

Change in olfactory function after septoplasty. A systematic review and meta-analysis*

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Abstract

Background: Septoplasty is one of the most frequently performed surgeries. However, there remains a question as to the effect of such intervention on the sense of smell. This study aims to examine the available evidence regarding the effect of septoplasty on the sense of smell.

Methods: A database search was performed using PubMed, ScienceDirect, Google Scholar and The Cochrane Library databases from January 1990 to February 2020. Search terms included smell, olfaction, odor, septum, septoplasty, and septorhinoplasty. A meta-analysis was performed with 12 studies that provided sufficient data on change in olfaction.

Results: 14 studies met the inclusion criteria, and 2 additional studies were included manually; comprising a total of 996 patients and 25 controls. Significant improvement in olfactory test scores was observed in all tests. Pre- and postoperative differences in means were 0.63 for BSIT, 0.80 for CCCRC test, 1.16 for odor threshold, 1.43 for odor discrimination, and 1.18 for odor identification.

Conclusions: Septoplasty seems to improve olfactory function. However, the outcome of this intervention is discrete and not equal for all patients, so further randomized trials are needed to confirm current findings.

Key words: smell, olfaction, nasal septum, septoplasty, Sniffin' Sticks, CCCRC, BSIT

Introduction

The sense of smell is responsible for perceiving and processing aromatic compounds. Inhaled particles pass through the nasal cavity to the neurons of the olfactory epithelium, which then synapse in the olfactory bulb. The signals are delivered to multiple locations in the brain where they are interpreted as recognizable odors^(1,2). Therefore, it is assumable to think that anatomical alterations of structures such as the nasal septum or the nasal turbinates can alter this airflow and compromise olfaction⁽³⁻⁶⁾. Nasal obstruction is one of the most common symptoms in sinonasal disease and can stem from various conditions such as rhinitis, turbinate hypertrophy, adenoid hypertrophy, or nasal masses⁽³⁾. However, nasal septum deviation is one of the most common, present in three-quarters of patients who consult for symptoms of nasal obstruction⁽⁷⁾ mostly in secondary and

tertiary centers. Likewise, olfactory dysfunction (OD) due to decreased nasal airflow is reported to be a common complaint among patients with nasal septum deviation^(4,8).

OD might have a negative impact on many aspects of daily life, including food appreciation, personal hygiene, social communication, and detection of environmental risks such as spoiled food or smoke⁽⁵⁾. Therefore, OD due to septal deviation can substantially affect the quality of life of patients. Septoplasty is the standard treatment for nasal septum deviation because it generally results in improved nasal airflow and resolution of nasal obstruction symptoms; however, it is not without side effects⁽⁹⁾. To date, there are multiple studies evaluating the effects of septal surgery on olfactory function and nasal symptoms with conflicting results^(2,4-6,10-21).

There are many tests designed for assessing olfaction objec-

tively. Each has a different methodology and scoring system with which the subjects are classified as anosmic, hyposmic, or normosmic^(22–24).

The purpose of this systematic review and meta-analysis is to examine the available evidence and assess the change in sense of smell of patients with septal deviation who undergo septoplasty.

Materials and Methods

The review protocol was based on the Preferred Reporting Items for Systematic Reviews and Meta-analyses guidelines.

Search strategy

A database search was performed using PubMed, Science Direct, Google Scholar, and the Cochrane Library encompassing records from January 1990 to February 2020. The search terms and details are shown in (Supplements, Table 1). Bibliographies of included studies were cross-referenced to manually identify additional articles.

Selection of studies

Inclusion criteria were research and review articles relating to: 1) any septoplasty technique with or without turbinate resection for nasal obstruction due to septal deviation; 2) assessment of olfaction before and after surgery without restrictions on follow-up time. Exclusion criteria were: 1) surgery for reasons other than nasal obstruction due to deviated septum (i.e. cosmetic, sinusitis, rhinitis, polyps, concha bullosa...); 2) lateralized assessment of olfaction (each nostril scored separately); 3) no available translation into English or Spanish language. Communications, case reports and case series, book chapters, encyclopedia entries, conference abstracts, letters and other correspondence, discussions, and scientific posters were excluded.

Outcomes

The main outcome was the mean change in olfactory test score after septoplasty. Other outcomes were change in percentage of patients classified as anosmic, hyposmic, and normosmic before and after surgery.

Assessment of risk of systematic bias

The methodological quality of the included studies was examined using the NIH Guidance for Assessing the Quality of Before-After (Pre-Post) Studies with No Control Group⁽²⁵⁾, which rates 12 domains as “yes”, “no”, or “not applicable”. The overall quality was valued as good, fair, or poor.

Data extraction

The following information was extracted from each included study and sorted in a Microsoft® Excel v16.0 spreadsheet: author

name, publication year, sample size, surgical technique/s performed, used test or questionnaire, mean follow-up time/s, pre- and post-op values, and reported results of the intervention. Characteristics of included studies are presented in Table 1. For studies with more than one branch, only the branches that met the inclusion criteria were included.

Summary of data

A qualitative synthesis was performed for the results from olfactory tests.

Statistical analysis

At this point, the authors of the included studies were contacted to obtain the correlation coefficient of the pre- and postoperative scores. For the rest of studies, the correlation coefficient used was the average of the reported coefficients (0.6543). To ensure that the results were robust enough, sensitivity analyses were conducted setting the correlation coefficient at 0.8 and 0.5. STATA® 16.0 (StataCorp., College Station, TX, USA) software was used for the meta-analysis. The difference in means in olfactory function was calculated for 2 studies reporting BSIT scores^(12,18), 4 reporting CCCRC scores^(6,13,15,20), 4 reporting all three subsets of the Sniffin' Sticks test^(4,5,14,21) and 2 reporting SST OI scores^(10,11). For clearer interpretation, the meta-analysis was conducted on a raw difference in means instead of a standardized one. The studies by Kokubo et al.⁽²¹⁾, Miyake et al.⁽¹⁶⁾, and Tutar et al.⁽²⁾ were excluded because no numerical values were reported for the olfactory test results. The study by Randhawa et al.⁽¹⁷⁾ was excluded because standard deviations were not reported. For studies with multiple follow up-times, the results from the longest follow-up time were used. For studies with more than one study branch, each branch was considered as a separate study. The test for heterogeneity was conducted using the I² statistic describing the percentage of variation across studies originating from heterogeneity rather than from chance. To calculate the effect sizes, a random effects model was used. However, because each test measures olfaction differently they can't be pooled together, so no overall effect size can be calculated. Publication bias was evaluated by funnel plot and Egger's regression test. The Duval and Tweedie trim-and-fill method was used to correct the effect size to account for potentially unpublished reports. Results were reported as mean, 95% CI, and P values.

Results

Six hundred and seventy records were identified, with 81 duplicates removed. 589 abstracts were screened, and 25 full-text articles reviewed, with 14 studies meeting the criteria for inclusion. Two additional articles were found after cross-referencing the bibliography of included studies. Figure 1 shows the flow diagram of the report selection.

Table 1. Characteristics of included studies.

Study	n	Type of surgery	Olfactory test	FU	Pre-op	Post-op
Damm et al. ⁽⁴⁾ , 2003	30	SP, IT	SST (OT, OD, OI)	4 m	7% were anosmic; 60% hyposmic and 33% normosmic	0% were anosmic; 20% hyposmic and 80% normosmic. Significant improvement of OT, OD, OI scores, albeit modest in OT
Pade et al. ⁽¹⁰⁾ , 2008	150	SP	SST OI	4 m	Mean identification score of 12.31 (2.29)	Mean identification score of 12.81 (2.02). Improvement in 13%, no change in 81%, and decreased function in 7%
Schriever et al. ⁽¹¹⁾ , 2013	44	SP, IT	SST OI	12 m	Mean identification score of 12.12 (3.7)	Mean identification score of 12.6 (2.52). No significant change in identification score
Dengiz et al. ⁽¹²⁾ , 2015	53	SRP	BSIT	4 w, 12 w	Mean preoperative score of 10.15 (1.30)	Change in mean BSIT score was not significant at 4 w, but became significant at 12 w postoperatively
Berkiten et al. ⁽¹³⁾ , 2016	50	SP	CCCRC	6 w	Mean score of 3.81 (0.89)	Statistically significant improvement
Dalgic et al. ⁽¹⁴⁾ , 2016	21 18	SP (ts) SP (m)	SST (OT, OD, OI)	1 w, 3 m	Mean composite score of hyposmia (29.6 group A (ts); 29.5 group B (m))	OT, OD and OI scores increased significantly at 3 m
Kilicaslan et al. ⁽¹⁵⁾ , 2016	37	SP	CCCRC	1 w, 6 w, 6 m, 1 y	2.7% had moderate hyposmia, 10.8% mild hyposmia, and 86.5% normosmia	Total olfaction worsened at 1 w, was the same as pre-op after 6 w, and improved after 6 m and 1 y. By 6 m all patients had become normosmic.
Miyake et al. ⁽¹⁶⁾ , 2016	110	SP, SRP	CCCRC	1 m, ≥3 m	Normosmia in 68.1% of patients, mild hyposmia in 6.4%, moderate hyposmia in 8.2%, severe hyposmia in 10%, anosmia in 7.3%	39.3% of patients improved, 46.4% remained the same, and 14.2% worsened 3 or more months after surgery
Randhawa et al. ⁽¹⁷⁾ , 2016	43	SRP	SST (OD)	12 w	NR	Significant change in SST score. 58% improved, 35% remained the same, 7% worsened
Haytoglu et al. ⁽¹⁸⁾ , 2017	41 (s) 75 (ns)	SP	BSIT	1 m, 3 m	Mean preoperative scores of 8.8 (1.2) in smokers (s) and 8.6 (1.2) in non-smokers (ns). No differences between smoking habit	BSIT scores worsened at 1 m, but improved above baseline at 3 m in both smokers and non-smokers
Turk et al. ⁽¹⁹⁾ , 2017	30	SP	SST (OT, OD, OI)	6 w	33.3% of patients were normosmic, 60.0% hyposmic, and 6.7% anosmic	Significant improvement in SST scores. 63.3% of patients were normosmic, 36.7% hyposmic, and none anosmic 6 w after surgery.
Aydogdu et al. ⁽⁶⁾ , 2019	25 14	SP (c) SP (e)	CCCRC	8 w	Mean preoperative scores of 3.01 (0.57) and 2.92 (0.49) for conventional and extracorporeal septoplasties, respectively	Significant improvement. No differences between surgical techniques
Elbistanli et al. ⁽²⁰⁾ , 2019	20	SP	CCCRC	1 m, 4 m	The mean score was 6.05 (0.9)	Worsening of olfaction at 1 m and recovering of baseline olfaction at 4 m.
Kokubo et al. ⁽²¹⁾ , 2019	34	SRP	UPSIT	4 w 12 w	79.4% of patients had normosmia, 14.7% mild hyposmia, and 5.9% moderate hyposmia	76.5% of patients had normosmia, 20.6% mild hyposmia, and 2.9% moderate hyposmia. No change in UPSIT score after 4 w nor 12 w postoperatively
Valsamidis et al. ⁽⁵⁾ , 2019	60/25 controls	SP	SST (OT, OD, OI)	6 m	31.66% were normosmic and 68.33% had olfactory deficits. Significantly lower scores than controls in all SST sub-tests	Significantly improved olfactory function, but still worse than controls
Tutar et al. ⁽²⁾ , 2020	141	SP	SST (OT, OD, OI)	1 w, 6 w, 6 m	1.4% were anosmic, 13.5% hyposmic, and 85.1% normosmic	0.7% of patients had anosmia at 1 w and 0% at 6 w and 6 m. 17.7% were hyposmic at 1 w, 6.4% at 6 w, and 2.8% at 6 m. 81.6% had normosmia at 1 w, 93.6% at 6 w, and 97.2% at 6 m.

n: number of patients, FU: follow-up, m: months, w: weeks, y: years. SP: Septoplasty, SRP: Septorhinoplasty, IT: surgery of the Inferior Turbinate. SST: Sniffin' Sticks Test, OT: Olfactory Threshold, OI: Olfactory Identification, Olfactory Discrimination, CCCRC: Connecticut Chemosensory Clinical Research Center test, BSIT: Brief Smell Identification Test, UPSIT: University of Pennsylvania Smell Identification Test. ts: transseptal sutures, m: merocele packing, c: classical septoplasty, e: extracorporeal septoplasty. Unless specified, values reported as mean (SD). NR: Not Reported.

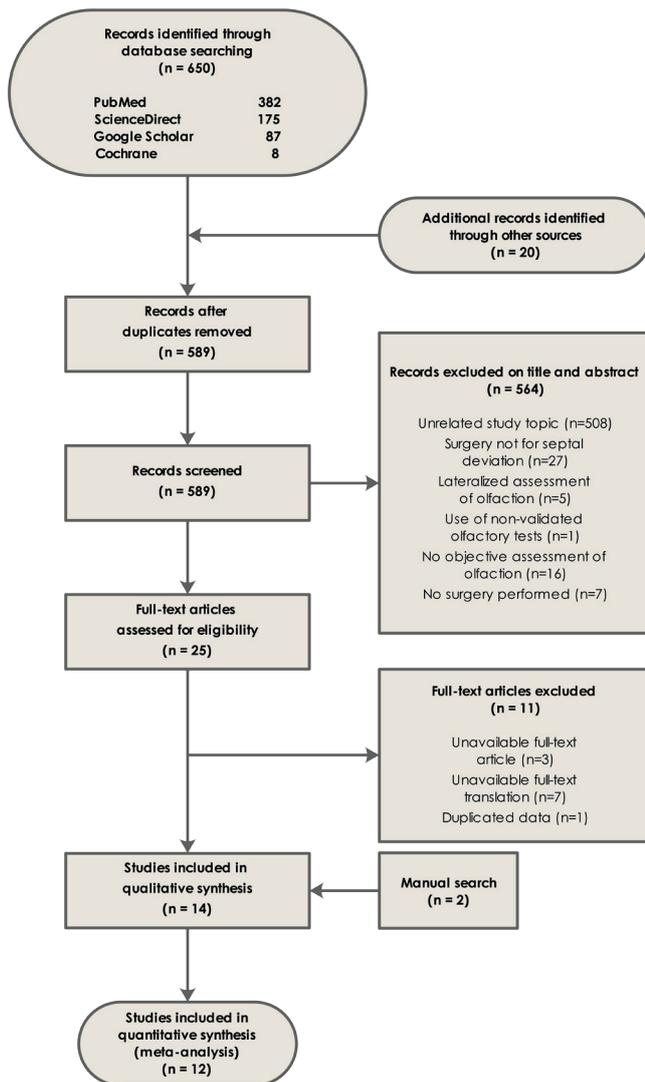


Figure 1. PRISMA Flow Diagram. QoL: quality of life.

Study characteristics

Of the 16 included studies, all were prospective observational studies except 2 (13%) randomized controlled trials^(2,14), 8 studies used some form of the Sniffin' Sticks Test^(2,4,5,10,11,14,17,19) 5 studies used the Connecticut Chemosensory Clinical Research Center (CCCRC) test^(6,13,15,20) and 2 used Brief Smell Identification Test (BSIT) to score olfaction^(12,18). One study used the University of Pennsylvania Smell Identification Test (UPSIT)⁽²¹⁾. The total sample size was 996 patients and 25 controls. Individual studies' size ranged from 30 to 150 patients, follow-up durations ranged from 1 week to 1 year, and patient ages ranged from 10 to 85 years.

Risk of systematic bias

Of the included 16 studies, methodologically 7 were considered to be good^(2,12,14,15,18,20,21), 7 were considered fair^(4-6,13,16,17,19), and 2 were considered poor^(10,11) (Table 2).

Olfactory tests

Seven studies included data on the distribution of patients as normosmic, hyposmic or anosmic. In 4 of those studies the majority of patients were normosmic^(2,15,16,21), whereas in 3 studies the majority of patients had OD before surgery^(4,5,19). The study by Valsamidis et al.⁽⁵⁾ included healthy subjects and found that patients with septal deviation had worse olfaction than controls. Of the 16 studies assessing olfactory function, 11 found a significant improvement in olfactory test scores after septoplasty^(2,4-6,12-15,17-19), whereas 5 found that septoplasty did not improve smell perception^(10,11,16,20,21). Furthermore, Pade et al.⁽¹⁰⁾, Miyake et al.⁽¹⁶⁾, and Randhawa et al.⁽¹⁷⁾ found worsening of olfactory function after septoplasty in 7%, 14.2%, and 7% of their patients, respectively.

BSIT

The total sample size of the included studies using the BSIT test was 169 patients. The calculated difference in means was 0.63 (95% CI, 0.43 to 0.84) (Figure 2A), which represents a 7.01% improvement over the average preoperative BSIT score. The heterogeneity was moderate ($I^2 = 45.64\%$). There was no significant publication bias (Egger's regression intercept $P = 0.2835$) and no trim-and-fill imputations were needed.

CCCRC

Studies using the CCCRC test had a total population of 166 patients. The pooled difference in means was 0.80 (95% CI, 0.26 to 1.33) (Figure 2B), which represents a 17.42% improvement over the average preoperative CCCRC score. However, the heterogeneity was very high ($I^2 = 96.54\%$). The Egger's regression test did not detect evidence of publication bias ($P = 0.9013$) and no trim-and-fill imputations were needed.

Sniffin' Sticks test

Only two of the included studies using the Sniffin' Sticks Test reported the combined score, 5,33 so each of the test's three subtests was analyzed separately. The number of patients administered the complete Sniffin' Sticks Test was 159, with 194 additional patients on the identification subtest. The pooled difference in means was 1.16 (95% CI, 0.55 to 1.76) for odor threshold (Figure 2C), 1.43 (95% CI, 0.86 to 2.00) for odor discrimination (Figure 2D), and 1.18 (95% CI, 0.74 to 1.63) for odor identification (Figure 2E). These differences represent a 20.30%, 13.62%, and 10.23% improvement over the average preoperative scores. Heterogeneity for the Sniffin' Sticks Test was significant, being highest in the identification subtest ($I^2 = 81.78\%$) and lowest in the discrimination subtest ($I^2 = 76.65\%$). Heterogeneity for the threshold test was $I^2 = 81.29\%$. There was no publication bias, as indicated by Egger's regression intercept P -values of 0.3592, 0.7087, and 0.6079 for the threshold, discrimination and

Table 2. Risk of bias within studies.

Item Stud	1	2	3	4	5	6	7	8	9	10	11	12	Total
Damm et al. ⁽⁴⁾ , 2003	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Pade et al. ⁽¹⁰⁾ , 2008	Yes	Yes	Yes	NA	NR	No	Yes	No	No	Yes	No	NA	Poor
Schriever et al. ⁽¹¹⁾ , 2013	Yes	Yes	Yes	NA	NR	No	Yes	No	No	Yes	No	NA	Poor
Dengiz et al. ⁽¹²⁾ , 2015	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Berkiten et al. ⁽¹³⁾ , 2016	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Dalgic et al. ⁽¹⁴⁾ , 2016	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Kilicaslan et al. ⁽¹⁵⁾ , 2016	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Miyake et al. ⁽¹⁶⁾ , 2016	Yes	Yes	Yes	NA	NR	No	Yes	No	No	Yes	Yes	NA	Fair
Randhawa et al. ⁽¹⁷⁾ , 2016	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Haytoglu et al. ⁽¹⁸⁾ , 2017	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Turk et al. ⁽¹⁹⁾ , 2017	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Aydogdu et al. ⁽⁶⁾ , 2019	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Elbistanli et al. ⁽²⁰⁾ , 2019	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Kokubo et al. ⁽²¹⁾ , 2019	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good
Valsamidis et al. ⁽⁵⁾ , 2019	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	No	NA	Fair
Tutar et al. ⁽²⁾ , 2020	Yes	Yes	Yes	NA	NR	No	Yes	No	Yes	Yes	Yes	NA	Good

NA: Not Applicable; NR: Not Reported

identification tests respectively. In no case were trim-and-fill imputations needed.

Sensitivity analysis

Because assumptions were made regarding the correlation coefficients of the pre- and postoperative scores, sensitivity analyses were performed setting the correlation coefficient at 0.8 and 0.5. The results obtained did not differ from the final results.

Discussion

Patients with septal deviation have reduced olfaction compared to the general population⁽⁵⁾, which may negatively affect their daily lives. OD is probably due to a restriction on the airflow by the deviated nasal septum and seems reasonable to think that correcting this anatomical defect would improve olfaction. However, the published data hasn't been able to conclusively provide evidence to support this claim and studies diverge on their findings.

Although surgery can be suggested as the treatment for some cases of OD, nasal surgery may harm olfactory epithelium and worsen olfactory function. This assertion is supported by studies showing that early preoperative ($\leq 6w$) scores had not improved or were even worse than before the septoplasty, but improved at later follow-up times^(12,14,15,18,20). Nasal surgical procedures may distort intranasal anatomy or cause direct trauma to the olfactory epithelium. Likewise, they could have indirect effects arising from pharmacological agents, mucosal edema or blood clots, causing potential damage to the olfactory nerve and com-

promise olfaction^(13,20). Since it can take several months for full recovery of the nasal structures after surgery⁽²⁶⁾, the evidence from studies with short follow-up times should be interpreted carefully.

Furthermore, a finding of some studies was that patients with more severe nasal obstruction had worse baseline olfaction and obtained more benefit from septoplasty compared to those patients with milder obstruction^(3,5). While this is not surprising, the high heterogeneity of assessing nasal obstruction makes it difficult to reliably determine the level of symptom severity that is indicative of significant benefit from septoplasty⁽²⁶⁾.

For those reasons, there is a need to identify prognostic factors for septoplasty and further research is required to find objective methods that better assess the outcome of surgical interventions on the nasal septum. Since it is paramount to any surgical intervention that benefits outweigh its risks, obtaining knowledge of outcome predictors will ensure better patient selection, such that both subjective and objective measures of success can be proven.

Septoplasty seems to improve olfactory function. However, the outcome of this intervention is discrete and not equal for all patients, so more researches are needed.

Limitations

The study is limited by the heterogeneity across studies, which complicates discerning the factors affecting the outcome of septoplasty. A source for this heterogeneity could be differences in patient characteristics such as age or severity of their

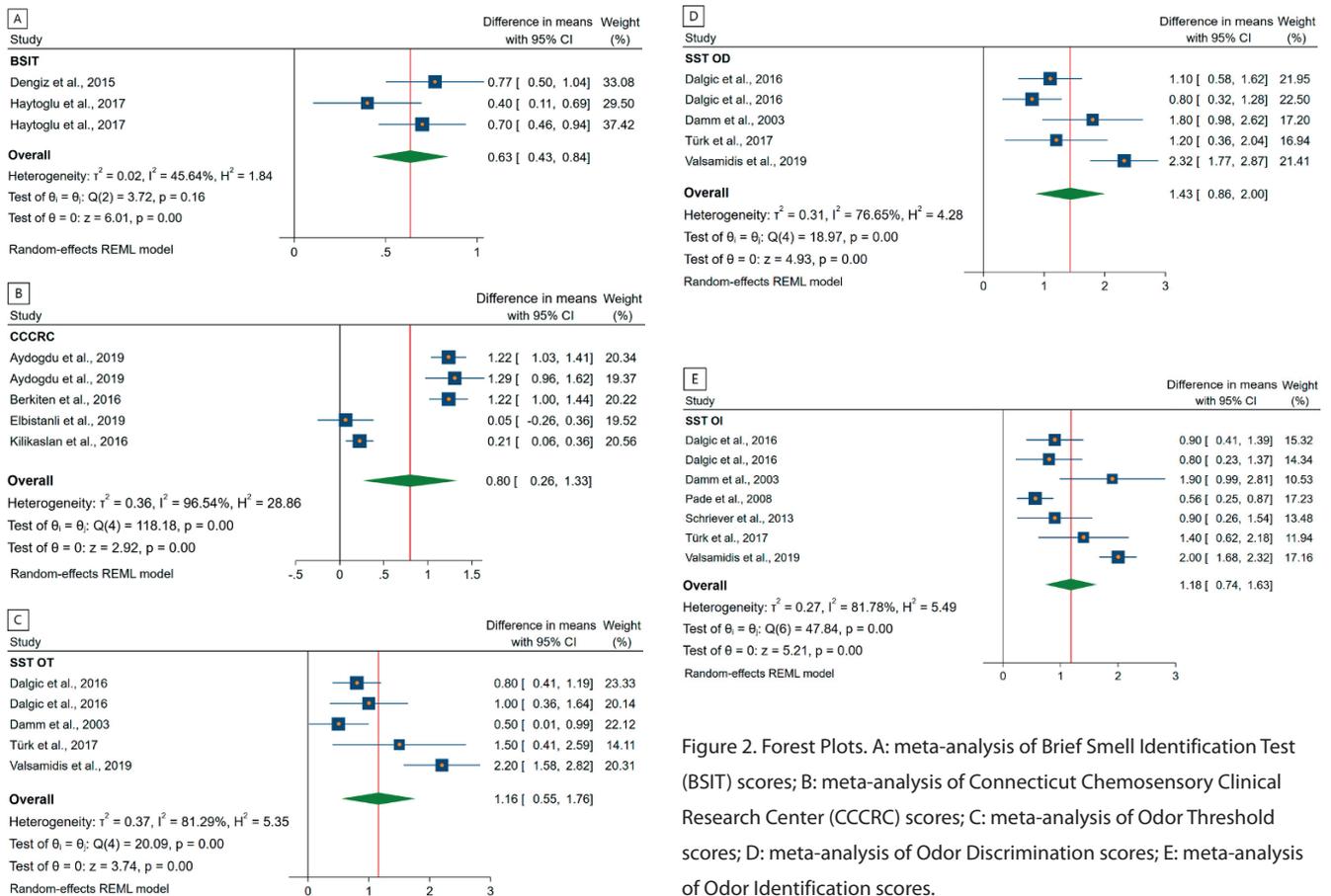


Figure 2. Forest Plots. A: meta-analysis of Brief Smell Identification Test (BSIT) scores; B: meta-analysis of Connecticut Chemosensory Clinical Research Center (CCCRC) scores; C: meta-analysis of Odor Threshold scores; D: meta-analysis of Odor Discrimination scores; E: meta-analysis of Odor Identification scores.

obstruction, as well as presence of concomitant causes of nasal obstruction or other rhinologic conditions. Also, the use of different septoplasty techniques among the included studies and non-standardization of the tools used to assess olfaction could be behind the observed heterogeneity.

Conclusion

Septal deviation compromises the sense of smell and septoplasty seems effective in resolving nasal obstruction due to septal deviation. This review shows that septoplasty has some effect on improving patient's olfaction, yet further randomized trials are

needed to confirm current findings.

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Authorship contribution

AG and PM: Literature review and search, prepare the manuscript. CL and IA: Prepare and review the manuscript

Conflict of interest

None.

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SUPPLEMENTARY DATA

Table S1. Detailed search strategy.

No.	Databases	Search Terms	Result Total = 650
1	PubMed	("Nasal Septum"[Mesh]) AND "Smell"[Mesh]) OR ((smell OR olfaction OR odor) AND (septoplasty OR septum OR septal))	381
2	ScienceDirect	((smell OR olfaction OR odor) AND septoplasty)	174
3	Google Scholar	Where my words occur: in the title of the article: 1. With all of the words: septoplasty With at least one of the words: smell olfaction olfactory odor 2. With all of the words: septal With at least one of the words: smell olfaction olfactory odor 3. With all of the words: septorhinoplasty With at least one of the words: smell olfaction olfactory odor 4. With all of the words: septum With at least one of the words: smell olfaction olfactory odor	15 + 43 + 6 + 23 = 87
4	Cochrane	In all text and ENT Cochrane group: (smell OR olfaction) AND septoplasty	8