       | Retrospective      | 45              | Sclerotherapy                  | Epistaxis frequency and severity                  | 95% Improved                                |
| Boyer H. et al., 2015        | Randomised         | 17              | Sclerotherapy                  | ESS                                              | Improved                                    |
| Esteban-Casado S. et al.,    | Cross-sectional    | 38              | Sclerotherapy and topical nasal| ESS, VAS, EQ-5D                                   | Improved                                    |
| 2019                         |                    |                 | propranolol                    |                                                   |                                              |
| Ghaferi B. et al., 2006      | Retrospective      | 18              | Bipolar cautery                | Need for recurrent intervention                   | Recurrent intervention in 50% at a mean follow-up of 2.3 years |
| Bergler W. et al., 1999      | Prospective        | 12              | APC                            | Epistaxis frequency and intensity                | Improved                                    |
| Pagella F. et al., 2006      | Prospective        | 36              | APC                            | Reported bleeding                                 | 100% reduction at 6 months                  |
| Pagella F. et al., 2013      | Retrospective      | 26              | APC                            | Epistaxis score                                  | Improved at 12 months                       |
| Jørgensen G. et al., 2011    | Prospective        | 30              | Laser                          | Epistaxis duration                               | Reduced at 1.5 and 6 months                 |
| Kuan E.C. et al., 2017       | Retrospective      | 20              | Laser                          | SNOT-22                                          | Improved at 1.5 months                      |
| Fiorella M.L. et al., 2012   | Retrospective      | 24              | Diode laser                    | Epistaxis frequency and severity                 | Improved                                    |
| Poje G. et al., 2017         | Retrospective      | 17              | Diode laser                    | Epistaxis frequency and severity                 |                                              |
| Papaspyrou G. et al., 2016   | Retrospective      | 38              | Nd:YAG laser +/- APC           | Need for recurrent intervention                   | 18% recurrent intervention at a mean follow-up of 3 years |
| Papaspyrou G. et al., 2017   | Prospective        | 45              | Nd:YAG laser +/- APC           | Need for recurrent intervention                   | 20-33% recurrent intervention at 3-10 years  |
| Joshi H. et al., 2011        | Case series        | 5               | Coblation                      | Epistaxis control                                | 80% of patients                             |
| Mortuaire G. et al., 2013    | Prospective        | 16              | Coblation                      | Epistaxis frequency and duration                 | Reduced frequency at 6 months              |
| Rotenberg B. et al., 2015    | Retrospective      | 37              | Coblation                      | ESS                                             | Improved at 6 months                        |
| Luk L. et al., 2014          | Prospective        | 11              | Coblation vs KTP laser         | ESS                                             | No difference in mean ESS                   |
| Ishibashi T. et al., 2003    | Case report        | 2               | Harmonic Scalpel               | Frequency of epistaxis                           | Improved                                    |
| Ichimura K. et al., 2006     | Retrospective      | 15              | Septodermoplasty               | Patient satisfaction                             | 100% satisfied                              |
| Lesnik G.T. et al., 2007     | Retrospective      | 9               | Septodermoplasty               | Epistaxis frequency, QoL, blood transfusions     | 100% improved QoL, blood transfusion reduced |
| Levine C.G. et al., 2008     | Retrospective      | 106             | Septodermoplasty               | QoL                                             | 62% responded and 86% improved              |
| Harvey R. et al., 2008       | Retrospective      | 33              | Septodermoplasty               | Frequency of KTP laser                           | Frequency of KTP laser treatment decreased (p = 0.012) |
| Rimmer J. et al., 2014       | Prospective        | 7               | Septodermoplasty               | Epistaxis frequency and severity                 | 100% improvement                            |
| Lee J.M. et al., 2019        | Prospective        | 7               | Temporary nasal occlusion with | ESS and clinical assessment of nasal cavity      | No significant ESS improvement (p = 0.179); clinical assessment of nasal cavity improved (p = 0.0088) at 1 month |
| Hosni A.A. et al., 1994      | Case series        | 2               | Nasal closure                  | Epistaxis frequency                              | Improved                                    |
Ishibashi and Takamatsu described the intraoperative application of the harmonic scalpel (an ultrasonic system) in two HHT patients; highlighting the lack of damage to the surrounding mucosa, but long-term results were not reported. Some authors described coblation in the treatment of HHT-related nosebleeds; this method has been shown to be a useful alternative to other surgical devices, improving nosebleed severity.

Septodermoplasty and nasal closure are considered useful in patients whose epistaxis fails to respond sufficiently to ablative therapies. According to several authors, septodermoplasty improves both quality of life and epistaxis severity. Septodermoplasty and nasal closure are considered useful in patients whose epistaxis fails to respond sufficiently to ablative therapies. According to several authors, septodermoplasty improves both quality of life and epistaxis severity.

Possible drawbacks of this study include the lack of objective outcome measures, its retrospective nature and the lack of a control group treated with different procedures (e.g. APC vs other techniques).

Conclusions

Recurrent epistaxis from telangiectasias of the nasal mucosa is a common manifestation present in more than 95% of HHT patients. Current therapies cannot permit a definitive resolution of the symptom, but only temporary control of nosebleeds and improvement in the quality of life. Herein, we describe our 20-year experience in the endoscopic surgical treatment of epistaxis in a large group of 323 HHT patients.

Endoscopic endonasal mini-invasive surgical options, and in particular APC, allowed good control of epistaxis without major complication and nasal packaging. These treatment modalities are repeatable, non-invasive and can be used as a first step even in patients with severe epistaxis. In conclusion, since no treatment could be considered definitive and the natural history of HHT-related nosebleeds is unknown, in our opinion the application of mini-invasive treatment modalities that permit preservation of nasal anatomy and physiology should be the approach of choice.

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Table III.

| Study               | Study design | No. of patients | Treatment       | Outcome of interest | Results                     |
|---------------------|--------------|----------------|-----------------|---------------------|-----------------------------|
| Ichimura K. et al., 2012 | Prospective  | 7              | Nasal closure   | Epistaxis cessation | 57% success                 |
| Richer S. et al., 2012 | Retrospective | 43             | Nasal closure   | Epistaxis cessation | 84% responded and 83% success |
| Lund V. et al., 2017  | Retrospective | 100            | Nasal closure   | Epistaxis cessation | 50% responded and 94% success |
| Andersen J.H. et al., 2020 | Retrospective | 10             | Nasal closure   | GBI                 | Overall GBI score of 38.05 with an average follow-up 66 months |

ESS: Epistaxis Severity Score; VAS: visual analogue scale; EQ-5D: EuroQol-5D scale; APC: Argon plasma coagulation; SNOT-22: Sinonasal outcome test-22; QoL: Quality of life; KTP: Potassium-titanium-phosphate; GBI: Glasgow benefit inventory.
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