

Original Article

Surgical Management of Rhinosinusitis in Onco-Hematological Patients

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Objectives. In onco-hematological diseases, the incidence of paranasal sinuses infection dramatically increase and requires a combination of medical and surgical therapy. Balloon dilatation surgery (DS) is a minimally invasive, tissue preserving procedure. The study evaluates the results of DS for rhinosinusitis in immunocompromised patients.

Methods. A retrospective chart review was conducted in 110 hematologic patients with rhinosinusitis. Twenty-five patients were treated with DS technique and 85 patients with endoscopic sinus surgery (ESS). We considered the type of anesthesia and the extent of intra- and postoperative bleeding. Patients underwent Sino-Nasal Outcome Test (SNOT-20) to evaluate changes in subjective symptoms and global patient assessment (GPA) questionnaire to value patient satisfaction

Results. Local anesthesia was employed in 8 cases of DS and in 15 of ESS. In 50 ESS patients, an anterior nasal packing was placed and in 12 cases a repacking was necessary. In the DS group, nasal packing was required in 8 cases and in 2 cases a repacking was placed (*P*=0.019 and *P*=0.422, respectively). The SNOT-20 change score showed significant improvement of health status in both groups. However the DS group showed a major improvement in 3 voices: need to blow nose, runny nose, and facial pain/pressure. The 3-month follow-up GPA questionnaire showed an higher satisfaction of DS group.

Conclusion. Balloon DS represents a potentially low aggressive treatment and appears to be relatively safe and effective in onco-hematologic patients. All these remarks may lead the surgeon to consider a larger number of candidates for surgical procedure.

Keywords. Rhinosinusitis, Balloon sinuplasty, Oncohematological disease

INTRODUCTION

In onco-hematological diseases, an high incidence of rhinosinusitis is present due to alteration of the immunological asset. Moreover the increasing use of aggressive and intensive cancer chemotherapeutic regimens, immunosuppressive therapy, transplan-

- Received May 3, 2013 Revision October 3, 2013 Accepted October 10, 2013
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tation and radiation therapy increase dramatically the incidence of paranasal sinuses infection [1,2]. Because of their higher risk, compared to the healthy population, to develop serious complications critically ill patients with both hematological malignancies and rhinosinusitis require a timely and appropriate treatment [3,4]. Prognosis is related to various factors, such as the type of infection, the speed of diagnosis and treatment, the complete neutrophil count normalization and the involvement of structures adjacent to the site of infection as the central nervous system, the orbit and the cavernous sinus [5].

Mucositis, induced by chemotherapy treatment, leads to an alteration of mucosal defense mechanism, mucociliary clearance failure, edema, crusting and scars that can alter the sinus ostial

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ventilation [6,7]. Sinus dysventilation can lead to a pathological sinus cycle, the first step of rhinosinusitis.

On the other hand, surgery in this immunocompromised patients is reserved only for critical patients because of the risk of general anesthesia, infections, bleeding due to thrombocytopenia and the overall clinical state lead the surgeon to a difficult decision [8].

In traditional endoscopic sinus surgery (ESS), the combination of enlargement of sinus ostia with throughbiting instruments that incise through highly vascular sinonasal mucosa, with an inefficient haemostatic mechanisms and a lack of physiologic immune reserve, make this kind of surgery in immunocompromised and critically ill patients a challenging endeavor [9].

Dilatation surgery (DS), on the contrary, is a minimally invasive and tissue preserving procedure that relies on sinus dilation by means of inflatable balloon without mucosal cutting or removing tissue [10]. The minimally invasive nature of the technique is proposed to minimize blood loss, to induce less postoperative scarring and to facilitate the management of postoperative patient [11].

The aim of our study is to compare the surgical outcomes between traditional endoscopic and DS for the treatment of sinusitis in immunocompromised patients.

MATERIALS AND METHODS

A retrospective and non-randomized chart review was conducted in 110 patients affected by rhinosinusitis. This study was approved by the Tor Vergata Hospital Institutional Review Board and all participants signed an informed consent agreement. The patients were selected in a group of 380 hematologic patients admitted to our hospital from June 2008 to June 2012. Preoperative assessment included physical examination, sinonasal endoscopy, baseline Sino-Nasal Outcome Test (SNOT-20) calculated as the mean of the total score obtained and all patients were studied by means of computed tomography (CT) as suggested by hematology and oncology guidelines [12]. The Lund-Mackay score (Evidence Level IIb) was used for the CT disease staging. It relies on a score of 0–2 dependent upon the absence, partial or complete cloudiness of each sinus system and of the ostiomeatal complex, deriving a maximum score of 12 per side [13].

The diagnosis of rhinosinusitis was confirmed in 112 cases out of 380. Two of them presented critical general health condition, so both surgical approach was reasonable. The other 110 patients had standard indication for surgery: a strictly defined rhinosinusitis diagnosis per Rhinosinusitis Task Force criteria (symptoms including but not restricted to nasal obstruction, sinus/facial pressure, nasal discharge, and congestion) that was unresponsive to maximal medical management. Maximal medical management included three to six weeks of antibiotic therapy, inhaled and/or systemic corticosteroids, decongestants as appropriate, and saline irrigations [14].

During preoperative time 71% of patients had a platelet count <50,000 (severe thrombocytopenia), 17% 50,000–100,000 platelets (moderate thrombocytopenia), 9% 100,000–150,000 platelets (mild thrombocytopenia) and 3% had a platelet count from 150,000 to 250,000. All patients were leukopenic: 68% of them were severely neutropenic (neutrophil count <500/mmc) and the other 32% were moderate or mild neutropenic.

Patients surgically treated were subdivided in two groups: group 1 included 25 patients treated with DS since balloon was available (2008) and group 2 included 85 patients treated with ESS (Table 1). The choice of the surgical technique was based on the availability of balloon in our clinic.

All patients undergoing surgery fill out pre- and postoperative SNOT-20 surveys to determine baseline symptom status (before treatment) and improvement (if any) after intervention. The SNOT-20 rates the severity of 20 symptoms on a six-point scale (from 0 [no problem] to 5 [problem as bad as it can be]). To assess the impact of treatment, the difference between pre- and postoperative (at 3 months) SNOT-20 scores was calculated to determinate the SNOT-20 change score. At 3 month postoperative follow-up, patients underwent a global patient assessment (GPA) questionnaire to value patient satisfaction. We ask a patient to rate on a scale of +5 (the best outcome possible) to -5 (the worst outcome possible) how they feel overall; also GPA assesses patient satisfaction by asking him, if given the choice, would choose to undergo the same procedure (yes/not sure/no).

Table 1. Different characteristics between the two groups

Characteristic	Dilatation surgery	Endoscopic sinus surgery
No.	25	85
Hematological diagnosis		
Acute leukemia	17	56
Lymphoma/myeloma	8	29
Sinus treated		
Maxillary	12	40
Frontal	8	17
Sphenoid	3	10
Multiple sinuses	2	18
Anesthesia		
Local	8	15
General	17	70
Anterior nasal packing		
Yes	8	50
No	17	25
Repacking		
Yes	2	12
No	23	73
Postoperative nasal debridement		
Yes	3	35
No	22	50
Outcome		
Death	2	10
Alive	23	75

Furthermore a total symptom score was constructed (1 point for each symptom if it was present for at least 6 hours a day and affecting daily life and no points if symptom did not affect daily life or was absent). Success and improvement were defined as a decrease in the total complaint score of 1 or more points at last visit; while total improvement was defined as total resolution of all complaints (i.e., symptom score of 0).

All statistical analyses were performed using SPSS ver. 17.0 (SPSS Inc., Chicago, IL, USA). Continuous data are displayed as mean \pm SD. Statistical significance was accepted when P<0.05. The Levine's test for equality of variances was used to determine statistically significant variances. The two independent samples test with Mann-Whitney U coefficient was used to compare the different results within each group. The χ^2 and the Fisher exact tests were used to test the association between categorical variables.

RESULTS

In 112 of 380 hematologic patients, an involvement of paranasal sinuses was present (29.4%). The infection was most common in patients with acute leukemia group compared to lymphoma or myeloma group (75 vs. 37).

Lund-Mackay scores, based on preoperative sinus CT scans, were comparable for the DS and ESS groups. P-value assessing presurgical Lund-Mackay scores between DS and ESS subjects found no significant difference (P=0.32) (Table 1).

Balloon catheters were used to treat 12 maxillary sinuses, 8 frontal sinuses, 3 sphenoid sinuses and 2 multiple sinuses. Of the 8 frontal sinuses, 3 were treated with hybrid procedure (opening of the posterior wall of the agger nasi or the anterior wall of the

ethmoidal bulla) (Table 1).

In the ESS group, 40 maxillary sinuses, 27 frontal sinuses, 10 sphenoid sinuses, and 8 maxillary-sphenoid sinuses (Table 1) were treated. Local anesthesia was employed in 8 of the 25 patients (32%) treated with DS and in 15 of the 85 patients treated with ESS (17.6%). General anesthesia was necessary in the others 17 patients treated with DS and the 70 patients treated with ESS (P=0.123) (Table 1).

No major adverse events were reported linked to th surgery techniques. In 50 ESS patients group treated an anterior nasal packing was placed at the end of the procedure and in 12 patient with a minor bleeding a new anterior nasal packing was necessary after 4 days. In the DS group nasal packing was required in 8 cases and in 2 cases a repacking was necessary (P=0.019 and P=0.422, respectively) (Table 1). A postoperative nasal debridement was necessary in 25 patients of ESS group and in 3 of DS group (P=0.08).

The t-test analysis revealed improvement in SNOT-20 scores following surgery in both dilatation and ESS group (P<0.001). The preoperative SNOT-20 scores were 58 ± 10.4 in DS patients and 56 ± 17 in patients treated with ESS, showing well-matched group. The mean 3-month postoperative SNOT-20 scores after DS (17.6 ± 11) was less than ESS postoperative SNOT-20 scores (19.6 ± 17.4). The SNOT-20 different score of both procedures showed clinically significant improvement of health status: 39.8 ± 13.2 for DS group and 36.2 ± 19.6 for ESS group. But also an improvement was recorded the DS group in 3 voices of SNOT-20: need to blow nose, runny nose, and facial pain/pressure (Fig. 1).

Table 2 is showed the overall improvement in reports of specific preoperative symptoms at final postsurgical visit as measured at 3 month postsurgical follow-up. Both group had improvement of symptoms during the follow-up period. 87.5% of patients who

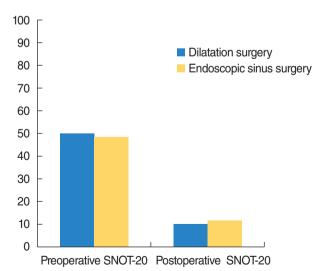


Fig. 1. Changes in Sino-Nasal Outcome Test (SNOT-20) results between 2 groups.

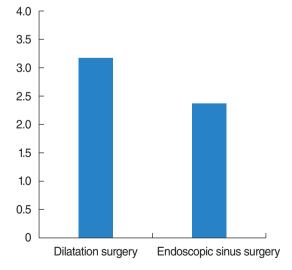


Fig. 2. Global patient assessment outcome between 2 groups.

Table 2. The patients in their group reporting improvement of sinus symptoms at 3 month follow-up

Variable	Dilatation surgery	Endoscopic sinus surgery	P-value
Symptom total	21 (87.5)	49 (61.2)	0.017
Facial pain	13 (54.1)	28 (35.0)	0.094
Sinus congestion	12 (50.0)	27 (33.7)	0.151
Rhinorrhea	14 (58.3)	32 (40.0)	0.114
Headaches	10 (41.6)	36 (45.0)	0.774

Values are presented as number (%).

underwent DS and 61.2% of patients who underwent ESS demonstrated a reduction of symptoms at 3 month postoperative follow-up (P=0.017) (Table 2).

The 3-month follow-up GPA form questionnaire recorded that 87.5% of DS patients responded with "yes" to having the same procedure done again, instead only 47.5% of post-ESS patients said "yes" (P=0.001) (Table 3). Additionally, 4.1% of DS patients responded with "no" and 8.3% were "not sure" when faced with the same question, versus 8.7% and 43.7% of the post-ESS patients, respectively (P=0.462 and P=0.002, respectively) (Table 3, Fig. 2).

DISCUSSION

In the last 20 years, as results of improved imaging and surgical techniques, sinus surgery has become more conservative [15,16]. To date ESS is the optimal surgical procedure for patients affected by sinusitis who not responding to medical management. A success rates between 75% and 95% is typically reported [17]. However major complications still occur in approximately 1.1% of surgery and minor complications, including bleeding, infection, periorbital emphysema, ecchymosis and synechiae, occur in 5.4% of cases [18]. The technical philosophy of ESS includes enlargement of sinus ostia and elimination of the surrounding mucosal contact areas. A constant growth of technical development has been a drive toward more delicate instrumentation that could help the surgeon to respect nasal mucosa. An enlargement of the surgical armamentarium was obtained introducing the DS tool.

This minimally invasive surgical procedure is able to dilate the sinus ostia preserving mucosa and saving the mucociliary transport function [19]. DS is a complementary tool for endoscopic nasal surgery and it is usually employed only in isolated sinusitis. In immunocompromised patients, DS seems to obtain multiple advantages and some remarks can be drawn.

In a critical mucosal condition, the intraoperative blood loss is insignificant. The lower number of anterior nasal packing and repacking confirm the low aggressiveness of the procedure. As a consequence, the development of crusting and synechiae is lower and postsurgical nasal debriment is necessary in a reduced number of patients. This is a very important goal since the general

Table 3. Global patient assessment experience outcome between 2 groups at 3 months

Variable	Dilatation surgery	Endoscopic si- nus surgery	P-value
Yes	21 (87.5)	38 (47.5)	0.001
No	1 (4.1)	7 (8.7)	0.462
Not sure	2 (8.3)	35 (43.7)	0.002
Death	1	5	

Values are presented as number (%).

condition of these patients are often poor and any medical invasive treatment should be avoided.

Local anesthesia was possible in a larger group of DS patients, although the higher complexity surgical procedure were performed in ESS group. All these remarks may lead the surgeon to consider a large number of candidates for this surgical procedure.

A balance between the surgery and the natural evolution of the paranasal infection is mandatory to prevent the critical evolution of the infective disease. A rapid spreading of the infective disease through the surrounding tissue lead to an aggressive paranasal surgery. Probably, the use of DS could avoid the progressive clinical distrectual worsening. DS allows to act timely on a damaged mucosa because of chemotherapy treatment. As a consequence, the restored mucociliary clearance led to a normal sinus cycle re-establishing the physiological sinus ventilation and arresting disease progression in immunocompromised patients.

In according with other authors, we demonstrated the efficacy of balloon sinuplasty as non inferior to ESS in treatment of rhinosinusitis [20]. Our results prove the safety and feasibility of this surgical procedure yet established in previous studies [13,21-25]. However we demonstrate the safety and efficacy of this technique also in onco-hematological patients with sinus dysfunction. In literature there is only one study analyzing the balloon sinuplasty technique in 5 immunocompromised patients [9]. All underwent sinus surgery with no complication, minimal blood loss, and no need for transfusion. To our knowledge only a case of surgical complications in hematological patients is reported: a septal hematoma, spontaneously resolved was descripted after dilatation of the sphenoid natural ostium by means of balloon [26,27].

DS represents a potentially low aggressive treatment and appears to be relatively safe, effective in onco-hematologic patients, minimizing mucosal and structural damage, period of healing and postoperative debridements.

Patients affected by hematological cancers showing an infectious involvement of the paranasal sinuses have a substantially higher risk than the general population to suffer from serious local and systemic complications. Therefore a timely medical and surgical treatment is necessary. DS represents a potentially low aggressive technique and appears to be relatively safe, effective and feasible, as demonstrated by procedural technical success, patient tolerability, and clinical outcomes. Minimizing mucosal

and structural damage, period of healing, postoperative debridements and using local anesthesia it should be considered a valid option in onco-hematologic patients.

CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

REFERENCES

- 1. Mazzone S, Passali FM, Di Girolamo S, Varakliotis T, Sibilia M. Sinusopatie nei pazienti oncoematologici. 99th Italian Congress of torhinolaryngology. 2012 May 23-26; Bari, Italy.
- 2. Rombaux P, Bertrand B, Eloy P. Sinusitis in the immunocompromised host. Acta Otorhinolaryngol Belg. 1997;51(4):305-13.
- 3. Galli J, Di Rienzo L, D'Ecclesia A, Motta S, Di Girolamo S. Difficulties in the clinical, radiological and therapeutic evaluation of the initial stage of mucormycosis of the rhinosinus. Acta Otorhinolaryngol Ital. 2000 Feb;20(1):47-53.
- 4. Castelnuovo P, Gera R, Di Giulio G, Canevari FR, Benazzo M, Emanuelli E, et al. Paranasal sinus mycoses. Acta Otorhinolaryngol Ital. 2000 Feb:20(1):6-15.
- 5. Morrison VA, Pomeroy C. Upper respiratory tract infections in the immunocompromised host. Semin Respir Infect. 1995 Mar;10(1): 37-50
- 6. Vadhan-Raj S. Management of chemotherapy-induced thrombocytopenia: current status of thrombopoietic agents. Semin Hematol. 2009 Jan;46(1 Suppl 2):S26-32.
- 7. Sonis ST. The pathobiology of mucositis. Nat Rev Cancer. 2004 Apr; 4(4):277-84.
- 8. Park AH, Muntz HR, Smith ME, Afify Z, Pysher T, Pavia A. Pediatric invasive fungal rhinosinusitis in immunocompromised children with cancer. Otolaryngol Head Neck Surg. 2005 Sep;133(3):411-6.
- 9. Wittkopf ML, Becker SS, Duncavage JA, Russell PT. Balloon sinuplasty for the surgical management of immunocompromised and critically ill patients with acute rhinosinusitis. Otolaryngol Head Neck Surg. 2009 Apr;140(4):596-8.
- 10. Levine HL, Sertich AP 2nd., Hoisington DR, Weiss RL, Pritikin J, PatiENT Registry Study Group. Multicenter registry of balloon catheter sinusotomy outcomes for 1,036 patients. Ann Otol Rhinol Laryngol. 2008 Apr;117(4):263-70.
- 11. Brown CL, Bolger WE. Safety and feasibility of balloon catheter dilation of paranasal sinus ostia: a preliminary investigation. Ann Otol Rhinol Laryngol. 2006 Apr;115(4):293-9.
- 12. Ruhnke M, Böhme A, Buchheidt D, Donhuijsen K, Einsele H, En-

- zensberger R, et al. Diagnosis of invasive fungal infections in hematology and oncology: guidelines of the Infectious Diseases Working Party (AGIHO) of the German Society of Hematology and Oncology (DGHO). Ann Hematol. 2003 Oct;82 Suppl 2:S141-8.
- 13. Fokkens WJ, Lund VJ, Mullol J, Bachert C, Alobid I, Baroody F, et al. European Position Paper on Rhinosinusitis and Nasal Polyps 2012. Rhinol Suppl. 2012 Mar;(23):1-298.
- 14. Rosenfeld RM, Andes D, Bhattacharyya N, Cheung D, Eisenberg S, Ganiats TG, et al. Clinical practice guideline: adult sinusitis. Otolaryngol Head Neck Surg. 2007 Sep;137(3 Suppl):S1-31.
- 15. Kennedy DW, Adappa ND. Endoscopic maxillary antrostomy: not just a simple procedure. Laryngoscope. 2011 Oct;121(10):2142-5.
- 16. Kutluhan A, Salvız M, Bozdemir K, Kanbak O, Ulu M, Yalcıner G, et al. The effects of uncinectomy and natural ostial dilatation on maxillary sinus ventilation: a clinical experimental study. Eur Arch Otorhinolaryngol. 2011 Apr;268(4):569-73.
- 17. Jakobsen J, Svendstrup F. Functional endoscopic sinus surgery in chronic sinusitis: a series of 237 consecutively operated patients. Acta Otolaryngol Suppl. 2000;543:158-61.
- 18. Rombout J, de Vries N. Complications in sinus surgery and new classification proposal. Am J Rhinol. 2001 Nov-Dec;15(6):363-70.
- 19. Levine H, Rabago D. Balloon sinuplasty: a minimally invasive option for patients with chronic rhinosinusitis. Postgrad Med. 2011 Mar; 123(2):112-8.
- 20. Achar P, Duvvi S, Kumar BN. Endoscopic dilatation sinus surgery (FEDS) versus functional endoscopic sinus surgery (FESS) for treatment of chronic rhinosinusitis: a pilot study. Acta Otorhinolaryngol Ital. 2012 Oct;32(5):314-9.
- 21. Bolger WE, Vaughan WC. Catheter-based dilation of the sinus ostia: initial safety and feasibility analysis in a cadaver model. Am J Rhinol. 2006 May-Jun;20(3):290-4.
- 22. Bolger WE, Brown CL, Church CA, Goldberg AN, Karanfilov B, Kuhn FA, et al. Safety and outcomes of balloon catheter sinusotomy: a multicenter 24-week analysis in 115 patients. Otolaryngol Head Neck Surg. 2007 Jul;137(1):10-20.
- 23. Thottam PJ, Haupert M, Saraiya S, Dworkin J, Sirigiri R, Belenky WM. Functional endoscopic sinus surgery (FESS) alone versus balloon catheter sinuplasty (BCS) and ethmoidectomy: a comparative outcome analysis in pediatric chronic rhinosinusitis. Int J Pediatr Otorhinolaryngol. 2012 Sep;76(9):1355-60.
- 24. Hopkins C, Noon E, Bray D, Roberts D. Balloon sinuplasty: our first year. J Laryngol Otol. 2011 Jan;125(1):43-52.
- 25. Friedman M, Schalch P, Lin HC, Mazloom N, Neidich M, Joseph NJ. Functional endoscopic dilatation of the sinuses: patient satisfaction, postoperative pain, and cost. Am J Rhinol. 2008 Mar-Apr;22(2):
- 26. Alexander AA, Shonka DC Jr., Payne SC. Septal hematoma after balloon dilatation of the sphenoid. Otolaryngol Head Neck Surg. 2009 Sep;141(3):424-5.
- 27. Kennedy DW. Sinus disease: guide to first-line management. Darien, CT: Health Communications; 1994. p. 10-5.