Changes in Sinus Membrane Thickness After Lateral Sinus Floor Elevation: A Radiographic Study

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Purpose: To radiographically monitor sinus membrane swelling after lateral sinus floor elevation surgery at short and long healing periods. Materials and Methods: For 26 patients seeking posterior maxillary implant-supported reconstruction, 32 lateral sinus floor elevations were performed using Piezosurgery. Sinus membranes were grafted using synthetic calcium phosphate bone substitutes, and graft volume was measured in cubic centimeters for each case. Cone beam computed tomography (CBCT) examination was conducted preoperatively in all patients and for each grafted sinus at 1 day (n = 8), 2 days (n = 9), 3 days (n = 8), or 7 days (n = 7) after surgery. Control CBCT was then performed for all patients at 3, 6, and 12 months after surgery. Sinus membrane thickness was measured on cross-sectional CBCT images at nine standardized points per sinus, before lateral sinus floor elevation and at all postoperative examinations. Results: Mean sinus membrane thickness was 0.73 mm before surgery, and 5 mm, 4.1 mm, 5.9 mm, and 7 mm, respectively, at 1, 2, 3, and 7 days after surgery. First week combined postoperative CBCT measurements of membrane thickness was 5.4 mm, then 1.3, 0.68, and 0.39 mm at 3, 6, and 12 months, respectively, after surgery. Membrane thickness significantly increased the first week after surgery and gradually decreased significantly at 3, 6, and 12 months in all groups (P < .001). First-week postoperative measurements showed a significant increase in membrane thickness at 3 days compared with the 1- and 2-day results (P < .001) and at 7 days compared with all other time points (P < .001). Membrane thickness at 2 days did not change significantly compared with 1-day measurements. Larger graft volume was positively correlated with an increase in membrane thickness after surgery at all time points (n = 32; r = 0.527; P < .001). Conclusion: After lateral sinus floor elevation surgery, transient swelling of sinus membrane is observed. It reaches a peak value 7 days after surgery and completely resolves over months. This swelling is correlated to the extent of sinus floor elevation. Key words: CBCT, lateral sinus floor elevation, mucosal thickness, posterior maxilla, sinus augmentation, sinus membrane

Sinus floor elevation was introduced to restore an atrophic posterior maxilla. This procedure showed high clinical success whether applied before or in conjunction with implant placement.

Different biomaterials can be predictably used for sinus augmentation such as autogenous bone in blocks from extraoral donor sites or from intraoral sites, allografts, xenografts, and synthetic biomaterials.

Rotary instruments were first used to perform sinus floor elevation, but nowadays, Piezosurgery is the technique of choice for lateral window opening and sinus membrane elevation. Because of their selective action on hard and soft tissues, ultrasonic instruments allow precise and effective osteotomy when preparing a lateral window without compromising soft tissue integrity, thus reducing sinus membrane perforation during surgery.

Radiographic technology is a major tool for lateral sinus floor elevation planning and grafted bone follow-up. Since its introduction, cone beam computed tomography (CBCT) has become the technique of choice for exploring
paranasal sinuses before lateral sinus floor elevation.\textsuperscript{14} CBCT was shown to be very effective for evaluating the presence of anatomical variations and pathologies inside the sinus\textsuperscript{16} and for measuring sinus membrane thickness.\textsuperscript{17}

Maxillary sinus lining membrane consists of a pseudostratified ciliated respiratory epithelium protecting it from exogenous agents by acting as a physical barrier and a mechanical clearing system.\textsuperscript{18} Following sinus membrane damage, initial postsurgical bleeding causes vasoactive substance release, leading to primary clot formation underneath it. It is followed by an inflammatory phase characterized by membrane transitory swelling that predominates at 48 hours and subsides over a period of several days.\textsuperscript{19}

Following lateral sinus floor elevation, radiographic studies have not shown substantial changes in sinus membrane thickness at long-term healing periods.\textsuperscript{20} Nevertheless, very few studies described early changes in the membrane after this procedure.

In a clinical study, Quirynen et al\textsuperscript{21} described a transient swelling of sinus membrane 1 week after crestal sinus floor elevation, which returned to normal after 3 weeks. Membrane thickness changes after sinus surgery are believed to occur because of the expression of swelling and postsurgical inflammatory reaction; the present radiographic study observed membrane thickness behavior after lateral sinus floor elevation at early and late healing times using CBCT.

**MATERIALS AND METHODS**

A prospective clinical study was conducted to assess clinical, histologic, and radiographic performance of eight different bone substitutes grafted into the maxillary sinus, and the behavior of sinus membrane after lateral sinus floor elevation was observed. Twenty-six patients seeking 32 lateral sinus floor elevations were included in this study. All treated patients had pneumatized sinuses with residual sinus floor heights of less than 5 mm. All patients were in good health, with no systemic disorders. All were accurately informed about the procedure, and all signed an informed consent form. The study was approved by the ethics committee at Saint Joseph University (Beirut, Lebanon).

Preoperative examination included CBCT as well as clinical examination. All surgeries were performed at the Oral Surgery Department, Faculty of Dental Medicine at Saint-Joseph University.

**Surgical Procedure**

Patients were asked to rinse with chlorhexidine digluconate solution (0.2%) for 1 minute approximately 10 minutes before surgery. Under localized anesthesia, a crestal incision and a full-thickness flap elevation were performed to expose the buccal wall of the maxillary sinus. Lateral window opening was performed using Piezosurgery (Mectron). With standard elevators and Piezosurgery inserts specific to this procedure, the sinus membrane was elevated. Metronidazole solution (0.5%) was used to wash the sinus and hydrate the graft as described by Choukroun et al.\textsuperscript{22} The sinus membrane was then grafted using synthetic calcium phosphate bone substitutes (Eurocoating, Trento, Italy), and the graft volume, measured in cubic centimeters, was noted for each case. Four types of synthetic calcium phosphate biomaterials were used in this study: pure phase HA, pure phase bTCP, biphasic HA/bTCP (60%/40%), and biphasic HA/TTCP (60%/40%). In each graft type, two different granulometries were used, round 500- to 1,000-µm granules and 1,200- to 5,000-µm aggregates of round granules. The eight different combinations of biomaterials and granule sizes were randomly and evenly used for all 32 sinus grafts included in this study, leading to four grafted sinuses per biomaterial type/granule size.

A collagen membrane was placed to cover the lateral window, and flaps were approximated and sutured. Patients were prescribed analgesics (ibuprofen 400 mg twice a day), local steroid and prophylactic antibiotic coverage (amoxicillin + clavulanic acid 2 g daily, or in case of allergy, clindamycin 600 mg daily) for 7 days, and oral rinses of 0.12% chlorhexidine gluconate for 15 days after implant placement. Patients were recalled for suture removal 7 days after surgery and were clinically and radiographically followed up until implant placement and final prosthetic reconstruction.

**Radiographic Examination**

CBCT was performed before surgery using the same Newtom VGI radiographic machine (QR) for all patients. After lateral sinus floor elevation, patients were randomly assigned to four different groups to perform CBCT for each grafted sinus at 1 day (n = 8), 2 days (n = 9), 3 days (n = 8), or 7 days (n = 7) after surgery. Postoperative control CBCT was then performed for all patients at 3 months, 6 months, and 12 months.

Sinus membrane thickness was measured to the nearest 0.1 mm at three different locations in the maxillary sinus by two different investigators (C.M. and A.M.). To define these locations, a horizontal line was drawn on the panoramic CBCT view along the maximal mesiodistal length of the sinus, and three measuring planes were selected at quarter, half, and three quarters that distance (Fig 1). For each of these locations, the corresponding cross-sectional CBCT image was retrieved and three different measurements were performed on it. To locate the measurement points, a first vertical line passing through the middle of the crest was drawn (midline); a middle (m) measurement of the sinus membrane thickness was taken on the intersection between the midline and sinus floor. A second horizontal line perpendicular to the midline was drawn 5 mm above
the sinus mucosa, and the intersections between this line and both palatal and buccal sinus walls, at two different points, were defined as palatal (p) and buccal (b). The sinus membrane thickness was then measured at these points (Fig 2). These measurements were repeated at the three sinus locations described earlier, leading to a total of nine standardized measuring points per sinus. Measurements were done before the lateral sinus floor elevation, at the first-week examinations (1, 2, 3, and 7 days), and at the 3-, 6-, and 12-month postoperative examinations.

The combined mean value of 1-, 2-, 3-, and 7-day results was calculated and defined as the first-week postoperative combined result.

Statistical Analyses
Statistical analyses were performed using the SPSS for Windows, version 18.0 software program (SPSS). The level of significance was set at $\alpha = .05$. Repeated-measures analysis of variance was conducted to explore significant differences in mean membrane thickness at baseline, first-week postoperative combined, 3 months, 6 months, and 12 months. This analysis was followed by Bonferroni multiple comparison.

One-way analysis of variance followed by Tukey (honest significant difference) post-hoc tests were also conducted to explore a significant difference in first week mean sinus membrane thickness among day 1, day 2, day 3, and day 7 values.

The relationship between graft volume and sinus membrane thickness for each time point was investigated using a Pearson product-moment correlation coefficient. Preliminary analyses were performed to ensure that normality, linearity, and homoscedasticity assumptions were not violated.

RESULTS
All 26 patients selected for the study underwent 32 lateral sinus floor elevations successfully and with no major complication.

Preoperative sinus membrane mean thickness as evaluated on CBCT was 0.73 mm.

First-week CBCT images indicated mean sinus membrane thickness of 5 mm, 4.1 mm, 5.9 mm, and 7 mm at 1 day, 2 days, 3 days, and 7 days, respectively. Mean membrane thickness increases significantly at 3 days compared with results at 1 day and 2 days ($P < .001$) and at 7 days compared with all other time points ($P < .001$). There was no significant change in membrane thickness at 2 days compared with 1-day results (Figs 3 and 4).

Mean sinus membrane thickness for the combined first-week postoperative measures was 5.4 mm, decreasing to 1.3 mm, 0.68 mm, and 0.39 mm at 3, 6, and 12 months, respectively (Figs 5 and 6). Sinus membrane thickness increased significantly the first week after surgery and then decreased significantly at 3, 6, and 12 months ($P < .001$). Mean sinus membrane thickness was significantly lower 12 months after surgery than before surgery ($P = .048$).

A positive correlation was noted between larger graft volume and membrane thickness immediately after surgery at all time points ($n = 32; r = 0.527; P < .001$).

No significant correlation was noted between sinus membrane thickness and type of synthetic bone substitute ($P = .359$) or between sinus membrane thickness and size of biomaterial particles ($P = .293$).
**Fig 3** (left) Mean Shneiderian membrane thickness at early postoperative periods.

**Fig 4** (below) Typical sinus membrane thickness as seen on CBCT cross-sectional images in different patients at (a) 1 day, (b) 2 days, (c) 3 days, and (d) 7 days after surgery.

**Fig 5** (left) Mean sinus membrane thickness over a 12-month period. Preop = preoperative; postop = postoperative.

**Fig 6** (below) Radiographic follow-up findings of changes in sinus membrane thickness for the same patient (a) before surgery and at (b) 7 days, (c) 3 months, (d) 6 months, and (e) 12 months after surgery.
DISCUSSION

In the present study, sinus membrane thickness was measured with CBCT. Although an automated Hounsfield threshold-based computed tomography evaluation has been used to predictably determine sinus membrane thickness, the present study measured sinus membrane thickness on CBCT images following linear measurements at two points. Because this technique may have some limitations with regard to measurement reproducibility, all calculations were done after agreement between two different evaluators.

Multiple studies showed large variability in sinus membrane thickness; some studies reported postmortem sinus mucosa measurements of a mean value of 0.8 mm in noninfected sinuses, whereas others described a mean value of 0.09 mm. This intervariability in sinus mucosa thickness was described by Janner et al in a radiographic study that showed a wide range of measurements varying between 0.16 and 34.61 mm.

Sinus mucosa thickness in this study varied between 0.1 and 3.6 mm, with a mean value of 0.7 mm. The present study enrolled only those patients who were suitable for sinus surgery with no evident sinus pathologies; therefore, preoperative sinus membrane thickness measurements were probably lower than those described in other studies, which manifestly enrolled patients regardless of pre-existing pathologic thickening of sinus mucosa. Moreover, many factors may influence sinus membrane thickness, such as gender, climate, smoking habits, allergies, and seasonal changes. Six of the 26 patients included in the present study underwent bilateral lateral sinus floor elevation and were among the 32 sinuses with follow-up data. Consequently, the aforementioned individual factors might be a source of bias in sinus mucosa behavior described in this article.

The main observation of the present study was a significant transient thickening of sinus membrane after lateral sinus floor elevation surgery. This thickening may be caused by a physiologic inflammatory reaction to surgical trauma involving bleeding and swelling at initial healing phases. Although nasal and paranasal mucosa reaction to different injuries such as infection, allergy, or surgical trauma have been described, sinus membrane reaction to lateral sinus floor elevation is poorly described in the literature. In 2002, Watelet et al clearly described the process of nasal and paranasal mucosa wound healing after injury. These authors divided this process into four overlapping phases: inflammation, cell proliferation, matrix deposition, and tissue remodeling. However, these recovery stages are described in the context of sinus surgery involving partial or total resection of sinus membrane, whereas in lateral sinus floor elevation, the sinus mucosa is kept intact and is only detached from underlying bone. Scala et al, after performing lateral sinus floor elevation in monkeys, sacrificed the animals at 4, 10, and 30 days and performed a histologic examination. The authors described the presence of a coagulum underneath the sinus membrane along with a dense inflammatory infiltrate inside this membrane at 4 days. Inflammatory infiltrate significantly reduced at 10 days, whereas at 30 days, the sinus mucosa appeared normal with only a few inflammatory cells.

Quirynen et al, in a radiographic study, were the first to describe early sinus mucosa behavior after crestal sinus floor elevation. They showed a significant swelling of sinus mucosa 1 week after surgery, which totally disappeared after 1 month. The radiographic results of the present study showed membrane thickening after lateral sinus floor elevation with a mean value of 5.4 mm during the first-week follow-up. This value is lower than the mean value of 6.7 mm measured by Quirynen et al. This difference may be explained by the fact that these authors only calculated 7 days’ results, whereas the present study also takes into account findings at 1, 2, and 3 days. When considering only 7-day postoperative values, results of this study showed a mean sinus mucosa thickness of 7 mm, which is similar to previously described findings.

The present study interestingly showed a significant increase in sinus membrane thickening at 3 and 7 days compared with the lower increase observed at 1 and 2 days after surgery. The lower values encountered at 1 and 2 days may be related to the early wound healing phase that involves primary clot formation, whereas the inflammatory phase starts at 48 hours. This inflammatory phase, characterized by a larger infiltration of polymorphonuclear cells inside the sinus membrane, is clearly demonstrated by significantly higher thickening of sinus membrane at 3 and 7 days, and reaching up to 10 times its original thickness.

In some instances, early hemorrhagic fill of the sinus that might occur after surgery may mimic pathologic or inflammatory thickening of sinus membrane observed on radiographs. Nevertheless, this radiographic finding shows a horizontal fill of the sinus, and is different from membrane swelling, which can also be observed on the lateral aspect of sinus walls. In the present study, lateral wall membrane swelling was taken into consideration by measuring membrane thickness at the palatal, middle, and buccal part of the sinus to obtain more reliable measurements.

To the best of the authors’ knowledge, this is the first longitudinal study monitoring sinus membrane swelling at different early healing periods, along with a follow-up period of 3, 6, and 12 months. In the present study, sinus mucosa at 3 months was significantly thicker than it was before surgery, whereas the results of other authors showed a return to normal of the sinus mucosa 1 month after surgery. To explain the apparent contradiction,
the previously cited study applied a crestal sinus approach, which involves less invasive surgery, and therefore minimal detachment of sinus mucosa. The present study, however, describes sinus mucosa behavior after sinus floor elevation using a lateral approach, which requires an extensive detachment of the membrane along with considerable flap elevation and a large bone access window.

A positive significant correlation between sinus graft volume and membrane thickening at early postoperative periods was observed in the present study. This observation supports the authors’ hypothesis that larger sinus elevation may lead to more pronounced membrane thickening even at 3 months after surgery.

The persistent thickening of sinus mucosa observed in this work may be the result of altered physiologic function after surgical trauma. Studies have reported histologic alterations of sinus mucosa after sinus surgery, leading to a decrease in its physiologic behavior mainly characterized by cleaning and clearance of the surface.29 After lateral sinus floor elevation, animal studies showed histologic changes in the sinus mucosa, characterized by reduction of mucociliary cells and increase of goblet cells.29,31 Clinical studies have shown mucociliary activity impairment in the detached part of this mucosa.32 Timmenga et al,33 in a human histologic study, described an increase of goblet cells at 3 and 9 months after lateral sinus floor elevation compared with preoperative histology. The authors explained this cellular modification as a physiologic adaptation that the sinus membrane undergoes over months. The same authors also examined the microbiology of sinus mucosa surface and described an increase in cultures with bacterial growth at 3 months after sinus floor elevation. The authors hypothesize that the maxillary sinus mucosal lining was affected after the surgical procedure, leading to the impairment of mucusal defense system mainly characterized by transitory reduction of sinus drainage.

Sinus membrane behavior at 3 months observed in the present study may be the result of the physiologic and microbiologic changes described before, leading to a temporary impaired function of sinus mucosa and causing a slight persistent but transitory inflammation.

The present study also indicated that sinus mucosa thickening was significantly reduced at 6 months compared with 3 months, reaching its preoperative normal value. This observation is in contradiction with those of other authors who described a significant difference in postoperative sinus thickness that persisted at 6 and 9 months compared with preoperative results.23,34 Moreover, sinus mucosa thickening was significantly reduced at 12 months compared with findings at 6 months, and showed a mean thickness of 0.12 mm that was also significantly lower than preoperative values.

This gradual decrease in membrane thickness that goes on until 1 year after surgery may be caused by the physiologic readaptation of sinus mucosa along with an enhanced activity of ciliary epithelium, leading to a better cleansing of sinus surface. This is in accord with Timmenga et al,33 who described a recovery of both microbiologic flora and histologic characteristics of sinus membrane at 9 months after surgery. Also, in an animal study, histologic examination 6 months after lateral sinus floor elevation showed a totally healthy aspect of sinus mucosa.35

Findings at 12 months surprisingly showed a significant reduction of membrane thickness compared with preoperative measurements. This may be explained by an improvement in sinus mucosa physiologic function related to the aforementioned histologic changes occurring in this mucosa after surgical trauma. On the other hand, Forsgren et al,36 in an animal study, described atrophic sinus membrane with a decreased thickness after complete sinus membrane removal. In the present work, sinus mucosa was not removed but only elevated. Nevertheless, this reduction in membrane thickness may also be caused by sinus mucosa atrophy after surgical trauma. Further histologic studies should be conducted to confirm both hypotheses.

The choice of grafting materials for sinus augmentation may play a role in membrane behavior at longer postoperative healing periods because it is obvious that a biomaterial that can evoke a higher immunologic response may lead to an inflammatory reaction. In a series of endoscopic observations at long healing periods, Aimetti et al37 described a slightly inflamed sinus mucosa over allografts and heterografts compared with totally healthy mucosa when the sinus was grafted with autogenous bone. In the present study, four different allografts calcium phosphate biomaterials were used, and data showed that these biomaterials did not generate an inflammatory response of the sinus membrane at long healing times because its thickness returned to normal at 6 months. Also, both grafting particle sizes used in this study did not lead to any differences in membrane behavior at short and long healing periods.

CONCLUSIONS

After lateral sinus floor elevation surgery, the sinus mucosa undergoes significant thickening. This thickening is believed to be a physiologic inflammatory reaction that is more pronounced at 3 and 7 days after surgery compared with the results at 1 and 2 days. Long-term observations revealed a slight but persistent swelling at 3 months, which completely disappeared at the 6-month follow-up when sinus mucosa reached its previous preoperative thickness. At 12 months, observations interestingly showed a significant reduction of membrane thickness compared with its preoperative size.
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REFERENCES