Maxillomandibular Advancement for OSA: A 25-year perspective

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ABSTRACT - Objective: The aim of this study was to evaluate the result of maxillomandibular advancement (MMA) for the treatment of obstructive sleep apnea (OSA) by a single surgeon. Materials and Methods: Patients that underwent MMA for the treatment of OSA over a 25-year period were included in the study. Patients who initially presented for revision MMA surgery were excluded. Demographics (e.g., age, gender, pre- and post-MMA body mass index [BMI]), pre- and post-MMA cephalometrics (e.g., sella-nasion-point A angle [SNA], sella-nasion-point B angle [SNB], posterior airway space base of tongue [PAS]) and pre- and post-MMA sleep study metrics (e.g., respiratory disturbance index [RDI], lowest desaturation [SpO₂nadir], oxygen desaturation index [ODI], total sleep time [TST], % TST Stage N3 sleep, % TST rapid eye movement [REM] sleep) were abstracted. MMA surgical success was defined as $a \ge 50\%$ reduction in RDI (or ODI) and post-MMA RDI (or ODI) < 20 events/hour. MMA surgical cure was defined as a post-MMA RDI (or ODI) < 5 events/hour. **Results:** A total of 1010 patients underwent MMA for the treatment of OSA. The mean age was 39.6 ± 14.3 years, and the majority were male (77%). Nine hundred forty-one patients with complete pre- and postoperative PSG data were analyzed. The mean ODI and RDI improved from 32.6 ± 27.4 to 7.7 ± 15.5 and 39.1 \pm 24.2 to 13.6 \pm 14.6 events per hour, respectively. The overall surgical success and surgical cure based on ODI was 79.4% and 71.9%, respectively. The overall surgical success and surgical cure based on RDI was 73.1% and 20.7%, respectively. Stratified by preoperative RDI showed older age, greater BMI were associated with greater preoperative RDI. Bivariate predictors of greater RDI reduction include younger age, female gender, lower preoperative BMI, higher preoperative RDI, greater BMI reduction postoperatively and greater change in SNA and PAS. Bivariate predictors of surgical cure based on RDI (RDI < 5) include younger age, female gender, lower preoperative RDI, and greater change in SNA and PAS. Bivariate predictor of RDI success (RDI < 20) include younger age, female gender, lower preoperative BMI, lower preoperative RDI, greater BMI reduction, greater increase in SNA, SNB and PAS postoperatively. Comparison of the first 500 patients and the later 510 patients demonstrate patients undergoing MMA have become younger, having lower RDI while achieving a better surgical outcome. Linear multivariate associations of greater percentage RDI reduction include younger age, greater percent change of SNA, greater preoperative SNA, lower preoperative BMI and higher preoperative RDI. Conclusions: MMA is an effective treatment to improve OSA, but the result can vary. Patient selection based on favorable prognostic factors and maximizing the advancement distance can improve outcomes.

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1. Introduction

Maxillomandibular advancement (MMA) has become a well-accepted treatment of obstructive sleep apnea (OSA) since the early reports over 40 years ago^{3,10,30}. However, despite numerous published case series on MMA results, prognostic factors are not well-established due to limited sample size. To improve the understanding of the impact of MMA on OSA, meta-analyses of pooled MMA data have been conducted. Patients with younger age and weight, lower apnea hypopnea index (AHI), and a greater degree of maxillary advancement were predictive of increased surgical success based on analysis of 320 subjects8. In a follow-up analysis of 518 subjects, preoperative AHI of fewer than 60 events per hour was shown to be the factor most strongly associated with surgical cure (AHI < 5)31. The rates of surgical success (AHI < 20) and cure (AHI < 5) were 389 (85.5%) and 175 (38.5%), respectively, among 455 patients with AHI data³¹. However, among the 68 patients with respiratory disturbance index (RDI) data, the impact on OSA was less dramatic. The rate of success and cure were 44 (64.7%) and 13 (19.1%), respectively.

Over the past 25 years, the authors have extensively studied the relationship between the airway and the maxillofacial skeleton. The principles of maximizing skeletal advancement, maintaining proper facial balance and occlusion while minimizing complications have been established¹¹⁻²⁹. This study was undertaken to assess the outcome of MMA for the treatment of OSA by a single surgeon (KKL).

2. Materials and Methods

Patients undergoing MMA for the treatment of OSA performed by a single surgeon (KKL) between 1997 and 2021 were retrospectively analyzed from prospectively collected clinical data. Patients with previous MMA that presented for revision surgery were excluded. The study was approved by the Stanford University review board (protocol ID # 15494) and was in accordance of the Declaration of Helsinki.

2.1. OSA Assessments

All patients underwent in-lab type-1 monitored polysomnography evaluations pre- and post-MMA that were performed per American Academy of Sleep Medicine's guidelines¹. The respiratory-dis-

turbance index (RDI) was calculated as the number of apneas (\geq 90% reduction in thermistor flow for \geq 10 seconds) plus hypopneas (\geq 30% reduction in nasal pressure amplitude accompanied by either a \geq 3% pulse oximeter (SpO₂) reduction or electroence-phalogram (EEG) arousal for \geq 10 seconds) per hour of sleep. The oxygen desaturation index (ODI) was calculated as the number of \geq 3% SpO₂ reductions from baseline per hour of sleep. Sleep apnea severity was defined as mild (5-14.9 events/hour), moderate (15-29.9 events/hour) or severe (\geq 30 events/hour).

2.2. Medical Record and Data Abstraction

Demographics (e.g., age, gender, pre- and post-MMA body mass index [BMI]), pre- and post-MMA cephalometrics (e.g., sella-nasion-point A angle [SNA], sella-nasion-point B angle [SNB], posterior airway space base of tongue [PAS]) and pre- and post-MMA sleep study metrics (e.g., RDI, SpO_2 -nadir, ODI, total sleep time [TST], %TST Stage N3 sleep, %TST rapid eye movement [REM] sleep) were abstracted. MMA surgical success was defined as a \geq 50% reduction in RDI (or ODI) and post-MMA RDI (or ODI) < 20 events/hour. MMA surgical cure was defined as a post-MMA RDI (or ODI) < 5 events/hour.

2.3. Analyses

All univariate and multivariate analyses were performed using Statistical Package for the Social Sciences version 22 for Windows (SPSS, Chicago, IL, USA). We compared categorical variables using a Fisher's exact test and continuous variables with a 2-tailed t-test (or a one-way analysis of variance [ANOVA] test as appropriate). The normality assumption for the ANOVA was assessed via the Kolmogorov-Smirnov test and a nonparametric test used for p-value calculations. Values are reported as mean ± standard deviation. A multivariate linear regression (dependent variable % reduction in RDI) was used to estimate the associations between baseline demographics, cephalometrics, and sleep study metrics, as well as changes in cephalometrics post-MMA, in relationship to improvements in sleep apnea metrics. We assessed each variable by stepwise backwards regression using a p-value cutoff ≤ 0.1 . To ensure uniformity between covariate scaling, z-scores of these variables were generated (each variable divided by its standard deviation) resulting in all regression beta values having scaled

Table 1. Pre and Post-MMA Characteristics*.

	Pre-MMA						
	Pre-IMIMA	Post-MMA	P-value				
Characteristics							
Age (n=1010)	39.6 ± 14.3	NA	NA				
Male Gender (%) (n=1010)	77.0%	NA	NA				
BMI (kg/m²)¶ (n=908)	27.7 ± 4.3	27.5 ± 4.3	<0.001				
Polysomography							
RDI (events/hour) (n=941)	39.1 ± 24.2	13.6 ± 14.6	<0.001				
RDI ≥ 30/hr (%) (n=941)	54.8%	8.5%	<0.001				
RDI ≥ 20/hr (%) (n=941)	77.1%	17.2%	<0.001				
SpO ₂ Nadir (%) [‡] (n=922)	83.4 ± 10.1%	88.9 ± 6.4%	<0.001				
ODI (desaturations/hour) (n=306)	32.6 ± 27.4	7.7 ± 15.5	<0.001				
ODI ≥ 30/hr (%) (n=306)	47.7%	8.5%	<0.001				
ODI ≥ 20/hr (%) (n=306)	55.9%	11.1%	<0.001				
Sleep Stages†	55.9%	11.1%	<0.001				
Total Sleep Time (min) (n=306)	363.9 ± 68.2	378.8 ± 67.5	0.002				
REM Sleep (%) (n=306)	10.6 ± 5.3%	16.9 ± 6.4%	<0.001				
Stage N3 Sleep (%) (n=306)	3.4 ± 5.3%	7.6 ± 7.7%	<0.001				
Cephalometrics							
SNA (degrees) (n=305)	80.1 ± 3.1	81.5 ± 3.2	<0.001				
SNB (degrees) (n=305)	77.1 ± 3.8	78.4 ± 3.3	<0.001				
PAS (mm) (n=305)	5.4 ± 2.2	8.4 ± 2.5	<0.001				
MP-H (mm) (n=305)	25.0 ± 7.1	22.1 ± 7.0	<0.001				
PNS-P (mm) (n= 305)	45.1 ± 4.9	30.9 ± 3.7	<0.001				
RDI Surgical Cure (%)# (n=941)	NA	20.7%	NA				
ODI Surgical Cure (%)# (n=306)	NA	71.9%	NA				
RDI Surgical Success (%)# (n=941)	NA	73.1%	NA				
ODI Surgical Success (%)# (n=306)	NA	79.4%	NA				

^{*} Plus-minus values are mean (or percent) ± standard deviation. Values in parenthesis are the number of patients evaluated. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space – base of tongue), PNS-P (distance of posterior nasal spine to the tip of the soft palate), RDI (respiratory disturbance index), REM (rapid eye movement), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SPO₂ (pulse oxyhemoglobin saturation). † Reported as a percent of total sleep time. ‡ The SPO₂ nadir is the lowest oxyhemoglobin saturation measured during sleep. Only patients with both pre and post-MMA SPO₂ nadir are included. # Surgical cure defined an RDI (or ODI) < 5 events/hour post-MMA Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA. ¶ Only patients with both pre and post-MMA BMI are included.

standardized units of the independent predictors. A two-tailed p-value < 0.05 was considered statistically significant.

3. Results

A total of 1010 patients underwent MMA for the treatment of OSA. The mean age was 39.6 \pm 14.3 years, and the majority were male (77%).

Nine hundred forty-one patients with complete pre- and postoperative PSG data were analyzed. The mean ODI and RDI improved from 32.6 ± 27.4 to 7.7 ± 15.5 and 39.1 ± 24.2 to 13.6 ± 14.6 events per hour, respectively. The overall surgical success and surgical cure based on ODI was 79.4% and 71.9%, respectively. The overall surgical success and surgical cure based on RDI was 73.1% and 20.7%, respectively (Table 1).

Stratified by preoperative RDI showed older age, greater BMI were associated with greater preoperative RDI (Table 2). Bivariate predictors of greater RDI reduction include younger age, female gender, lower preoperative BMI, higher preoperative RDI, greater BMI reduction postoperatively and greater change in SNA and PAS (Table 3). Bivariate predictors of surgical cure based on RDI (RDI < 5) include younger age, female gender, lower preoperative RDI, and greater change in SNA and PAS (Table 4). Bivariate predictor of RDI success (RDI < 20 include younger age, female gender, lower preoperative BMI, lower preoperative RDI, greater BMI reduction, greater increase in SNA, SNB and PAS postoperatively (Table 5). Comparison of the first 500 patients and the later 510 patients demonstrate patients undergoing MMA have become younger, having lower RDI while achieving a better surgical outcome (Table 6).

Linear multivariate associations of greater percentage RDI reduction include younger age, greater percent change of SNA, greater preoperative SNA, lower preoperative BMI and higher preoperative RDI (Fig. 1).

4. Discussion

The results of this study demonstrate that patients with younger age, lower BMI, lower RDI and female gender achieve a better surgical outcome as measured by PSG parameters. Furthermore, a larger advancement distance (percentage increase of SNA and SNB) and a greater impact on airway size (larger increase in PAS) resulted in a greater reduction in RDI in achieving surgical success.

The application of MMA in treating OSA has evolved over the past 25 years¹¹⁻²⁹. MMA has changed from a last stage operation of the phased surgical protocol to a primary, single stage treat-

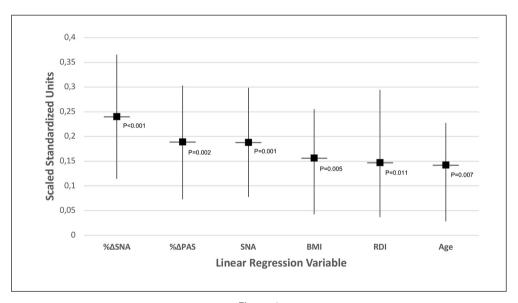


Figure 1

Multivariate Associations with Percent RDI Reduction Post MMA. Linear regression model with percent reduction in RDI (continuous variable) as dependent variable. All regression beta values are scaled standardized units of the independent predictors. Regression beta point estimate with 95% confidence intervals shown for each included variable. Initial model variables included age, pre-MMA BMI, male gender, pre-MMA RDI, pre-MMA SNA, pre-MMA SNB, pre-MMA PAS, post-MMA BMI percent change, post-MMA percent change SNA, post-MMA percent change SNB, and post-MMA percent change PAS. Final model developed utilizing backwards elimination (p-value < 0.10 cutoff). Predictors of increased percent RDI reduction post MMA in final model include greater positive percent change in SNA (β = 0.240, p < 0.001), greater positive percent change in PAS (β = 0.189, p = 0.002), greater pre-MMA SNA angle (β = 0.188, p = 0.001), lower pre-MMA BMI (β = 0.154, p = 0.005), greater pre-MMA RDI (β = 0.147, p = 0.011), and younger age (β = 0.142, p = 0.007). Model R2 is 0.201 (adjusted for number of predictors). Including various interaction terms (age*RDI, BMI*RDI, Male*RDI, SNA*RDI, SNB*RDI, PAS*RDI, Male*BMI, Age*Male) and/or polynomial variables (age2, BMI2, RDI2, RDI3) did not improve data fit and were not included in final model. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), PAS (posterior airway space –base of tongue), RDI (respiratory disturbance index), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), % Δ PAS (percent PAS reduction), and % Δ SNA (percent SNA reduction).

Table 2. Patient Characteristics Stratified by Pre-MMA RDI*.

		Dueluet			
Characteristics	< 30/hour (n=425)	30-59.9/hour (n=338)	≥ 60/hour (n=178)	P-value [†]	
Pre-MMA					
Age (years)	33.7 ± 13.9 (425)	44.5 ± 12.3 (338)	45.7 ± 12.2 (178)	<0.001	
Male (%)	69.9% (425)	81.4% (338)	91.6% (178)	<0.001	
BMI (kg/m²)	26.0 ± 2.5 (412)	27.8 ± 4.1 (338)	31.4 ± 5.6 (175)	<0.001	
BMI > 30 kg/m ² (%)	5.9% (412)	20.1% (338)	52.2% (175)	<0.001	
Cephalometrics					
SNA (degrees)	80.2 ± 2.7 (60)	80.6 ± 3.2 (117)	79.8 ± 3.2 (136)	0.118	
SNB (degrees)	77.3 ± 3.6 (60)	77.3 ± 3.9 (117)	76.9 ± 3.8 (135)	0.765	
PAS (mm)	6.3 ± 2.4 (60)	5.5 ± 2.2 (111)	4.9 ± 2.0 (135)	0.602	
Polysomnography					
RDI (events/hour)	19.2 ± 5.9 (425)	42.7 ± 9.0 (338)	79.7 ± 15.8 (178)	<0.001	
SpO ₂ Nadir (%)	88.9 ± 5.4% (421)	82.9 ± 8.2 (331)	71.4 ± 10.9 (177)	<0.001	
ODI (desaturations/hour)	7.4 ± 6.9 (60)	19.9 ± 14.9 (111)	54.3 ± 24.7 (135)	<0.001	
REM Sleep (% TST)	13.5 ± 5.6% (60)	11.1 ± 4.5% (111)	8.8 ± 5.2 (135)	0.003	
Stage III/IV Sleep (% TST)	6.8 ± 7.4% (60)	3.4 ± 4.6% (111)	1.8 ± 3.7% (135)	0.002	
Post-MMA					
BMI (kg/m²)	25.9 ± 2.6 (379)	27.7 ± 4.2 (308)	31.4 ± 5.6 (172)	<0.001	
Percent Change in BMI (%)	-0.7 ± 1.3% (378)	-0.7 ± 1.2% (303)	-0.5 ± 3.5% (172)	0.443	
Cephalometrics					
SNA (degrees)	80.6 ± 2.9 (59)	81.7 ± 2.9 (111)	81.8 ± 3.5 (135)	0.039	
Change in SNA (degrees)	0.004 ± 0.02 (59)	0.01 ± 0.02 (111)	0.02 ± 0.03 (135)	<0.001	
SNB (degrees)	77.5 ± 0.03 (59)	78.5 ± 0.03 (111)	78.7 ± 0.03 (135)	0.055	
Change in SNB (degrees)	0.003 ± 0.01 (59)	0.01 ± 0.02 (111)	0.02 ± 0.02 (135)	<0.001	
PAS (mm)	8.2 ± 2.6 (59)	8.5 ± 2.3 (111)	8.5 ± 2.6 (135)	0.639	
Change in PAS (mm)	1.9 ± 2.2 (59)	3.0 ± 2.1 (111)	3.7 ± 2.4 (135)	<0.001	
Polysomnography					
RDI (events/hour)	8.2 ± 6.1 (425)	15.2 ± 11.7 (338)	23.2 ± 24.9 (178)	<0.001	
Percent Change in RDI (%)	-56.0 ± 35.3% (425)	-63.8 ± 27.6% (338)	-70.0 ± 32.1% (178)	<0.001	
RDI 5-14.9/hour (%)	58.6% (425)	47.3% (338)	36.0% (178)	<0.001	
RDI 15-29.9/hour (%)	11.5% (425)	29.3% (338)	25.3% (178)	<0.001	
RDI ≥ 30/hour (%)	0.1% (425)	4.0% (338)	4.4% (338)	<0.001	
SpO, Nadir (%)	91.8 ± 3.9% (418)	88.5 ± 4.5% (332)	83.3 ± 9.3% (177)	<0.001	
Percent Change in SpO ₂ Nadir (%)	3.5 ± 5.5% (417)	7.6 ± 10.9% (328)	18.5 ± 17.5% (177)	<0.001	
REM Sleep (% TST)	17.1 ± 6.1% (60)	16.2 ± 6.2% (111)	17.1 ± 6.7 (135)	0.829	
Percent Change in REM Sleep (%)	70.7 ± 216.3% (60)	81.9 ± 187.5% (111)	207.8 ± 346.6% (135)	0.558	
Stage III/IV Sleep (% TST)	10.9 ± 8.6% (60)	6.6 ± 6.2% (111)	6.9 ± 8.0% (135)	0.002	
Percent Change in Stage III/IV (%)	221.8 ± 495.7% (60)	235.7 ± 473.1% (111)	335.7 ± 567.3 (135)	0.868	
ODI (desaturations/hour)	2.2 ± 4.5 (60)	5.2 ± 9.5 (111)	12.1 ± 20.7 (135)	<0.001	
RDI Surgical Cure (%) [‡]	29.6% (425)	12.1% (338)	15.7% (178)	<0.001	
ODI Surgical Cure (%)	86.7% (60)	73.9% (111)	63.7% (135)	0.004	
RDI Surgical Success (%) [‡]	73.2% (425)	74.0% (338)	66.3% (178)	0.148	
ODI Surgical Success (%)	85.0% (60)	82.9% (111)	80.7% (135)	0.759	

^{*} Mean (or percent) ± standard deviation. The number of patients is shown in parentheses. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space – base of tongue), REM (rapid eye movement), RDI (respiratory disturbance index), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SpO₂ (pulse oxyhemoglobin saturation). † P-value assesses whether mean differences between the groups are statistically significant. ‡ Surgical cure defined an RDI (or ODI) < 5 events/hour post-MMA. Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA.

Table 3. Patient Characteristics Stratified by Percent Reduction in RDI Post-MMA*.

	Percent RDI Reduction						
Characteristics	≥ 75.0% (n=367; 39.0%)	50-74.9% (n=362; 38.5%)	25-49.9% (n=120; 12.8%)	< 24.9% (n=92; 9.8%)	P-value [†]	P-value [‡]	
Pre-MMA							
Age (years)	38.8 ± 14.3 (367)	38.4 ± 14.4 (362)	41.9 ± 14.4 (120)	46.8 ± 10.0 (92)	<0.001	<0.001	
Age > 50 years (%)	24.8% (367)	25.7% (269)	33.3% (120)	42.4% (92)	0.003	0.001	
Male (%)	80.4% (367)	72.1% (362)	80.0% (120)	90.2% (92)	0.001	0.031	
BMI (kg/m²)	28.3 ± 4.8 (363)	26.4 ± 3.0 (343)	27.9 ± 3.8 (116)	30.0 ± 5.4 (91)	0.048	0.030	
BMI > 30 kg/m² (%)	24.5% (363)	9.3% (343)	24.1% (116)	40.7% (91)	<0.001	0.002	
Cephalometrics							
SNA (degrees)	80.0 ± 3.1 (164)	80.3 ± 2.8 (75)	80.9 ± 3.2 (26)	80.0 ± 3.5 (48)	0.696#	1.000	
SNA < 80 degrees (%)	41.5% (164)	30.7% (75)	34.6% (26)	41.7% (48)	0.404	0.554	
SNB (degrees)	76.8 ± 4.0 (164)	77.4 ± 3.7 (74)	77.4 ± 3.7 (22)	77.4 ± 3.7 (46)	0.219#	0.818	
SNB < 80 degrees (%)	72.6% (164)	74.3% (74)	81.8% (22)	76.1% (46)	0.805	0.392	
PAS (mm)	5.1 ± 2.2 (164)	5.8 ± 2.1 (74)	5.4 ± 2.0 (22)	5.7 ± 2.4 (46)	0.076#	0.354	
Polysomnography							
RDI (events/hour)	47.6 ± 27.3 (367)	32.0 ± 17.7 (362)	33.8 ± 22.8 (120)	39.8 ± 25.0 (92)	<0.001	0.049	
SpO ₂ Nadir (%)	81.3 ± 10.6% (362)	85.9 ± 8.1% (357)	86.0 ± 7.7% (120)	78.4 ± 13.1% (90)	0.521	0.214	
ODI (desaturations/hour)	36.1 ± 28.3 (164)	23.6 ± 20.4 (74)	31.5 ± 32.1 (22)	35.2 ± 29.1 (46)	0.484	0.998	
Post-MMA							
BMI (kg/m²)	28.2 ± 4.9 (346)	26.3 ± 3.1 (320)	28.0 ± 3.9 (104)	30.0 ± 5.5 (89)	0.027	0.026	
Percent Change in BMI (%)	-0.8 ± 1.9 (345)	-0.7 ± 1.2 (315)	-0.6 ± 1.1 (104)	-0.1 ± 3.7 (89)	0.005#	0.008	
Cephalometrics							
SNA (degrees)	82.3 ± 3.0 (163)	80.9 ± 2.9 (74)	80.8 ± 3.0 (22)	80.1 ± 3.5 (46)	<0.001#	<0.001	
SNA < 80 degrees (%)	17.8% (163)	21.6% (74)	36.4% (22)	41.3% (46)	0.004	0.001	
Percent Change in SNA (%)	3.0 ± 3.8% (163)	0.9 ± 2.2% (74)	0.2 ± 1.1% (22)	0.2 ± 1.2% (46)	<0.001	<0.001	
SNB (degrees)	78.8 ± 3.2 (163)	78.1 ± 3.1 (74)	78.1 ± 3.3 (22)	77.5 ± 3.8 (46)	0.016#	0.103	
SNB < 80 degrees (%)	57.7% (163)	68.9% (74)	81.8% (22)	76.1% (46)	0.023	0.016	
Percent Change in SNB (%)	2.7 ± 3.6% (163)	1.0 ± 2.7% (74)	0.2 ± 1.1% (22)	0.2 ± 1.2% (46)	<0.001	<0.001	
PAS (mm)	9.1 ± 2.2 (163)	8.4 ± 2.4 (74)	7.0 ± 2.6 (22)	6.8 ± 2.6 (46)	<0.001#	<0.001	
Percent Change in PAS (%)	114.4 ±109.9% (163)	59.8 ± 62.8% (74)	32.9 ± 33.2% (22)	28.8 ± 53.6 (46)	<0.001	<0.001	
Change in PAS (mm)	4.1 ± 2.2 (163)	2.6 ± 2.0 (74)	1.6 ± 1.6 (22)	1.1 ± 1.9 (46)	<0.001#	<0.001	
Polysomnography							
RDI (events/hour)	6.2 ± 4.5 (367)	11.6 ± 6.6 (362)	20.8 ± 14.6 (120)	41.4 ± 24.2 (92)	<0.001	<0.001	
Percent Change in RDI (%)	-86.0 ± 7.0% (367)	-63.0 ± 7.1% (362)	-38.6 ± 7.0% (120)	12.8 ± 42.2% (92)	<0.001	<0.001	
SpO ₂ Nadir (%)	89.8 ± 3.9% (360)	90.4 ± 4.8% (358)	88.3 ± 6.4% (119)	80.9 ± 11.8% (90)	<0.001	<0.001	
ODI (desaturations/hour)	1.4 ± 2.3 (164)	5.3 ± 9.2 (74)	16.7 ± 17.7 (22)	29.5 ± 25.6 (46)	<0.001	<0.001	
RDI Surgical Cure (%)§	45.5% (367)	7.7% (334)	0.0% (120)	0.0% (92)	<0.001	<0.001	
ODI Surgical Cure (%)	92.7% (164)	73.0% (74)	22.7% (22)	19.6% (46)	<0.001	<0.001	
RDI Surgical Success (%)§	98.6% (367)	90.1% (362)	0.0% (120)	0.0% (92)	<0.001	<0.001	
ODI Surgical Success (%)	97.0% (164)	89.2% (66)	50.0% (22)	34.8% (46)	<0.001	<0.001	

^{*}Mean (or percent) ± standard deviation. The number of patients analyzed is shown in parentheses. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space – base of tongue), RDI (respiratory disturbance index), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SPO₃ (pulse oxyhemoglobin saturation). † P-value assesses whether differences between the groups are statistically significant (Weighted linear trend for continuous data; Fisher's exact 2-sided p-value used for categorical data). ‡ P-value assesses whether difference between ≥ 75% vs < 24.9% groups are statistically significant (Tukey [Levene's p > 0.05] or Games-Howell test for linear; Fisher's exact 2-sided for categorical data). § Surgical cure defined as an RDI (or ODI) < 5 events/hr post-MMA. Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA. # Levene's test was not statistically significant (p > 0.05) for the one-way ANOVA comparison, suggesting the variances are homogenous. || Welch's ANOVA p-value < 0.05.

Table 4. Predictors of Post-MMA OSA Surgical Cure*.

	Post MMA			Post MMA		
Predictor	Cure (RDI < 5) (n=195; 20.7%)	No Cure (RDI ≥ 5) (n=746; 79.3%)	P-value [‡]	Cure (ODI < 5) (n=220; 71.9%)	No Cure (ODI ≥ 5) (n=86; 28.1%)	P-value [‡]
Pre-MMA						
Age (years)	32.6 ± 13.8 (195)	41.8 ± 13.7 (746)	<0.001	45.8 ± 11.7 (220)	51.3 ± 10.9 (86)	<0.001
Age > 50 years (%)	11.8% (195)	32.2% (746)	<0.001	37.7% (220)	58.1% (86)	0.001
Male (%)	70.8% (195)	80.0% (746)	<0.001	92.7% (220)	98.8% (86)	0.048
BMI (kg/m²)	27.1 ± 4.6 (193)	27.9 ± 4.2 (720)	0.051	29.8 ± 5.7 (220)	32.5 ± 5.5 (86)	<0.001
BMI > 30 kg/m ² (%)	17.1% (193)	21.3% (720)	0.228	40.9% (220)	66.3% (86)	<0.001
Cephalometrics						
SNA (degrees)	79.8 ± 3.4 (66)	80.3 ± 3.0 (247)	0.376	80.2 ± 3.0 (219)	80.0 ± 3.5 (86)	0.635
SNA < 80 degrees (%)	42.4% (66)	37.2% (247)	0.477	37.3% (220)	43.0% (86)	0.364
SNB (degrees)	76.7 ± 4.1 (66)	77.3 ± 3.7 (247)	0.292	77.0 ± 3.9 (220)	77.4 ± 3.6 (86)	0.471
SNB < 80 degrees (%)	72.7% (66)	74.6% (247)	0.753	73.2% (220)	76.7% (86)	0.564
PAS (mm)	5.4 ± 2.1 (66)	5.4 ± 2.3 (240)	0.938	5.4 ± 2.3 (220)	5.3 ± 2.0 (86)	0.658
Polysomnography						
RDI (events/hour)	31.0 ± 25.2 (195)	41.2 ± 23.5 (746)	<0.001	52.9 ± 27.5 (220)	63.4 ± 24.1 (86)	0.001
RDI ≥ 20 events/hr (%)	55.4% (195)	82.7% (746)	<0.001	86.4% (220)	100.0% (86)	<0.001
SpO ₂ Nadir (%)	85.5 ± 9.1% (193)	82.9 ± 10.2% (736)	0.001	75.0 ± 10.0 (220)	69.4 ± 10.4 (86)	<0.001
ODI (desaturations/hour)	27.0 ± 27.7 (66)	34.2 ± 27.2 (240)	0.066	28.5 ± 26.8 (220)	43.1 ± 26.2 (86)	<0.001
ODI ≥ 20 events/hr (%)	43.9% (66)	59.2% (240)	0.035	49.1% (220)	73.3% (86)	<0.001
Post-MMA						
BMI (kg/m²)	27.1 ± 4.7 (181)	27.8 ± 4.3 (678)	0.055	29.7 ± 5.7 (219)	32.4 ± 5.5 (86)	<0.001
% Change in BMI (%)	-0.9 ± 2.5% (181)	-0.6 ± 1.7% (672)	0.108	-0.1 ± 0.7 (219)	-0.1 ± 0.3 (86)	0.401
Cephalometrics						
SNA (degrees)	82.0 ± 3.0% (66)	81.4 ± 3.2 (239)	0.202	82.0 ± 3.0 (219)	80.5 ± 34.7 (86)	0.001
SNA < 80 degrees (%)	19.7% (66)	24.7% (239)	0.513	18.7% (219)	36.0% (86)	0.002
Change in SNA (degrees)	2.1 ± 3.0 (66)	1.3 ± 2.3 (239)	0.030	1.8 ± 2.7 (219)	0.5 ± 1.5 (86)	<0.001
SNB (degrees)	78.4 ± 3.2 (66)	78.4 ± 3.4 (239)	0.913	78.5 ± 3.2 (219)	78.0 ± 3.6 (86)	0.230
SNB < 80 degrees (%)	62.1% (66)	65.7% (239)	0.662	62.1% (219)	72.1% (86)	0.111
Change in SNB (degrees)	1.7 ± 2.6 (66)	1.2 ± 2.3 (239)	0.138	1.5 ± 2.5% (219)	0.6 ± 1.8% (86)	<0.001
PAS (mm)	9.1 ± 2.1 (66)	8.3 ± 2.6 (239)	0.013	8.9 ± 2.3 (219)	7.3 ± 2.6 (86)	<0.001
Change in PAS (mm)	3.7 ± 2.3 (66)	2.9 ± 2.4 (239)	0.024	3.5 ± 2.4 (219)	2.0 ± 2.0 (86)	<0.001
Polysomnography						
RDI (events/hour)	2.6 ± 1.2 (195)	16.4 ± 15.2 (746)	<0.001	9.0 ± 7.5 (220)	42.4 ± 25.7 (86)	<0.001
% Change in RDI (%)	-86.1 ± 11.0% (195)	-55.0 ± 33.2% (746)	<0.001	-76.6 ± 30.9% (220)	-29.4 ± 37.6% (86)	<0.001
RDI < 20 events/hr (%)	100.0% (195)	88.8% (746)	<0.001	94.1% (220)	17.4% (86)	<0.001
SpO ₂ Nadir (%)	91.6 ± 3.3% (192)	88.3 ± 6.8% (735)	<0.001	87.2 ± 3.1% (220)	74.8 ± 9.8% (86)	<0.001
% Change in SpO ₂ Nadir (%)	8.4 ± 11.4% (192)	7.7 ± 12.2 (730)	0.479	18.4 ± 16.9% (220)	9.1 ± 15.4% (86)	<0.001
ODI (desaturations/hour)	0.4 ± 0.7 (66)	9.7 ± 17.0 (240)	<0.001	1.1 ± 1.4 (220)	24.6 ± 21.4 (86)	<0.001
% Change in ODI (%)	-95.8 ± 16.7% (66)	-43.8 ± 225.0% (240)	<0.001	-87.3 ± 34.1% (220)	27.7 ± 3.6% (86)	0.004
RDI Surgical Success (%)†	100.0% (195)	66.1% (746)	<0.001	92.7% (220)	16.3% (86)	<0.001
ODI Surgical Success (%)†	98.5% (66)	74.2% (240)	<0.001	93.6% (220)	43.0% (86)	<0.001

^{*} Surgical cure defined as an RDI (or ODI) < 5 events/hour post-MMA. Mean (or percent) ± standard deviation. The total number of patients is shown in parentheses. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space – base of tongue), RDI (respiratory disturbance index), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SpO₂ (pulse oxyhemoglobin saturation). † Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA. ‡ Equal variances not assumed for p-value calculation. Fisher's exact 2-sided p-value used for categorical data.

Table 5. Predictors of Post-MMA OSA Surgical Success*.

Bradistor	Post MMA			Post MMA			
Predictor	RDI Success (n=688; 73.1%)	No Success (n=253; 26.9%)	P-value [‡]	ODI Success (n=243; 79.4%)	No Success (n=63; 20.6%)	P-value [‡]	
Pre-MMA							
Age (years)	38.2 ± 14.3 (688)	44.2 ± 12.9 (253)	<0.001	46.3 ± 12.0 (243)	51.5 ± 9.9 (63)	0.001	
Age > 50 years (%)	24.4% (688)	37.5% (253)	<0.001	40.7% (243)	54.0% (243)	0.065	
Male (%)	75.7% (688)	84.6% (253)	0.003	93.0% (243)	100.0% (63)	0.028	
BMI (kg/m²)	27.2 ± 4.1 (668)	29.0 ± 4.6 (245)	<0.001	30.1 ± 5.8 (243)	32.3 ± 5.7 (63)	0.010	
BMI > 30 kg/m ² (%)	15.7% (668)	33.1% (245)	<0.001	44.4% (243)	61.9% (63)	0.016	
Cephalometrics							
SNA (degrees)	80.2 ± 3.0 (219)	80.0 ± 3.5 (94)	0.631	80.1 ± 3.0 (243)	80.0 ± 3.6 (63)	0.865	
SNA < 80 degrees (%)	37.9% (219)	39.4% (94)	0.801	38.3% (243)	41.3% (63)	0.666	
SNB (degrees)	77.1 ± 3.9 (219)	77.3 ± 3.6 (88)	0.637	77.0 ± 3.8 (243)	77.5 ± 3.8 (63)	0.353	
SNB < 80 degrees (%)	72.5% (218)	78.4% (88)	0.315	74.5% (243)	73.0% (63)	0.872	
PAS (mm)	5.3 ± 2.2 (218)	5.5 ± 2.2 (88)	0.461	5.4 ± 2.3 (243)	5.3 ± 2.0 (63)	0.696	
Polysomnography							
RDI (events/hour)	38.1 ± 23.5 (688)	41.8 ± 26.0 (253)	0.049	52.9 ± 27.5 (220)	63.4 ± 24.1 (86)	0.954	
RDI ≥ 20 events/hr (%)	76.5% (688)	78.7% (253)	0.541	86.4% (220)	100.0% (86)	0.056	
SpO ₂ Nadir (%)	84.1 ± 9.4% (688)	81.6 ± 11.3% (250)	0.002	75.0 ± 10.0 (220)	69.4 ± 10.4 (86)	0.253	
ODI (desaturations/hour)	30.8 ± 26.6 (218)	37.0 ± 28.9 (88)	0.084	28.5 ± 26.8 (220)	43.1 ± 26.2 (86)	0.991	
ODI ≥ 20 events/hr (%)	54.6% (218)	59.1% (88)	0.525	49.1% (220)	73.3% (86)	0.088	
Post-MMA							
BMI (kg/m²)	27.1 ± 4.2 (630)	29.0 ± 4.8 (229)	<0.001	30.0 ± 5.7 (242)	32.1 ± 5.7 (63)	0.010	
% Change in BMI (%)	-0.8 ± 1.7% (624)	-0.4 ± 2.4% (229)	0.031	-4.2 ± 2.1% (242)	-3.7 ± 9.5% (63)	0.774	
Cephalometrics							
SNA (degrees)	82.1 ± 2.8 (217)	80.1 ± 3.6 (88)	<0.001	81.9 ± 3.0 (242)	80.2 ± 3.5 (63)	0.001	
SNA < 80 degrees (%)	17.5% (217)	38.6% (88)	<0.001	19.4% (242)	39.7% (63)	0.001	
Change in SNA (degrees)	1.9 ± 2.7 (217)	0.3 ± 1.1 (88)	<0.001	1.8 ± 2.7 (242)	0.2 ± 0.9 (63)	<0.001	
SNB (degrees)	78.7 ± 3.1 (217)	77.7 ± 3.7 (88)	0.027	78.6 ± 3.2 (242)	77.7 ± 3.6 (63)	0.097	
SNB < 80 degrees (%)	60.8% (217)	75.0% (88)	0.024	62.4% (242)	74.6% (63)	0.077	
Change in SNB (degrees)	1.6 ± 2.5 (217)	0.4 ± 1.4 (88)	<0.001	1.6 ± 2.5 (242)	0.2 ± 1.2 (63)	<0.001	
PAS (mm)	9.0 ± 2.2 (217)	7.1 ± 2.6 (88)	<0.001	8.9 ± 2.3 (242)	6.8 ± 2.4 (63)	<0.001	
Change in PAS (mm)	3.7 ± 2.2 (217)	1.6 ± 2.0 (88)	<0.001	3.5 ± 2.3 (242)	1.5 ± 1.7 (63)	<0.001	
Polysomnography							
RDI (events/hour)	7.9 ± 4.7 (688)	29.0 ± 20.3 (253)	<0.001	11.4 ± 11.2 (243)	45.3 ± 28.2 (63)	<0.001	
% Change in RDI (%)	-75.2 ± 13.4% (688)	-23.9 ± 39.0% (253)	<0.001	-74.9 ± 30.1% (243)	-18.6 ± 37.9 (63)	<0.001	
RDI < 20 events/hr (%)	100.0% (688)	36.0% (253)	<0.001	85.6% (243)	22.2% (63)	<0.001	
SpO ₂ Nadir (%)	90.5 ± 3.7% (678)	84.9 ± 9.5% (249)	<0.001	86.3 ± 4.3% (243)	73.8 ± 11.1% (63)	<0.001	
% Change in SpO ₂ Nadir (%)	8.9 ± 12.5% (673)	4.9 ± 10.1% (249)	<0.001	18.8 ± 16.4% (243)	4.1 ± 13.9% (63)	<0.001	
ODI (desaturations/hour)	1.5 ± 2.3 (218)	23.0 ± 22.4 (88)	<0.001	2.4 ± 3.8 (243)	28.2 ± 24.4 (63)	<0.001	
% Change in ODI (%)	-90.0 ± 22.4% (218)	31.8 ± 359.1% (88)	0.002	-91.8 ± 11.6% (243)	87.0 ± 414.0% (63)	0.001	
RDI Surgical Success (%)†	28.3% (688)	0.0% (253)	<0.001	26.7% (243)	1.6% (63)	<0.001	
ODI Surgical Success (%)†	93.6% (688)	18.2% (88)	<0.001	84.8% (243)	22.2% (63)	<0.001	

^{*} Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA. Mean (or percent) ± standard deviation. The total number of patients is shown in parentheses. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space -base of tongue), RDI (respiratory disturbance index), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SpO₂ (pulse oxyhemoglobin saturation). † Surgical cure defined as an RDI (or ODI) < 5 events/hour post-MMA. ‡ Equal variances for linear data not assumed for p-value calculation. Fisher's exact 2-sided p-value used for categorical data.

Table 6. First 500 vs Last 510 MMA Characteristics*.

	First MMA N=500	Last MMA N=510	P-value				
Characteristics							
Age	44.8 ± 13.0 (500)	34.6 ± 13.7 (510)	<0.001				
Male Gender (%)	84.8% (500)	69.4% (510)	<0.001				
PreMMA BMI (kg/m²)	29.3 ± 5.1 (495)	25.8 ± 1.8 (481)	<0.001				
PostMMA BMI (kg/m²)	29.2 ± 5.1 (491)	25.5 ± 1.7 (423)	<0.001				
Percent Change BMI	-0.3 ± 2.2% (489)	-1.0 ± 2.3% (419)	<0.001				
PreMMA Polysomography							
RDI (events/hour)	48.0 ± 26.3 (498)	28.4 ± 16.8 (507)	<0.001				
RDI ≥ 30/hr (%)	69.9% (498)	36.7% (507)	<0.001				
SpO ₂ Nadir (%) [‡]	78.6 ± 11.2% (491)	88.8 ± 4.4% (495)	<0.001				
ODI (desaturations/hour)	32.6 ± 27.4 (306)	-	NA				
ODI ≥ 30/hr (%)	47.7% (306)	-	NA				
PostMMA Polysomography							
RDI (events/hour)	17.1 ± 18.3 (484)	9.9 ± 7.8 (458)	<0.001				
RDI ≥ 30/hr (%)	13.4% (484)	3.3% (458)	<0.001				
Percent Change RDI (%)	-59.0 ± 40.3% (484)	-64.0 ± 21.4% (457)	0.017				
SpO ₂ Nadir (%) [‡]	86.0 ± 7.3% (479)	92.1 ± 2.7% (449)	<0.001				
Percent Change SpO ₂ Nadir (%)	11.5 ± 15.5% (476)	3.9 ± 3.8 (449)	<0.001				
ODI (desaturations/hour)†	7.7 ± 15.5 (306)	-	NA				
ODI ≥ 30/hr (%)	8.5% (306)	-	NA				
Percent Change ODI (%)	-55.0 ± 2.0 (306)	-	NA				
PreMMA Cephalometrics†							
SNA (degrees)	80.2 ± 3.1 (321)	-	NA				
SNB (degrees)	77.1 ± 3.8 (306)	-	NA				
PAS (mm)	5.4 ± 2.2 (306)	-	NA				
PostMMA Cephalometrics [†]							
SNA (degrees)	81.5 ± 3.2 (305)	-	NA				
SNB (degrees)	78.4 ± 3.3 (305)	-	NA				
PAS (mm)	8.4 ± 2.5 (305)	-	NA				
RDI Surgical Cure (%)#	17.8% (484)	23.9% (457)	0.024				
ODI Surgical Cure (%)#,†	71.9% (306)	-	NA				
RDI Surgical Success (%)#	66.7% (484)	77.9% (457)	<0.001				
ODI Surgical Success (%)#,†	79.4%	-	NA				

^{*} Plus-minus values are mean (or percent) ± standard deviation. Values in parenthesis are the number of patients evaluated. Abbreviations: BMI (body mass index), MMA (maxillomandibular advancement), ODI (oxygen desaturation index), PAS (posterior airway space – base of tongue), PNS-P (distance of posterior nasal spine to the tip of the soft palate), RDI (respiratory disturbance index), REM (rapid eye movement), SNA (sella-nasion-Point A angle), SNB (sella-nasion-Point B angle), SpO₂ (pulse oxyhemoglobin saturation). † ODI and cephalometric data only available on a subset of MMA patients that were exclusively within the first 500 subjects studied. ‡ The SpO₂ nadir is the lowest oxyhemoglobin saturation measured during sleep. Only patients with both pre- and post-MMA SpO₂ nadir are included. # Surgical cure defined an RDI (or ODI) < 5 events/hour post-MMA. Surgical success defined as the percent of subjects with an RDI (or ODI) < 20/hour and a ≥ 50% reduction in the RDI (or ODI) post-MMA.

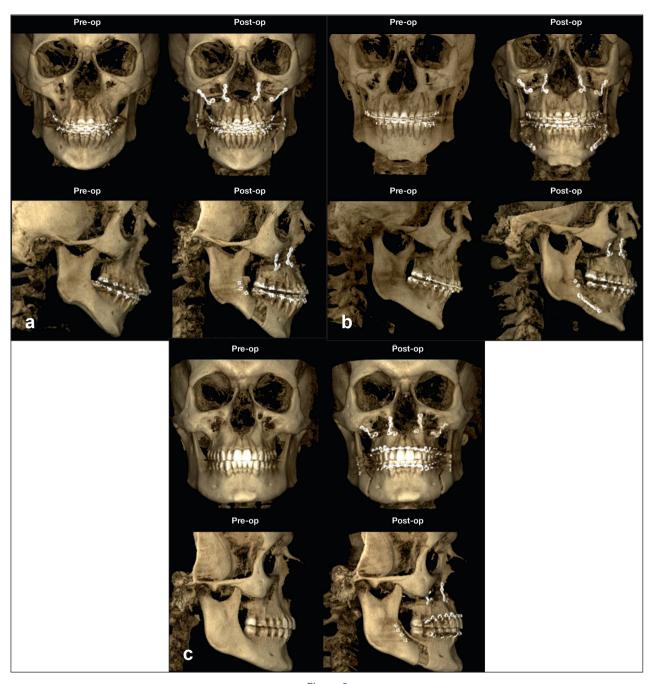


Figure 2

(a-c) Pre- and post-MMA imaging demonstrating different surgical plans but the main objective of maximizing the advancement within the physiologic limitation of the soft tissues is maintained.

ment option in properly selected patients. We have seen a trend of younger patients with less severe OSA increasingly opting for MMA as the treatment of choice. The improved surgical outcome with greater percentage reduction in RDI and surgical success/cure in patients with younger age, lower RDI and lower BMI are quite evident when comparing the first 500 patients versus the later 510 patients (Table 6). The differences in the patient population reflected the clinical insights gained over the years in the patient selection for MMA. The improved outcome is also likely related to the advances in surgical expertise and technique in the more recently treated patients.

The RDI was chosen as one of the primary outcome variables. This is due to the improved understanding of the impact of respiratory effort-related arousals (RERA) in causing sleep fragmentation affecting daytime function and quality of life^{1,6,7}. The results clearly demonstrate that MMA significantly reduces the RDI in the majority of the patients where the overall surgical success and surgical cure based on RDI were 73.1% and 20.7%, respectively. The RDI outcome compares favorably to the meta-analysis from 2016 with the rate of success and cure of 44 (64.7%) and 13 (19.1%), respectively³¹. This could be related to the small sample size in the meta-analysis with only 68 patients as well as the heterogeneity of the pooled data³¹. The comparison of the first 500 patients to the more recent 510 patients also showed that the outcome has improved in the rate of surgical success (77.9% versus 66.7%) and surgical cure (23.9% versus 17.8%). Again, the improved outcome could be related to better patient selection and surgical execution.

The maxillofacial anatomy as a risk factor in patients with OSA based on cephalometric data has been published extensively since the initial report by Guilleminault's group in 19869. The analysis on cephalometric measurements in this study showed that a greater increase in SNA, SNB and PAS postoperatively resulted in improved outcome (Table 5). This finding suggests that greater advancement distance results in greater OSA improvement. Therefore, it must be emphasized that the goal of MMA is not to "normalize" cephalometric measurements since many patients with "normal" measurements have OSA¹⁸ but rather, it is to maximize the advancement distance in optimizing airway expansion. This objective must

be adhered to because OSA remained even when advancement was maximized. It is recognized that lower OSA severity, neurocognitive symptoms and cardiovascular sequelae remain^{2,4,5}. Clearly, maximizing the advancement must be balanced with the alteration of facial esthetics. Therefore, the surgical plan must be a collaborative decision between the surgeon and the patient. A comprehensive discussion of prognostic factors and the impact of advancement on airway and esthetics is essential so an ideal surgical plan can be formulated based on a collaborative effort between the patient and the surgeon (Fig. 2).

5. Conclusions

MMA is an effective treatment to improve OSA, but the result can vary. Patient selection based on favorable prognostic factors and maximizing the advancement distance can improve outcomes.

Links of interest

The authors declare that they have no interest in the data published in this article.

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