Introduction

Headache may result when pressure exerted on the nasal mucosa in the contact areas by anatomical variations, polyps, or mucosal swelling initiates the release of substance P, which triggers pain impulses in afferent C fibers [1]. With the rapid development of medical imaging and endoscopic techniques, abnormal nasal anatomy has been found to have multiple manifestations, including agger nasi cells, Haller cells, pneumatization of the middle turbinate, middle turbinate abnormal curve, uncinate process abnormalities (hyperplasia, deviation, and pneumatization), nasal septum deviation, ethmoid bulla and maxillary sinus hypoplasia [2]. Contact points refer to intranasal contact between opposing mucosal surfaces. In 1980, in their experience with middle turbinectomy, Morgenstein and Krieger [3] discovered that deformity of turbinate anatomy could cause headache and were the first investigators to propose the concept of contact point headache.

The concept of contact point headache has been widely cited by the Society of Otolaryngology-Head and Neck Surgery and has been applied in clinical practice [4]. Mucosal contact headache was recently added as secondary headache disorder in the second edition of the International Classification of Headache Disorders (ICHD-2),4 where it is described as being characterized by intermittent pain localized in the periorbital and medial canthal or temporozygomatic regions, associated with evidence of mucosal contact points by nasal endoscopy or Computed Tomography (CT) imaging. Research has shown that different anatomical variations, especially septal spurs and abnormal turbinates, can be the cause of intranasal mucosal contacts [5]. Pneumatized superior turbinate process or Haller’s cell has been reported to cause migraine-type headaches [2].
scans of the nose and paranasal sinuses of a child with recurrent attacks of unilateral headache showed a spur in contact with the left inferior turbinate [6] however, whether the type of nasal anatomic abnormalities associates with incidence of headache remain undefined. Surgical treatment has been recommended for the patients with refractory primary headaches associated with mucosal contact point [7,8]. However, surgery for removal of nasal contact points in patients with facial pain or headache may not be unnecessary [9,10].

The purpose of this study was: (1) to investigate the differences in incidence of mucosal contact points within the nasal cavity in subjects with and without headache; (2) to investigate whether surgical treatment is necessary for the patients who fulfill most part of the diagnostic criteria for mucosal contact point headache.

Materials and Methods

Participants

This study was approved by the Institutional Review Board of Beijing Millennium Monument Hospital affiliated with Capital Medical University. In this observational study, 185 healthy volunteers who had undergone physical examination and were patients at the Ear, Nose and Throat clinic or the Department of Neurology between September 2005 and July 2010 were recruited by convenience sampling. Of these, a headache group included 78 patients with a single symptom of refractory headache for more than 2 years. A non-headache group included 107 patients who had visited the Ear, Nose and Throat clinic but without headache. Patients with nasal-borne diseases, including sinusitis, nasal polyps, nasal cavity and paranasal sinus tumors, and medical conditions that could induce headache (e.g., hypertension) were excluded. All participants provided signed informed consent and underwent sinus high-resolution CT scan.

High-resolution CT (HRCT)

The Siemens Sensation 16-slice spiral CT was used (Siemens, Erlangen, Germany), and the scan baseline was parallel to the orbitomeatal line. The scanning parameters included collimation: 0.75mm x 16, pitch 0.55. Three-dimensional reconstruction of images was created. Slice thickness was 0.75mm, slice interval was 0.5mm, the matrix was 512 × 512, the window level was 400HU, and window width was 1800HU, which together ensured the continuity of image information. The same researcher reviewed the images for more than three times at different times (test-retest) to ensure consistency of results of the nasal cavity and sinus anatomical structures. The incidence of various nasal anatomical abnormalities was calculated, and the difference in the incidence of mucous contact points was compared between the headache group and non-headache group.

Diagnostic criteria of mucosal contact point headache

In our study, the diagnosis of mucosal contact point headache referred to the diagnostic criteria of “Mucosal contact point headache” proposed by Headache Classification Committee of the International Headache Society [4].

A. Intermittent pain localized to the periorbital or medial canthal or temporozygomatic regions fulfilling criteria C and D.

B. Clinical, nasal endoscopic and/or CT imaging evidence of mucosal contact points without acute sinusitis.

C. Evidence that the pain can be attributed to mucosal contact based on at least one of the following:

1. Pain corresponds to gravitational variations in mucosal congestion as the patient moves between upright and recumbent positions.

2. Abolition of pain within 5 minutes after diagnostic topical application of local anesthesia to the middle turbinate using placebo or other controls.

3. Pain resolves within 7 days, and does not recur, after surgical removal of mucosal contact points.

Intranasal endoscopic surgery

During the same study period, a total of 25 additional patients (other than 185 healthy volunteers) from the Ear, Nose and Throat clinic who had refractory headache symptoms and were diagnosed as nasal mucosal contact by high-resolution CT without acute sinusitis underwent intranasal endoscopic correction of nasal anatomic abnormalities. Those patients were originally diagnosed as having migraine (9 patients), tension headache (3 patients) and vascular headache (1 patient). However, before surgery, those patients also fulfilled the diagnostic criteria A to C of “Mucosal contact point headache,” including abolition of pain with local anesthesia within 5 min after diagnostic topical application of local anesthesia using placebo or other controls [4]. Among these patients, 17 were males and 8 were females. Mean age was 31.9 years. The course of disease ranged from 2 to 20 years. All participants provided signed informed consent and underwent both nasal endoscopy and sinus high-resolution CT scan examination to confirm the diagnosis. Pre- and post-operative CT scan images of the sinuses were collected and the pain scores (Visual Analogue Scale, VAS) were recorded pre- and post-operatively as previously described [11].

Surgical procedure

All 25 patients underwent functional nasal endoscopic surgery under general anesthesia. The basic principle of surgical treatment was that it should be minimally invasive, that is, the objective was to remove the mucosal contact point while preserving normal anatomical structures of the nasal cavity. Septoplasty was performed for patients with nasal septum deviation, and resection of the spinous process was done for patients with contact between the spinous process of the nasal septum and the lateral wall of the nasal cavity. For patients with nasal septum deviation combined with abnormal anatomy of the middle turbinate, septoplasty was performed first and then turbinoplasty was carried out. Resection of the superior turbinates or ethmoidectomy was performed for patients with contact between the nasal septum and the olfactory cleft. External relocation and radiofrequency ablation of the inferior turbinate were carried out for patients with inferior turbinate hypertrophy.

Statistical analysis

Comparability between the patients with and without headache was tested using Chi-square test for categorical variables and data were represented by numbers (n) and percentage (%). Paired t test was used to determine differences in Visual Analogue Scale (VAS) pre- and post-operatively and data were represented as mean ± Standard Deviation (SD). All statistical assessments are two sided and
evaluated at the 0.05 level of significance. All statistical analyses were performed using SPSS 15.0 statistics software (SPSS Inc, Chicago, IL, USA).

Results
The incidence of nasal anatomic abnormality among all subjects is shown in Table 1. The most common nasal anatomic abnormality was the nasal septum deviation (41.1%) (Figure 1), followed by pneumatization of middle turbinate (32.4%) (Figure 2). Some patients showed more than one abnormality and were included as two or more cases in Table 1. For example, one patient had hypertrophy of bilateral inferior turbinate in contact with the nasal septum, and pneumatization of the bilateral superior turbinate in contact with the nasal septum (Figure 3). Anatomical abnormalities were either unilateral (Figure 4) or bilateral (Figure 2).

Table 2 represents comparisons of abnormal nasal mucosa contact between patients with headache and without headache, including 67 patients in the headache group who had abnormal nasal mucosa contact points between the headache and non-headache groups.

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Table 1: Incidence of nasal anatomic abnormality determined by sinus high-resolution CT.

<table>
<thead>
<tr>
<th>Nasal anatomic abnormality</th>
<th>Number of cases showing abnormality</th>
<th>Number of sites showing abnormality</th>
<th>Incidence</th>
</tr>
</thead>
<tbody>
<tr>
<td>Deviation of nasal septum (NS)</td>
<td>76</td>
<td>76</td>
<td>41.1%</td>
</tr>
<tr>
<td>Pneumatization of middle turbinate</td>
<td>60</td>
<td>83</td>
<td>32.4%</td>
</tr>
<tr>
<td>Pneumatization of superior turbinate</td>
<td>40</td>
<td>71</td>
<td>21.6%</td>
</tr>
<tr>
<td>Abnormal curve of middle turbinate (MT)</td>
<td>37</td>
<td>65</td>
<td>20.0%</td>
</tr>
<tr>
<td>Pneumatization of nasal septum</td>
<td>30</td>
<td>30</td>
<td>16.2%</td>
</tr>
<tr>
<td>Hypertrophy of inferior turbinate</td>
<td>17</td>
<td>30</td>
<td>9.2%</td>
</tr>
<tr>
<td>Haller's cell</td>
<td>10</td>
<td>12</td>
<td>5.4%</td>
</tr>
<tr>
<td>Pneumatization of uncinate process (UP)</td>
<td>7</td>
<td>10</td>
<td>3.8%</td>
</tr>
<tr>
<td>Hypertrophy of middle turbinate</td>
<td>5</td>
<td>8</td>
<td>2.7%</td>
</tr>
<tr>
<td>Node of nasal septum</td>
<td>3</td>
<td>3</td>
<td>1.6%</td>
</tr>
<tr>
<td>Hypertrophy of uncinate process</td>
<td>3</td>
<td>5</td>
<td>1.6%</td>
</tr>
<tr>
<td>Pneumatization of Concha nasalis suprema</td>
<td>2</td>
<td>4</td>
<td>1.1%</td>
</tr>
<tr>
<td>Abnormal curve of superior turbinate</td>
<td>2</td>
<td>4</td>
<td>1.08%</td>
</tr>
<tr>
<td>Nasal septum dislocation</td>
<td>1</td>
<td>1</td>
<td>0.5%</td>
</tr>
<tr>
<td>Medially bent uncinate process</td>
<td>1</td>
<td>2</td>
<td>0.5%</td>
</tr>
<tr>
<td>Hypertrophy of agger nasi</td>
<td>1</td>
<td>2</td>
<td>0.5%</td>
</tr>
</tbody>
</table>

\(^a\) Number of patients showing anatomical abnormalities. The same patient showing more than one abnormality was considered as two or more cases.  
\(^b\) Number of sites showing anatomical abnormalities. Some of the cases showing bilateral abnormality were considered as two sites.  
\(^c\) Incidence was the number of cases divided by total number of patients.

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Table 2: Comparisons of abnormal nasal mucosa contact between patients with headache and without headache.

<table>
<thead>
<tr>
<th></th>
<th>Headache (n=78)</th>
<th>Non-headache (n=107)</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Total number of patients showing abnormal nasal mucosa contact (including bony structure and soft structure)</td>
<td>67 (85.9%)</td>
<td>86 (80.4%)</td>
<td>0.327</td>
</tr>
<tr>
<td>Bony structure contact</td>
<td>40 (51.3%)</td>
<td>58 (54.2%)</td>
<td>0.694</td>
</tr>
<tr>
<td>Left olfactory fissure contact</td>
<td>36 (46.2%)</td>
<td>56 (52.3%)</td>
<td>0.406</td>
</tr>
<tr>
<td>Anatomically abnormal middle turbinate contacting with surrounding structure</td>
<td>11 (14.1%)</td>
<td>9 (8.4%)</td>
<td>0.218</td>
</tr>
<tr>
<td>Anatomically abnormal uncinate process and ethmoid bulla contacting with surrounding structure</td>
<td>4 (5.1%)</td>
<td>1 (0.9%)</td>
<td>0.201</td>
</tr>
<tr>
<td>Deviation of nasal septum contacting with lateral nasal wall</td>
<td>43 (55.1%)</td>
<td>43 (40.2%)</td>
<td>0.044*</td>
</tr>
<tr>
<td>Spur pricks of nasal septum contacting with lateral nasal wall</td>
<td>9 (11.5%)</td>
<td>10 (9.3%)</td>
<td>0.628</td>
</tr>
</tbody>
</table>

\(^*\) Indicates a significant difference between the 2 group, P<0.05.
mucosal contact, both bony structure and soft structure contact; and 11 patients (14.1%) who showed abnormal middle turbinate in contact with surrounding nasal mucosa. A total of 86 (80.4%) subjects in the non-headache group showed abnormal nasal mucosa contact after CT scan examination. Significant differences were found in the incidence (%) of nasal septal deviation with a contact point on the lateral nasal wall between patients with and without headache (55.1% vs. 40.2%, P=0.044) (Table 2). No significant differences were shown between headache and non-headache groups in the other abnormalities.

Among 25 patients receiving surgical treatment, 10 cases had spur pricks of nasal septum associated middle turbinate and inferior turbinate contact with surrounding structure, 6 cases had deviation of nasal septum associated middle turbinate and inferior turbinate contact. Other abnormalities included olfactory fissure contact (4 cases), pneumatization of middle turbinate (4 cases), and abnormal curve of middle turbinate (3 cases) and hypertrophy of turbinate (3 cases). The same patient showing more than one abnormality was considered as two or more cases. Figure 5 showed CT image before and after correction of intranasal anatomic abnormalities. After the operation, pain scale scores (VAS) decreased significantly from 60.60±20.02 (pre-operation) to 26.20±27.92 (post-operation) (P<0.001) (Figure 6). Eleven patients (44%) recovered completely from headache symptoms (VAS score 0) 7 days after the surgery and did not recur after the surgery, suggesting that those patients also fulfilled the diagnostic criteria D of "Mucosal contact point headache" proposed by Headache Classification Committee of the International Headache Society [4]. Eight patients (32%) showed partial recovery, and 6 patients (24%) had the same intensity of pain. Among those 6 patients, 2 patients had complete headache relief (VAS score 0) during the first year after surgery but the headache recurred during the follow-up period (12-18 months post-operative).
Discussion

We investigated whether nasal mucosal contact points caused by nasal anatomic abnormalities could be a cause of refractory headache. We observed mucosal contact points in 85.9% of patients with refractory headache but contact points were also found in 80.4% of subjects without headache. The most common nasal anatomical abnormality was nasal septum deviation, and a significant difference was found in the incidence of deviated septum and contact points on the lateral nasal wall between those with headache and those without, suggesting an association between these conditions and headache. However, the incidence of other types of mucosal contact points was not significantly different between the two groups, suggesting that contact points at the spinous process over the septum, in the olfactory cleft area and those caused by anatomic abnormalities of the middle turbinate, uncinate process, and ethmoid bulla might not be associated with headache.

The average post-operative pain scale scores of 25 patients with headache and mucosal contact points decreased significantly after corrective surgery, suggesting that mucosal contact points may be associated with headache. Only eleven patients (44%) had recovered completely from headache symptoms at 7 days after the surgery and headache had not recurred after the surgery, suggesting that only those patients had fulfilled the diagnostic criteria A to D of "Mucosal contact point headache" proposed by the Headache Classification Committee of the International Headache Society.[4] There were still 24% (6/25) of patients who had fulfilled the diagnostic criteria A to C but failed to recover from headache. In addition, we must consider that 80% of subjects showing abnormal nasal mucosa contact on CT scan examination did not have headache symptoms, and approximately 14% of subjects with headache did not have abnormal nasal mucosa contact, suggesting that headache had multiple causes. In fact, those 25 patients who had received surgical treatment in our study also suffered from migraine, tension headache and vascular headache. Thus, the etiology and pathology behind those types of headache might be exceptionally diverse. Taken together, our findings suggest that mucosal contact points do not result in refractory headache in most cases, and mucosal contact point headaches may not be accurately diagnosed before surgery.

Contact points are acknowledged as one of the causes of secondary headache or an aggravating factor for primary headache.[4] and are also reported to trigger primary headache (migraine, cluster, tension-type) [7]. A previous study found that 74% (73/99) of subjects with various primary headaches had underlying rhinologic abnormalities [12], suggesting that subjects with primary headache might have to be examined by both neurologists and otolaryngologists. However, our study observed 76% of recovery or partial recovery from headache after the surgery, which was consistent with previous reports. Relief of mucosal contact point headache was reported in 90% of patients after endonasal surgery, with complete relief in 43% and improvement in 47% [2]. In the report of Clerico et al., 79% of patients with refractory primary headaches responded to endoscopic sinus surgery with alleviation of pain severity or headache frequency [7]. Similarly, based on comparisons of pre- and post-operative pain questionnaires, 83% of patients in another study had recovered from headaches and 8% had significant relief post-operatively [8]. Removal of contact points in patients with rhinogenic mucosal contact point headache was effective only in carefully selected patients [9]. Surgery for removal of nasal contact points in patients with facial pain is usually unnecessary since prevalence is the same in patients with or without facial pain, indicating that a more central process is responsible [10]. Thus, for some patients with refractory primary headaches, mucosal contact points may not be the only or major cause of the headache, and surgical correction of nasal anatomic abnormalities may not be the proper treatment.

Results of the present study revealed that a greater percentage of patients (55.1%) had refractory headache than subjects without headache (40.2%) had nasal septum deviation and a contact point on the lateral nasal wall. Nasal septum deviation accompanied by nasal ventilation disorders, in which the sinuses may be under negative pressure, could result in headache [13]. Currently there is no direct evidence that substance P is produced in response to mucosal contact points. However, Blumenthal suggested that mucous contact points promoted release of substance P by local sensory nerve endings, which may act on the nasal mucosa, causing headache [14]. In migraine patients, mucosal contact points may provide additional input, decreasing the threshold for migraine attacks [15]. Thus, a contact point on the lateral nasal wall may stimulate the local sensory nerve or nasal mucosal receptors, which release substance P both centrally and peripherally and can cause headache.

Contact point headache patients may seek medical help at the Department of Neurology, where they may be diagnosed with migraine, cluster headache and tension headache. Although they may undergo long-term pharmacologic treatment, treatment effects may be poor. In the present study, among 78 patients with refractory headache, many had taken pain medicine for more than two years without alleviating their headache symptoms. For patients who fail to respond to conservative treatment, sinus CT scans should be a routine examination for patients with headache, even in the absence of sinus conditions [1]. In addition, Welge-Leusen, et al. [16] suggests that these patients should be checked at the Department of Otolaryngology-Head and Neck Surgery in addition to the Department of Neurology. Careful nasal endoscopy is essential to determine whether mucosal contact points are present. The pathogenesis of headache is complex and diverse and mucosal contact points are not the only cause of refractory headache.

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Based on our results, we suggest that a more strict and scientific standard should be developed for selecting contact point headache surgery cases and long-term follow-up should be carried out to assess the short-, mid- and long-term recovery rates, efficacy rates and recurrence rates of surgical resection of mucosal contact points for treatment of headache. Further studies are necessary to demonstrate the significance and necessity of surgical treatment.

In conclusion, nasal septum deviation with a contact point between the deviated nasal septum and the lateral nasal wall is not necessarily associated with headache. Patients who have fulfilled the diagnostic criteria A to C of “Mucosal contact point headache” proposed by the Headache Classification Committee of the International Headache Society may still fail to recover from headache after corrective surgery.

Acknowledgement

Thank my mentor Yin Jinshu for giving me guidance in the course of my research, Dr. Peng Hong for surgeries support, thanks all members of our team, there would be no such research without you, thank you.

References