

## Original Article

# Prediction of lip response to orthodontic treatment using a multivariable regression model

Amin Shirvani<sup>1</sup>, Saeid Sadeghian<sup>1</sup>, Safieh Abbasi<sup>2</sup>

<sup>1</sup>Torbinejad Dental Research Center and Department of Orthodontics, <sup>2</sup>Dental Students Research Center, School of Dentistry, Isfahan University of Medical Sciences, Isfahan, Iran

## ABSTRACT

**Background:** This was a retrospective cephalometric study to develop a more precise estimation of soft tissue changes related to underlying tooth movement than simple relationship between hard and soft tissues.

**Materials and Methods:** The lateral cephalograms of 61 adult patients undergoing orthodontic treatment (31 = premolar extraction, 31 = nonextraction) were obtained, scanned and digitized before and immediately after the end of treatment. Hard and soft tissues, angular and linear measures were calculated by Viewbox 4.0 software. The changes of the values were analyzed using paired *t*-test. The accuracy of predictions of soft tissue changes were compared with two methods: (1) Use of ratios of the means of soft tissue to hard tissue changes (Viewbox 4.0 Software), (2) use of stepwise multivariable regression analysis to create prediction equations for soft tissue changes at superior labial sulcus, labrale superius, stomion superius, inferior labial sulcus, labrale inferius, stomion inferius (all on a horizontal plane).

**Results:** Stepwise multiple regressions to predict lip movements showed strong relations for the upper lip (adjusted  $R^2 = 0.92$ ) and the lower lip (adjusted  $R^2 = 0.91$ ) in the extraction group. Regression analysis showed slightly weaker relations in the nonextraction group.

**Conclusion:** Within the limitation of this study, multiple regression technique was slightly more accurate than the ratio of mean prediction (Viewbox4.0 software) and appears to be useful in the prediction of soft tissue changes. As the variability of the predicted individual outcome seems to be relatively high, caution should be taken in predicting hard and soft tissue positional changes.

**Key Words:** Cephalometry, orthodontics, regression

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### Address for correspondence:

Dr. Safieh Abbasi, Dental Students Research Center, School of Dentistry, Isfahan University of Medical Sciences, Isfahan, Iran.  
E-mail: soofi\_abbasi@yahoo.com

## INTRODUCTION

Prediction is an important part of orthodontic treatments. For many years, cut-and-paste techniques of cephalometric acetate tracing have been used to visualize treatment objectives. Recently, computer programs allow the clinicians to analyze and predict the treatment changes.

Many investigators analyzed the facial form to relate the soft tissue changes to the underlying hard tissue. Since the 1980s, various systems including Orthographic, Dolphin, Quick Ceph, and Viewbox have been introduced in the literature.<sup>[1-5]</sup> Software

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developers have used linear relations for soft to hard tissue movements.<sup>[6]</sup> Most researchers have stated that the predictions based on the ratios between soft tissue movement to the movement of the underlying hard tissue cannot be made accurately.<sup>[1,4,7-9]</sup> So attempts to find the response of the soft tissue with the help of statistical methods such as multivariable regression analysis have been made. By using statistical methods, we can explain many factors that can affect lips movement after premolar extraction such as initial lip thickness.<sup>[10-15]</sup>

Most studies have used ratios to explain lip response to incisor retraction in premolar extraction cases. Ratios of maxillary incisor retraction to movements of labrale superius (Ls) reported to vary from 1.2:1 to 3.2:1, and for lower lip ranging from 0.4:1 to 1.8:1.<sup>[16-19]</sup> Talass *et al.*<sup>[20]</sup> showed that multiple regression model explained only 49% of the variability in upper lip retraction. Caplan and Shivapuja<sup>[17]</sup> stated they can explain 70.1% and 42% of the variations in lip response to premolar extraction therapy with regression models. Brock *et al.*<sup>[21]</sup> found that 60% of the variability in upper lip retraction at superior labial sulcus (Sls) could be explained using multivariable regression. Denis and Speidel<sup>[22]</sup> stated that the predictions of lower lip movements are twice as accurate as simple ratios.

As the data for soft tissue responses to four premolars extraction and specially nonextraction treatments are limited, and most studies were accomplished on the African-American, Caucasian or Southeast Asian populations, So the purpose of this study was first to identify soft tissue profile changes associated with the movement of the incisors by using linear ratios (Viewbox 4.0 software) and regression models in an Iranian adult population. Second, the comparison of accuracy of these two methods with actual values.

## MATERIALS AND METHODS

### Sample size and subjects

The sample consisted of 62 orthodontic patients (17 male, 45 female) with a mean age of 23.6 years that were selected from patients treated in orthodontic clinics at Afzal and Ghaedi university affiliated clinics of Isfahan University of Medical Sciences, Isfahan, Iran. Thirty-one treated with four first premolar extractions and the remaining with nonextraction treatment.

The selection criteria for those patients were as follows:

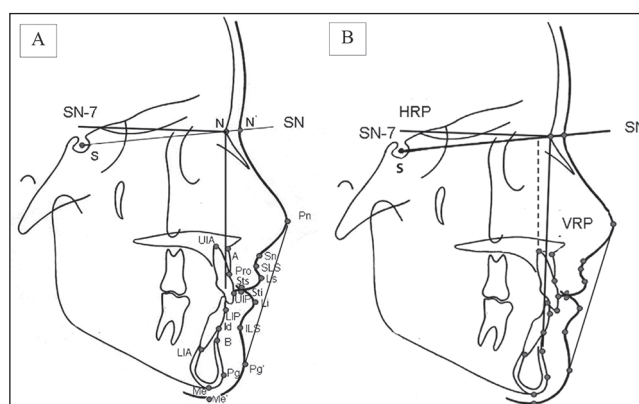
1. Belonging to an Iranian ethnic group with a minimum age of 18 years at the beginning of treatment to reduce growth effects.
2. Availability of a good quality lateral cephalograms.
3. No syndromes or craniofacial anomalies.
4. No orthogenetic or cosmetic surgery.

### Cephalometric procedure and measurements

The radiographs and a transparent millimeter ruler for calibration were digitally scanned using a flatbed scanner at 300 DPI resolutions (HP Scanjet G4050, China). Radiographic images were imported into the software (Viewbox imaging software, version 4.0, dhal, Kifissia, Greece) and were digitized by one of the investigators. Magnification was corrected prior to data analysis. Since in this software magnification tool did not guess the real size correctly, so for calibrating radiographs, the real length of the ruler in each radiograph was entered in the image resized plane and then the magnification was calculated with software. Twelve hard and 11 soft tissue landmarks were identified on each digitized cephalogram [Figure 1a]. Then 12 linear and angular measurements [Table 1] were computed according to the reference lines that shown in Figure 1b. These reference lines were used in previous studies.<sup>[23]</sup>

### Statistical analysis

*T*-test was used to evaluate the differences between the pre and post treatment in extraction and nonextraction groups. A  $P = 0.05$  was used as the minimal level of statistical significance. Stepwise multiple regression analysis was performed to detect whether the variables were helpful to predict soft tissue changes. Six dependent variables were



**Figure 1:** (a) Hard and soft tissue landmarks, (b) horizontal (HRP) and vertical (VRP) reference planes.

**Table 1: Linear and angular measurements from cephalograms**

Linear and angular measurements	Definition
Soft tissue thickness at Ls	Horizontal distance (mm) from the U1P to Ls
Soft tissue thickness at Sls	Horizontal distance from (mm) the A-point to Sls
Soft tissue thickness at Li	Horizontal distance (mm) from the L1P to Li
Soft tissue thickness at IIs	Horizontal distance from (mm) the B-point to IIs
Total facial height	Vertical distance (mm) from N' to Me'
ILG	Vertical distance (mm) from Sts to Sti
Ls to E-Line	Horizontal distance (mm) from E-Line to Ls
Sls to E-Line	Horizontal distance (mm) from E-Line to Sls
Li to E-Line	Horizontal distance (mm) from E-Line to Li
IIs to E-Line	Horizontal distance (mm) from E-Line to IIs
IMPA (°)	Lower incisor angulation to mandibular plane
Interincisal angle (°)	The Ant-posterior angle made by the intersection of the long axis of the maxillary central incisor with the mandibular central incisor

ILG: Interlabial gap; IMPA: Incisor mandibular plane angle; IIs: Inferior labial sulcus; Ls: Labrale superius; Li: Labrale inferius; Sls: Superior labial sulcus; U1p: Upper incisor point; L1p: Lower incisor point; N':Nasion; Me':Menton; E-line:line is formed by joining tip of the nose and soft tissue pogonion.

selected for regression analyses; post treatment location of Sls, Ls, stomion superius (Sts), inferior labial sulcus (IIs), labrale inferius (Li) and stomion inferius (Sti) (all in horizontal plane). All measurements were taken from pretreatment cephalograms. The independent variables were chosen according to the most significance and the influence on the outcome of treatment mentioned in the previous studies.<sup>[13,15,24]</sup>

Predictions were carried out firstly with Viewbox 4.0 software, and secondly using equations produced by stepwise regression analysis on a small sample ( $n = 5$ ) in both extraction and nonextraction groups. Finally, these data were compared with the actual values on the post treatment cephalograms. A series of 20 subjects was reassessed 2 months after initial digitization. Error of methods were calculated by Dahlberg formula:<sup>[25]</sup>

The error of the method<sup>2</sup> =  $\Sigma d^2/2n$ .

If the error of the method was not  $>0.5$ , the reliability in cephalometric tracing would be acceptable.

## RESULTS

Table 2 shows the means and standard deviations (SDs) of the horizontal changes of the hard and soft tissue landmarks and the mean differences between the pretreatment and post treatment stages. Significant level was set at 0.05.

Table 3 shows the means and SDs and Dahlberg errors of the horizontal changes for hard and soft tissue variables (in mm). Significant level was set at 0.05. The error of the method was not  $>0.5$ , so

reliability of cephalometric tracing was acceptable. Significant differences in hard tissue variables were noted between the stages for the variables involving the upper and lower incisors. No significant differences were found in the skeletal variables which show all the subjects were adult patients. In the soft tissue variables, significant differences were detected between the stages for the variables involving upper and lower lips.

### Extraction group

For the hard tissue variables, significant differences were detected in the horizontal changes of Pro, Id, L1p, and U1p. Significant differences in the soft tissue variables were found in the horizontal changes of the upper and lower lips landmarks. The upper and lower lips at Ls and Li retracted by an average  $1.86 \pm 3.19$  mm and  $1.5 \pm 2.42$  mm, giving a ratio of 2:1 and 1.8:1, respectively. The mean changes for Ls and Li to E-Line were  $1.12 \pm 4.35$  mm and  $2.49 \pm 1.85$  mm, respectively, that showed retraction movements of lips.

Stepwise multiple regression analysis was performed to predict the soft tissue profile changes. The results of the stepwise regression analysis [Table 4] suggested that the final position of Sti (adjusted  $R^2 = 0.96$ ), Li and Sls (adjusted  $R^2 = 0.95$ ) had strong relations with underlying structures. For the horizontal changes of upper and lower lips, incisors' facial point differences had the most effect. The other soft and hard tissue landmarks appeared to be more variable. The results indicated that the upper and lower lips could be strongly predicted, and there were a small difference between them.

**Table 2: Comparison of horizontal hard and soft tissue landmark movements (mm) of extraction and nonextraction group**

Landmarks	Before				After				Difference				P	
	Extraction		Nonextraction		Extraction		Nonextraction		Extraction		Nonextraction		Extraction	Nonextraction
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD		
Me-H	21.08	11.35	16.65	9.10	21.72	8.55	16.18	8.41	-0.64	10.63	0.47	5.92	0.497	0.161
U1P-H	-5.85	7.02	-3.97	5.50	-1.99	4.97	-3.67	4.78	3.85	4.68	0.29	3.71	0.01	0.03
U1A-H	6.70	4.53	6.34	4.29	5.87	3.95	7.15	4.22	-0.82	4.07	0.80	3.11	0.285	0.16
Pro-H	-2.59	4.66	-1.53	4.73	-0.38	4.43	-1.04	4.32	2.21	2.25	0.49	3.37	0.01	0.04
Pg-H	12.30	6.93	8.85	8.53	12.15	7.80	10.14	7.96	-0.15	5.30	1.28	5.61	0.876	0.212
L1p-H	0.31	5.57	1.12	6.12	3.12	5.35	1.47	5.62	2.81	2.88	0.34	3.36	0.01	0.03
L1A-H	14.67	7.96	13.16	7.37	13.52	6.11	14.31	6.86	-1.15	7.77	1.15	4.23	0.43	0.139
Id-H	3.03	4.94	3.59	6.40	4.79	5.60	4.16	6.07	1.76	3.10	0.57	3.52	0.005	0.374
B-point-H	9.19	5.74	7.63	7.31	9.87	6.16	8.80	6.75	0.68	5.07	1.17	4.12	0.475	0.124
A-point-H	2.96	3.29	2.47	3.49	3.38	3.71	3.30	3.95	0.42	2.79	0.83	2.87	0.425	0.116
Sts-H	-7.55	7.67	-6.10	5.12	-3.83	4.73	-5.54	4.81	3.72	5.50	0.56	3.76	0.01	0.03
Sti-H	-6.77	8.04	-5.01	5.66	-2.80	5.27	-4.47	5.09	3.97	5.93	0.54	4.08	0.01	0.05
Sn-H	-17.09	12.03	-16.01	4.57	-17.25	3.73	-15.59	4.12	0.16	10.80	0.41	3.28	0.168	0.482
Sls-H	-15.70	11.03	-14.24	4.68	-12.09	4.01	-13.65	4.19	3.61	9.54	0.58	3.10	0.01	0.04
Pg'-H	-0.82	5.93	-4.22	8.56	-0.10	7.00	-4.21	7.24	0.72	3.74	0.01	5.11	0.305	0.351
lIs-H	-2.34	5.38	-3.55	7.34	-0.38	5.68	-2.82	6.59	1.95	3.68	0.72	4.43	0.008	0.04
Li-H	-14.98	11.78	-13.50	6.33	-10.66	5.60	-12.90	5.57	4.32	9.42	0.60	3.96	0.02	0.06
Ls-H	-14.38	11.85	-17.59	5.56	-10.0	4.98	-17.06	4.85	4.38	10.19	0.52	3.70	0.02	0.05
N'-H	-6.03	7.53	-4.83	1.40	-6.64	1.51	-4.81	1.69	0.61	7.23	0.02	2.00	0.31	0.736
Me'-H	17.56	12.81	11.88	10.09	17.06	9.43	11.80	9.19	0.5	13.60	0.08	5.85	0.556	0.297

SD: Standard deviation; U1p: Upper incisor Point; U1A: Upper incisor Apex; Pro: Prosthion; Pg: Pogonion; L1p: Lower incisor point; L1A: Lower incisor apex; Id: Infradental; Sn: Subnasal; lIs: Inferior labial sulcus; Pg': Soft pogonion; Ls: Labrale superius; Li: Labrale inferius; Sls: Superior labial sulcus; N': Soft Nasion; Me': Soft menton.

**Table 3: Dental and soft tissue changes due to extraction and nonextraction treatments**

Variables	Before				After				Difference				P		Dahlberg error
	Extraction		Nonextraction		Extraction		Nonextraction		Extraction		Nonextraction		Extraction	Non-extraction-	
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD			
Interincisal angle	118.1	10.26	131.51	11.98	136.86	11.68	122.70	9.59	18.76	10.74	-8.81	14.84	0.01	0.002	0.5
IMPA	96.38	9.32	92.48	7.89	86.86	10.21	98.32	7.51	-9.52	6.53	5.83	8.09	0.001	0.01	0.31
lIs to E-line	6.32	4.62	7.15	2.46	6.39	2.03	6.94	2.77	0.07	3.70	-0.21	2.27	0.913	0.606	0.4
Ls to E-line	3.26	5.12	3.99	2.20	4.38	2.70	3.92	2.45	1.12	4.35	-0.07	2.51	0.882	0.876	0.4
Li to E-line	-1.17	3.86	1.61	2.75	1.31	3.73	1.36	3.26	2.49	1.85	-0.24	3.52	0.01	0.702	0.25
Sls to E-line	11.07	7.56	11.10	2.39	12.11	2.14	11.11	2.66	1.04	7.14	0.00	2.04	0.436	0.993	0.39
Interlabial gap	3.67	2.31	3.54	2.10	2.58	1.41	3.17	1.81	-1.09	2.90	-0.37	2.66	0.052	0.445	0.31
Anterior face height	143.6	77.50	128.60	16.02	143.1	12.08	130.69	19.51	-0.5	76.54	2.08	10.73	0.251	0.288	0.42
Lip thick at Sls	20.49	11.76	18.18	2.95	17.70	3.29	18.38	3.29	-2.79	11.37	0.20	2.69	0.197	0.682	0.40
Lip thick at Ls	14.61	6.04	14.31	2.57	14.19	2.40	14.00	2.41	-0.41	6.22	-0.30	1.97	0.721	0.394	0.39
Lip thick at Li	14.65	8.69	13.51	2.45	12.88	2.45	12.65	2.06	-1.76	8.38	-0.86	2.01	0.266	0.023	0.5
Lip thick at lIs	12.46	6.24	11.49	1.73	11.04	1.67	11.96	1.82	-1.41	6.18	0.46	1.50	0.229	0.094	0.46

SD: Standard deviation; IMPA: Incisor mandibular plane angle; lIs: Inferior labial sulcus; Ls: Labrale superius; Li: Labrale inferius; Sls: Superior labial sulcus; E-line: line is formed by joining tip of the nose and soft tissue pogonion.

When the regression models were tested on the 5 post treatment patients, some variations were found between the actual values compared to the predicted values of multivariable models and Viewbox4.0 software. The differences between the actual and predicted mean values of variables are shown in Table 5.

### Nonextraction group

We found the ratio of 2.8:1 and 1.9:1 between upper and lower incisors and lips, respectively. This was with significant changes in inter incisal angle and incisor mandibular plane angle. The mean changes for Ls and Li to E-Line

**Table 4: Stepwise multivariable regression model for upper and lower lip in extraction and nonextraction groups**

Dependent variables	R	Adjusted R <sup>2</sup>	Constant	Prediction equation			
				1 <sup>st</sup>	2 <sup>nd</sup>	3 <sup>rd</sup>	4 <sup>th</sup>
Extraction group							
Sls	0.97	0.95	-1.72	0.94 (Sls-x)	0.29 (ILG)	0.29 (ILG)	
Ls	0.96	0.91	-6.51	0.61 (Ls-x)	0.56 (Pro-x)	0.56 (Pro-x)	
Sts	0.96	0.91	0.04	0.93 (Sts-x)			
Ils	0.92	0.83	0.25	1.05 (Ils-x)			
Li	0.97	0.95	-5.67	0.58 (Li-x)	0.48 (L1p-x)	0.48 (L1p-x)	-0.94 (ld-x)
Sti	0.98	0.96	-3.76	0.02 (Sts-x)	1.88 (L1p-x)	1.88 (L1p-x)	
Non-extraction group							
Sls	0.91	0.81	-3.94	0.67 (Sls-x)	-0.65 (U1dif-x)	0.37 (Pro-x)	
Ls	0.92	0.83	-4.57	0.70 (Ls-x)	-0.78 (U1dif-x)	0.47 (Pro-x)	
Sts	0.94	0.88	-2.65	0.47 (Sti-x)	-0.87 (U1dif-x)	0.60 (Pro-x)	
Ils	0.94	0.88	0.16	0.86 (Ils-x)	-0.79 (L1dif-x)		
Li	0.95	0.91	-0.31	0.97 (Li-x)	-0.89 (L1dif-x)		
Sti	0.97	0.93	-2.69	0.51 (L1p-x)	-1.03 (L1dif-x)	0.44 (Sts-x)	

Prediction equations: Y (dependent variable) = constant + (1<sup>st</sup>) + (2<sup>nd</sup>) + (3<sup>rd</sup>) + (4<sup>th</sup>), for all values given;  $P < 0.05$ ; Sls: Superior labial sulcus; Ls: Labrale superius; Sts: Stomion superius; Ils: Inferior labial sulcus; Li: Labrale inferius; Sti: Stomion inferius; ILG: Interlabial Gap; Pro: Prosthion; L1p: lower incisor Point; U1dif: Upper incisors movements after treatment; L1dif: Lower incisors movements after treatment.

**Table 5: Result of testing the prediction equations and Viewbox software on 5 new subjects: Differences of means between the result derived from the prediction equations and Viewbox and actual results on the posttreatment cephalograms**

Group	Means (mm)					
	Sts-H	Sti-H	Sls-H	Ils-H	Li-H	Ls-H
Extraction						
Viewbox 4.0	-6.62	-6.34	-14.14	0.32	-14.52	-18.14
Regression model	-11.09	-4.14	-13.42	-0.08	-12.29	-17.46
Real values	-8.97	-3.90	-12.98	-0.76	-12.38	-16.52
Real values – Regression model	2.12	0.24	0.44	-0.68	-0.09	0.94
Real values – Viewbox 4.0	-2.35	2.44	1.24	-1.08	2.14	1.62
Nonextraction						
Viewbox 4.0	-8.48	-7.38	-18.82	-5.48	-16.0	-20.24
Regression model	-9.83	-9.20	-17.90	-6.85	-17.67	-20.93
Real values	-9.92	-10.08	-16.80	-8.62	-17.64	-22.32
Real values – Regression model	-0.09	-0.88	1.10	-1.77	0.03	-1.39
Real values – Viewbox 4.0	-1.35	-2.70	2.02	-3.14	-1.64	-2.08

Sls: Superior labial sulcus; Ls: Labrale superius; Sts: Stomion superius; Ils: Inferior labial sulcus; Li: Labrale inferius; Sti: Stomion inferius

were  $-0.07 \pm 2.51$  mm and  $-0.25 \pm 3.52$  mm, respectively.

The results of the regression analysis [Table 4] suggested that the final position of Sti (adjusted  $R^2 = 0.93$ ) and Li (adjusted  $R^2 = 0.91$ ), had a strong association with associated landmarks. Like extraction group, incisors' facial point differences had the most effect on the horizontal changes of upper and lower lips.

## DISCUSSION

The known ratios of soft to hard tissue movements are different in each individual, and only mean values are currently being used in the prediction programs. So we can suppose that in more instances, inaccuracy of prediction is inevitable.

Behrents<sup>[26]</sup> stated that facial growth could be detected well up to adulthood. Bishara *et al.*<sup>[27]</sup> also suggested that in females the most significant soft tissue changes occurs between the age of 10 and 15. Subtelny<sup>[28]</sup> and Vig and Cohen<sup>[29]</sup> noted that soft tissue changes will become complete after menarche. So we reduced the effects of growth and ethnic by selecting an Iranian adult patient especially women with a minimum age of 18 at the start of treatment which made the sample size relatively small.

For a closer look at the effect of extraction and nonextraction treatments on the soft tissue response, the following discussion is presented in two separate sections.

### Extraction group

The need for extraction depends on many factors such as crowding, incisor proclination, and dentofacial esthetics.<sup>[30]</sup> In our study, the ratio between Ls and upper incisors was 2:1. Comparing the studies is difficult due to the methodological differences. For the lower lip, this ratio was 1.8:1 that falls between the ratios reported from studies with extraction, which ranged from 1.9:1 to 0.4:1.<sup>[16,17]</sup>

Like other studies, the lip thickness at Ls and Li reduced with incisors retraction.<sup>[13]</sup> This contradicts with the reports of Hasstedt<sup>[31]</sup> and Ricketts,<sup>[32]</sup> that upper lip thickness increased with incisor retraction. It is clear when lip posture is relaxed, lip thickness does not increase during the retraction of incisors.

Six dependent variables were selected to perform stepwise multivariable regression analysis. More points might be helpful to explain some of the variability in lip responses. The correlation between Sti (adjusted  $R^2 = 0.96$ ), Li and SIs (adjusted  $R^2 = 0.95$ ) were high and similar to previous studies.<sup>[24,33,34]</sup> Post treatment position of Sti was also found to be affected by pretreatment Sts. This point may indicate the importance of lip competency and taking the pretreatment cephalograms in a reproducible relaxed lip posture.<sup>[15]</sup>

The multiple regression prediction equations for upper lip retraction explained 92% of the variability. Our equations were simpler than previously reported.<sup>[15,35]</sup> Previous multiple regression prediction equation has explained from 42% to 56% of the variability in the upper lip response to incisor retraction.<sup>[17,21]</sup> The stronger association could have been due to the strict selection criteria and reducing confounding factors. Brock *et al.*<sup>[21]</sup> explained 60% variability in SIs retraction. Talass *et al.*<sup>[20]</sup> and Ramos *et al.*<sup>[36]</sup> found the horizontal movement of prosthion and pretreatment upper lip thickness were important predictor factors.

The multivariable prediction equation for lower lip retraction was able to explain 91% of the variability. Retraction of the lower lip was similar to the upper lip, which disagrees with others reporting that upper lip retraction is less predictable because of the complex anatomy of the upper lip.<sup>[13,20]</sup> However, in this study, difference between upper and lower lips were small and it can be negligible. Veltkamp *et al.*<sup>[14]</sup> showed that using simple ratios only approximately 50% of the variation in soft tissue response can be explained. According them, multivariable regression model increased predictive accuracy by up to 40%. Our findings agree with this and seem to be able to predict soft tissue response after premolar retraction with a higher degree of accuracy than using Viewbox 4.0 software.

Because it was not possible to test the prediction equations on a sufficiently large group, we used 5 patients out of the original study. However, the sample

size was small, but differences between means were relatively small, and it was possible to predict the soft tissue response within clinically useful ranges [Table 3]. Our findings were similar to Kneafsey *et al.* study.<sup>[15]</sup>

### Nonextraction group

In this study, we desired to assess the soft tissue response in extraction and nonextraction patients. Except dental measurements, both groups were similar in hard and soft tissue measurements. Forward tipping of the incisor was noted after treatment.

Similar to extraction group, regression models were more accurate than Viewbox 4.0 prediction. The multiple regression equations for upper and lower lips explained 84% and 90% of the variability, respectively, that were less than what we got in extraction group that can be explained by more personal variability in soft tissue response. Until now, there is no adequate data on nonextraction treatments.

Soft tissue activities during imaging may confound soft tissue measurements. In our study, estimating the effect of this was difficult, but we attempted to have the lips relaxed during the actual exposure as mentioned before.<sup>[37,38]</sup> Previous studies of lip profile changes following orthodontic treatments had used Caucasian,<sup>[39,40]</sup> African American,<sup>[16]</sup> and Asian<sup>[18,41]</sup> races. Our study is unique as no data exists for the Iranian population, and it may provide useful information for clinician who is treating Iranians with similar malocclusions.

Prediction using programs such as Viewbox has two obvious downsides. First, the ratios that have been used are based on the average data collected from different samples. Second, using this method, it was supposed that there is a linear and fixed relation between soft and hard tissues. These factors could result in less accuracy in comparison to regression models. Until now, there are few studies on the accuracy of the cephalometric prediction using prediction programs.<sup>[5]</sup> Our study had many limitations such as nonhomogeneous studied population and not controlling the effect of treatment variables such as the method of space closure. Since all radiographic images in this research did not have enough quality, we try to select better contrast images. However, it was possible to have some cases with not enough detectable borders of soft tissues. Future studies could be carried out with a larger sample size that has more uniform pretreatment characteristic and more controlled treatment variables.

This study showed that the soft tissue response could be predicted more accurately by multivariable regression models and in the future, these models might benefit to improve software programs.

## CONCLUSION

Within the limitation of this study, the following conclusions can be made from this study:

1. The relationships between the lip changes and hard tissue changes are strong in both extraction and nonextraction groups, horizontally.
2. Incisors movements showed the strongest relationship to upper and lower lips.
3. With the regression models, it was possible to explain 92% and 91% of the variations in soft tissue response for upper and lower lips retraction, respectively. These models produced slightly weaker prediction in nonextraction group.
4. Predicting the soft tissue changes could be accomplished by using multivariable regression analysis. This method was more accurate than using simple ratios (Viewbox 4.0 software).
5. Prediction of soft tissue changes following orthodontic treatments showed more variability in the nonextraction group.
6. The variability of the predicted hard and soft tissue individual outcome seems to be relatively high.

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## Conflicts of interest

The authors of this manuscript declare that they have no conflicts of interest, real or perceived, financial or non-financial in this article.

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