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The Cervical Vertebral Maturation (CVM) Method for the Assessment of Optimal Treatment Timing in Dentofacial Orthopedics

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The present study introduces a further modified version of the Cervical Vertebral Maturation (CVM) method for the detection of the peak in mandibular growth, based on the analysis of the second through fourth cervical vertebrae in a single cephalogram. The morphology of the bodies of the second (C2 –odontoid process), third (C3), and fourth (C4) cervical vertebrae were analyzed in 6 consecutive cephalometric observations (T₁ through T₆) of 30 orthodontically untreated subjects. Observations for each subject consisted of two consecutive cephalograms comprising the interval of maximum mandibular growth (as assessed by means of the maximum increment in total mandibular length, Condylion –Gnathion: Co-Gn), together with two earlier consecutive cephalograms and two later consecutive cephalograms. The analysis consisted of both visual and cephalometric appraisals of morphological characteristics of the three cervical vertebrae. The construction of this new modified version of the CVM method was based on the results of both ANOVA for repeated measures with post hoc Scheffé's test ($P < 0.05$) and discriminant analysis. The new clinically improved CVM method is comprised of six maturational stages (cervical stage 1 through cervical stage 6, ie, CS1 through CS6). CS1 and CS2 are prepeak stages; the peak in mandibular growth occurs between CS3 and CS4. CS6 is recorded at least 2 years after the peak. The use of the CVM method enables the clinician to identify optimal timing for the treatment of a series of dentoskeletal disharmonies in all three planes of space.
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In the organization, differentiation, development, and growth of any somatic structure, time plays a crucial role in determining the final morphological and dimensional result. In orthodontics and dentofacial orthopedics, it is becoming increasingly evident that the timing of the treatment onset may be as critical as the selection of the specific treatment protocol, as will be discussed below. By beginning a protocol at the individual patient's optimal maturational stage, the most favorable response with the least potential morbidity can be anticipated.

The issue of optimal timing for dentofacial orthopedics is linked intimately to the identification of periods of acceler-

ated growth that can contribute significantly to the correction of skeletal imbalances in the individual patient. Cephalometric investigations on longitudinal samples have identified a pubertal spurt in mandibular growth that is characterized by wide individual variations in onset, duration, and rate.¹⁻⁶ Individual skeletal maturity can be assessed by means of several biologic indicators: increase in body height¹⁻³; skeletal maturation of the hand and wrist⁷⁻¹⁰; dental development and eruption^{8,11,12}; menarche or voice changes^{9,13,14}; and cervical vertebral maturation.^{15,16} The biologic indicators of skeletal maturity refer mainly to somatic changes at puberty, thus emphasizing the strict interactions between the development of the craniofacial region and the modifications in other body regions.

The reliability and efficiency of a biologic indicator of skeletal maturity can be evaluated with respect to several fundamental requisites.¹⁷ An "ideal" biologic indicator of individual mandibular skeletal maturity should be characterized by at least five features.

1. Efficacy in detecting the peak in mandibular growth. The method should present with a definite stage or

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phase that coincides with the peak in mandibular growth in the majority of subjects.

2. No need for additional x-ray exposure.
3. Ease in recording.
4. Consistency in the interpretation of the data. The inter-examiner error in the appraisal of the defined stages or phases should be as low as possible.
5. Usefulness for the anticipation of the occurrence of the peak. The method should present with a definable stage or phase that occurs before the peak in mandibular growth in the majority of subjects.

The main features of the Cervical Vertebral Maturation (CVM) method as described previously by Franchi and co-workers¹⁸ included:

1. In nearly 95% of North American subjects, a growth interval in CVM coincides with the pubertal peak in both mandibular growth and body height.
2. The cervical vertebrae are available on the lateral cephalogram that is used routinely for orthodontic diagnosis and treatment planning.
3. The appraisal of the shape of the cervical vertebrae is straightforward.
4. The reproducibility of classifying CVM stages is high (>98% by trained examiners).
5. The method is useful for the anticipation of the pubertal peak in mandibular growth.

A subsequent study by our group¹⁹ provided a few improvements of the original CVM analysis to make the method easier and applicable to the vast majority of patients:

1. A more limited number of vertebral bodies was used to perform the staging (as suggested by Hassel and Farman²⁰). In particular, the method included only those cervical vertebrae (C2, C3, and C4) that can be visualized when a protective radiation collar is worn by the patient.
2. Definitions of stages were not based on a comparative assessment of between-stage changes, so that stages can be identified easily on a single cephalogram.

A series of investigations performed in different parts of the world have confirmed the validity of the CVM method, mostly by comparing it with the hand and wrist method. Pancherz and Szyska found that the cervical vertebral maturation method has level of reliability comparable to the hand and wrist method.²¹ By replacing the hand-wrist method with the CVM method, an additional radiograph can be avoided, thus reducing the patient's total radiation dose. Grave and Townsend also have confirmed the validity of the CVM method in Australian aborigines.²²

The aim of the present article is to present a further modified and refined version of the CVM method and its validity for the appraisal of mandibular skeletal maturity in the individual patient in light of the findings of recent studies in which the CVM method has been used to assess optimal timing for the treatment of malocclusions in the transverse, sagittal, and vertical planes of space.

Subjects and Methods

The total sample ($n = 706$) that comprises the cephalometric files of the University of Michigan Elementary and Secondary School Growth Study was evaluated.²³ Due to the longitudinal nature and aim of the present investigation, subjects with less than six consecutive annual cephalometric observations ($n = 492$) were excluded from the study. Total mandibular length (Co-Gn) was measured on the longitudinal sets of lateral cephalograms for each of the 214 remaining subjects at yearly intervals. The lateral cephalograms were analyzed by means of a digitizing tablet (Numonics, Lansdale, PA) and digitizing software (Viewbox, version 3.0, D. Halazonetis, Athens, Greece). The maximum increase in Co-Gn between two consecutive annual cephalograms was used to define the peak in mandibular growth at puberty in the individual subjects. Two consecutive cephalograms comprising the interval of maximum mandibular growth, together with two earlier consecutive cephalograms and two later consecutive cephalograms, had to be available for each subject and were included in the study. This limited the investigation to 30 subjects (18 males, 12 females).

The morphology of the bodies of the second (C2 –odontoid process), third (C3), and fourth (C4) cervical vertebrae were analyzed in the six consecutive annual observations (T_1 through T_6). The analysis consisted of both visual and cephalometric appraisals of morphological characteristics of the cervical vertebrae.

Visual analysis. The morphology of the three cervical vertebrae (C2, C3, C4) on the six consecutive cephalograms (T_1 through T_6) was evaluated by visual inspection. Two investigators (LF and TB) performed the appraisal independently. The percentage of interexaminer agreement was 96.7%. Two sets of variables were analyzed:

1. Presence or absence of a concavity at the lower border of the body of C2, C3, and C4; and
2. Shape of the body of C3 and C4. Four basic shapes were considered:
 - trapezoid (the superior border is tapered from posterior to anterior);
 - rectangular horizontal (the heights of the posterior and anterior borders are equal; the superior and inferior borders are longer than the anterior and posterior borders);
 - squared (the posterior, superior, anterior, and inferior borders are equal); and
 - rectangular vertical (the posterior and anterior borders are longer than the superior and inferior borders).

Cephalometric analysis. On the lateral cephalograms, the following points for the description of the morphologic characteristics of the cervical vertebral bodies were traced and digitized (Fig 1):

- C2p, C2 m, C2a: the most posterior, the deepest, and the most anterior points on the lower border of the body of C2.

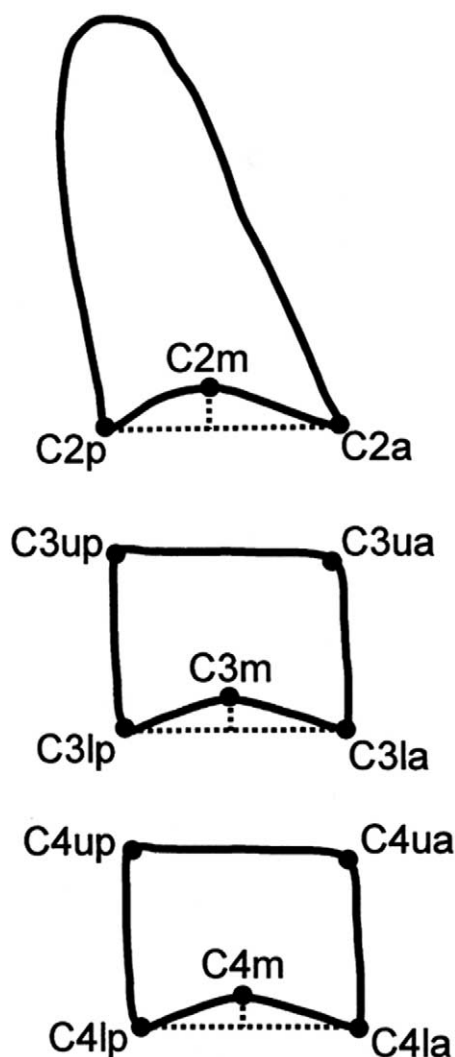


Figure 1 Cephalometric landmarks for the quantitative analysis of the morphologic characteristics of the vertebral bodies of C2, C3, and C4.

C3up, C3ua: the most superior points of the posterior and anterior borders of the body of C3.

C3lp, C3 m, C3la: the most posterior, the deepest, and the most anterior points on the lower border of the body of C3.

C4up, C4ua: the most superior points of the posterior and anterior borders of the body of C4.

C4lp, C4 m, C4la: the most posterior, the deepest, and the most anterior points on the lower border of the body of C4.

For the location of landmarks, the indications described by Helsing were adopted partially.²⁴ With the aid of these landmarks, the following measurements were performed:

C2Conc: a measure of the concavity depth at the lower border of C2 (distance from the line connecting C2p and C2a to the deepest point on the lower border of the vertebra, C2 m).

C3Conc: a measure of the concavity depth at the lower

border of C3 (distance from the line connecting C3lp and C3la to the deepest point on the lower border of the vertebra, C3 m).

C4Conc: a measure of the concavity depth at the lower border of C4 (distance from the line connecting C4lp and C4la to the deepest point on the lower border of the vertebra, C4 m).

C3BAR: ratio between the length of the base (distance C3lp-C3la) and the anterior height (distance C3ua-C3la) of the body of C3.

C3PAR: ratio between the posterior (distance C3up-C3lp) and anterior (distance C3ua-C3la) heights of the body of C3.

C4BAR: ratio between the length of the base (distance C4lp-C4la) and the anterior height (distance C4ua-C4la) of the body of C4.

C4PAR: ratio between the posterior (distance C4up-C4lp) and anterior (distance C4ua-C4la) heights of the body of C4.

Statistical analysis. The significance of the prevalence rates for the morphologic characteristics of the cervical vertebrae was evaluated at each observation time by means of the chi-squared test with Yates' correction ($P < 0.05$). Descriptive statistics were obtained for total mandibular length and for vertebral cephalometric measures at each of the six consecutive observations (T_1 through T_6). The differences between the mean values for all the computed variables at the six consecutive stages were tested for significance by means of ANOVA for repeated measurements with post hoc Scheffé's test ($P < 0.05$).

The cephalometric measurements of the bodies of the cervical vertebrae at each interval between consecutive cephalograms were analyzed by means of a multivariate statistical approach, discriminant analysis, to identify those vertebral morphologic variables mostly accounting for the differences between two consecutive observations. A stepwise variable selection (forward selection procedure) was performed with the goal of obtaining a model with the smallest set of significant cephalometric variables (F to enter and to remove = 4). Finally, the classifying power of selected cephalometric variables was tested. All statistical computations were performed by means of computer software (SPSS for Windows, version 10.0.0, SPSS, Inc., Chicago, IL).

Results

The findings of the visual analysis of the morphologic characteristics of cervical vertebrae (C2, C3, C4) are reported in Table 1. The features of the examined vertebrae at the six consecutive observations can be summarized as follows:

T_1 . The lower border of C2 is flat in the vast majority of subjects at this stage; a concavity is evident at the lower border of C2 in only 7% of the individuals examined, a percentage that is not significant. The concavity is absent at the lower borders of both C3 and C4 in 100% of the subjects. The bodies of both C3 and C4 are trapezoid in shape.

Table 1 Results of Qualitative Analysis of Cervical Vertebral Characteristics at the Six Consecutive Observations

	T ₁		T ₂		T ₃		T ₄		T ₅		T ₆	
	Yes	No	Yes	No	Yes	No	Yes	No	Yes	No	Yes	No
Concavity at the lower border of C2	2 (6.7%)	28 (93.3%)	24 (80%)	6 (20%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)
Concavity at the lower border of C3	0 (0%)	30 (100%)	2 (6.7%)	28 (93.3%)	28 (93.3%)	2 (6.7%)	28 (93.3%)	2 (6.7%)	29 (96.7%)	1 (3.3%)	30 (100%)	0 (0%)
Concavity at the lower border of C4	0 (0%)	30 (100%)	0 (0%)	30 (100%)	3 (10%)	27 (90%)	26 (86.7%)	4 (13.3%)	29 (96.7%)	1 (3.3%)	30 (100%)	0 (0%)
C3 shape: trapezoid	30 (100%)	0 (0%)	29 (96.7%)	1 (3.3%)	23 (76.7%)	7 (23.3%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)
C4 shape: trapezoid	30 (100%)	0 (0%)	26 (86.7%)	4 (13.3%)	15 (50%)	15 (50%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)
C3 shape: rectangular horiz.	0 (0%)	30 (100%)	1 (3.3%)	29 (96.7%)	7 (23.3%)	23 (76.7%)	30 (100%)	0 (0%)	12 (40%)	18 (60%)	0 (0%)	30 (100%)
C4 shape: rectangular horiz.	0 (0%)	30 (100%)	4 (13.3%)	26 (86.7%)	15 (50%)	15 (50%)	30 (100%)	0 (0%)	14 (46.7%)	16 (53.3%)	0 (0%)	30 (100%)
C3 shape: squared	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	18 (60%)	12 (40%)	15 (50%)	15 (50%)
C4 shape: squared	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	16 (53.3%)	14 (46.7%)	16 (53.3%)	14 (46.7%)
C3 shape: rectangular vert.	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	15 (50%)	15 (50%)
C4 shape: rectangular Vert.	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	0 (0%)	30 (100%)	14 (46.7%)	16 (53.3%)

T₂. A concavity is present at the lower border of C2 in 80% of the subjects. The observation at T₂ is characterized also by the absence of a concavity at the lower borders of C3 (with the nonsignificant exception of 7% of the subjects) and of C4. Both C3 and C4 still are trapezoid in shape, with the nonsignificant exceptions of 3% and 13% of the subjects showing rectangular horizontal bodies for C3 and C4, respectively.

T₃. A concavity is present at the lower border of C2 (100% of the subjects) and of C3 (with the nonsignificant exception of 7% of the subjects). No concavity is present at the lower border of C4 (with the nonsignificant exception of 10% of the cases). The shape of both C3 and C4 may be either trapezoid or rectangular horizontal.

T₄. This observation is characterized by the presence of a concavity at the lower borders of C2, C3 (with the nonsignificant exception of 7% of the cases), and C4 (with the nonsignificant exception of 13% of the cases). The bodies of both C3 and C4 now are rectangular horizontal in shape (100% of the subjects).

T₅. A concavity is present at the lower borders of C2, C3 (with the nonsignificant exception of 3% of the cases), and C4 (with the nonsignificant exception of 3% of the cases). The body of C3 is rectangular horizontal in 40% of the cases and squared in the remaining subjects. The body of C4 is rectangular horizontal in 47% of the cases and squared in the remaining subjects.

T₆. A concavity is present at the lower borders of all the three examined cervical vertebrae. The body of C3 is squared in 50% of the cases and rectangular vertical in the remaining 50% of the cases. The body of C4 is squared in 53% of the cases and rectangular vertical in the remaining subjects.

Descriptive statistics for the cephalometric measurements of vertebral morphologic characteristics are reported in Table 2, together with the statistical comparisons between consecutive observations. No significant differences for any of the measurements were assessed between T₁ and T₂ with the exception of a significant increase in the depth of the concavity at the lower border of the second cervical vertebra (C2Conc). The depth of the concavities at the lower borders of both the second (C2Conc) and the third (C3Conc) cervical vertebra is significantly greater at T₃ when compared with T₂. In the transition from T₂ to T₃, the height of the anterior border of both C3 and C4 increases significantly, thus leading to significant decrements in the ratio between the heights of the posterior and anterior borders of the vertebral bodies (C3PAR and C4PAR).

At T₄, the depth of the concavity at the lower border of C4 (C4Conc) becomes significantly greater than at T₃. In the transition from T₃ to T₄, the height of the anterior borders of both C3 and C4 increases significantly again, thus leading to significant decreases both in the ratio between the heights of the posterior and anterior borders of the vertebral bodies (C3PAR and C4PAR) and in the ratio between the length of the base and the anterior height of the vertebral bodies (C3BAR and C4BAR). On average, C3PAR and C4PAR now

Table 2 Results of Quantitative Analysis: Descriptive Statistics and Statistical Comparisons (ANOVA for Repeated Measurements with Post Hoc Scheffe's test) on the Measurements at the Six Consecutive Cephalometric Observations.

	T ₁			T ₂			T ₃			T ₄			T ₅			T ₆		
	Mean	SD	SE	Mean	SD	SE	Mean	SD	SE	Mean	SD	SE	Mean	SD	SE	Mean	SD	SE
Age (months)	104.67	15.12	2.76	116.40	14.84	2.71	128.73	14.99	2.74	141.17	14.42	2.63	153.30	14.41	2.63	166.00	13.83	2.53
Co-Gn (mm)	107.85	5.33	0.97	110.32	5.68	1.04	112.63	5.78	1.06	118.05*	5.83	1.06	119.63	5.91	1.08	121.77	5.93	1.08
C2Conc (mm)	0.44	0.46	0.08	0.76*	0.49	0.09	1.15*	0.44	0.08	1.58*	0.37	0.07	1.91	0.40	0.07	2.23	0.48	0.09
C3Conc (mm)	0.01	0.41	0.07	0.36	0.49	0.09	0.95*	0.53	0.10	1.36	0.64	0.12	1.85	0.60	0.11	2.40*	0.68	0.12
C4Conc (mm)	0.03	0.31	0.06	0.12	0.29	0.05	0.31	0.39	0.07	1.07*	0.53	0.10	1.77*	0.60	0.11	2.28*	0.63	0.11
C3PAR (ratio)	1.35	0.15	0.03	1.26	0.16	0.03	1.16*	0.12	0.02	0.98*	0.08	0.01	0.98	0.06	0.01	0.99	0.06	0.01
C3BAR (ratio)	1.85	0.24	0.04	1.77	0.23	0.04	1.61*	0.15	0.03	1.39*	0.14	0.02	1.20*	0.15	0.03	1.03*	0.13	0.02
C4PAR (ratio)	1.34	0.13	0.02	1.25	0.14	0.03	1.15*	0.17	0.03	1.01*	0.06	0.01	1.00	0.07	0.01	0.97	0.05	0.01
C4BAR (ratio)	1.83	0.25	0.05	0.71	0.22	0.04	1.59	0.21	0.04	1.36*	0.12	0.02	1.19*	0.18	0.03	1.04	0.12	0.02

*Indicates statistical significance with respect to the preceding observation.

have a ratio of approximately 1:1, an indication that both C3 and C4 vertebral bodies are rectangular horizontal in shape.

T₅ and T₆ are characterized by decrements of the ratio between the length of the base and the anterior height of the vertebral bodies (C3BAR and C4BAR). The mean values for these measurements indicate that the vertebral bodies become progressively more squared in shape. At T₆, one third of the cases show a rectangular vertical shape of one or both C3 and C4 vertebral bodies.

Discriminant analysis revealed that the forming concavity at the lower border of C2 can account for 80% of the differences between T₁ and T₂. The depth of C3Conc becomes the discriminant variable between T₂ and T₃ with a classifying power of 75%. The difference in the posteroanterior ratio of C3 (C3PAR) together with the depth of the concavity at the lower border of C4 (C4Conc) are the discriminant factors between T₃ and T₄ (classifying power equal to 85%). C3PAR in association with both the ratio between the length of the base and the anterior height of C3 (C3BAR) and C4Conc are able to discriminate between T₄ and T₅ in 88% of the cases. The ratios for C3 (C3BAR and C3PAR) together with the depth of the concavity at the lower border of C2 (C2Conc) are the discriminant variables between T₅ and T₆ in 80% of the cases.

Discussion

The modifications in the size and shape of the cervical vertebrae in growing subjects have gained increasing interest during the past few decades as a biological indicator of individual skeletal maturity. One of the main reasons for the increasing popularity of the method is that the analysis of cervical vertebral maturation is performed on the lateral cephalogram, a type of film used routinely in orthodontic diagnosis. The objective of the present investigation was to provide a refinement of the method through the definition of six stages (cervical stages 1 to 6) for a more practical application in dentofacial orthopedics, and more specifically:

- a direct appraisal of the skeletal maturity of the mandible in relation to the morphological features of the cervical vertebrae;
- an evaluation of the morphological features of the cervical vertebral bodies restricted to those that are visible on the lateral cephalogram even when a protective collar is worn, as originally proposed by Hassel and Farman²⁰;
- a definition of the cervical vertebral morphology at each developmental stage that allows the clinician to apply the CVM method on the basis of the information derived from a single cephalogram. The assessment of individual stages in cervical vertebral maturation through the comparative analysis of between-stage changes should be avoided.

The anatomical features of the second (odontoid process), third, and fourth cervical vertebrae were evaluated here as visualized on lateral cephalograms in a time interval ranging on average from 2 years before to 2 years after the peak in mandibular growth. The description of the consecutive

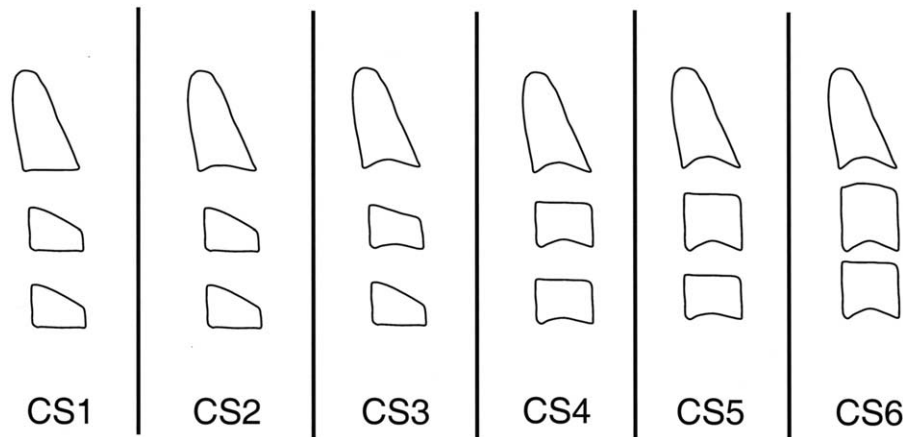


Figure 2 Schematic representation of the stages of cervical vertebrae according to the newly modified method.

stages in vertebral development consisted of a noncomparative definition of morphological characteristics at each observation.

The findings of both the visual (qualitative) and cephalometric (quantitative) analyses revealed that a statistically significant discrimination can be made between the initial two stages in cervical vertebral maturation only according to the difference in depth of the concavity at the lower border of the second cervical vertebra. A definite concavity at the lower border of C2 is present in 80% of the subjects at cervical stage 2.

The appearance of a visible concavity at the lower border of the third cervical vertebra is the anatomic characteristic that mostly accounts for the identification of the stage immediately preceding the peak in mandibular growth (cervical stage 3). The distinction among Cvs 4, Cvs 5, and Cvs 6 as defined in the former CVM method is possible only by using the shape of the bodies of C3 and/or C4 as a discriminant factor.¹⁸

Stages of Cervical Vertebral Maturation

The stages of cervical vertebral maturation in the modified version of the method presented here are illustrated diagrammatically in Fig 2. The six stages are defined as follows:

Cervical stage 1 (CS1, Fig 3). The lower borders of all the three vertebrae (C2-C4) are flat. The bodies of both C3 and C4 are trapezoid in shape (the superior border of the vertebral body is tapered from posterior to anterior). The peak in mandibular growth will occur on average 2 years after this stage.

Cervical stage 2 (CS2, Fig 4). A concavity is present at the lower border of C2 (in four of five cases, with the remaining subjects still showing a cervical stage 1). The bodies of both C3 and C4 are still trapezoid in shape. The peak in mandibular growth will occur on average 1 year after this stage.

Cervical stage 3 (CS3, Fig 5). Concavities at the lower borders of both C2 and C3 are present. The bodies of C3 and C4 may be either trapezoid or rectangular horizontal in shape.

The peak in mandibular growth will occur during the year after this stage.

Cervical stage 4 (CS4, Fig 6). Concavities at the lower borders of C2, C3, and C4 now are present. The bodies of both C3 and C4 are rectangular horizontal in shape. The peak in mandibular growth has occurred within 1 or 2 years before this stage.

Cervical stage 5 (CS5, Fig 7). The concavities at the lower borders of C2, C3, and C4 still are present. At least one of the bodies of C3 and C4 is squared in shape. If not squared, the body of the other cervical vertebra still is rectangular horizontal. The peak in mandibular growth has ended at least 1 year before this stage.

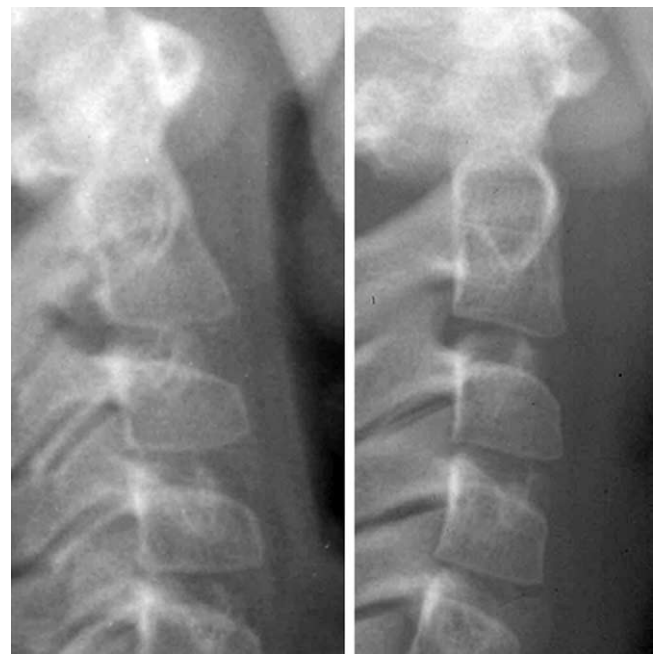


Figure 3 Cervical stage 1 (CS1): two clinical examples.



Figure 4 Cervical stage 2 (CS2): two clinical examples.

Cervical stage 6 (CS6, Fig 8). The concavities at the lower borders of C2, C3, and C4 still are evident. At least one of the bodies of C3 and C4 is rectangular vertical in shape. If not rectangular vertical, the body of the other cervical vertebra is squared. The peak in mandibular growth has ended at least 2 years before this stage.



Figure 5 Cervical stage 3 (CS3): two clinical examples.



Figure 6 Cervical stage 4 (CS4): two clinical examples.

Application to Dentofacial Orthopedics

The clinical application of the method to dentofacial orthopedics becomes relevant for those treatment protocols that benefit from the inclusion of the period of accelerated mandibular growth. CVM method can be useful as a maturational index to detect the optimal time to start treatment of mandibular deficiencies by means of functional jaw orthopedics.^{25,26} It has been demonstrated that the effectiveness of functional treatment of Class II skeletal disharmony depends strongly on the biological responsiveness of the condylar cartilage, which in turn is related to the growth rate of the mandible.²⁷



Figure 7 Cervical stage 5 (CS5): two clinical examples.



Figure 8 Cervical stage 6 (CS6): two clinical examples.

When CS1 or CS2 are diagnosed in the individual patient with mandibular deficiency, the clinician can wait at least one additional year for a radiographic reevaluation aimed to start treatment with a functional appliance. The appearance of a definite concavity at the lower border of C2 indicates that the growth spurt is approaching, that is, that the year of the peak will start approximately 1 year after this stage. CS3 represents the ideal stage to begin functional jaw orthopedics, as the peak in mandibular growth will occur within the year after this observation. In the sample examined here, total mandibular length exhibited an average increase of 5.4 mm in the year following CS3, a significantly greater increment when compared with the growth interval from CS1 to CS2 (about 2.5 mm), from CS2 to CS3 (again about 2.5 mm), and to the postpeak between-stage intervals (1.6 mm and 2.1 mm for the intervals from CS4 to CS5 and from CS5 to CS6, respectively).

Treatment Timing for Class II Malocclusion

An emerging fundamental concept underlying Class II correction is that this type of intervention should be undertaken when the likelihood for a maximum growth response is high, that is, during the circumpubertal growth period. A series of short-term studies has demonstrated statistically and clinically significant correction of the Class II dentoskeletal relationships when either functional appliances or fixed appliances in combination with Class II elastics are used during the circumpubertal period (Table 3). When Class II malocclusion is treated too early (therapy starting at CS1 and completed before the interval of peak velocity in mandibular growth, ie, before CS3), the net difference in supplementary growth of the mandible (expressed cephalometrically by the measurement Co-Pg or Co-Gn) in the treated samples versus untreated controls ranges between 0.4 mm and 1.8 mm (Table 3). On the contrary, when intervention in a Class II pa-

Table 3 Analysis of the Literature Regarding Treatment Timing for Class II Malocclusion*

The Effect of Treatment Timing on Supplementary Elongation of the Mandible in Class II Treatment		
Pre-Pubertal Class II Treatment (treatment ends before the pubertal peak in mandibular growth)		Pubertal Class II Treatment (treatment includes the pubertal peak in mandibular growth)
Study	Appliance	Net increase in mandibular length over untreated controls
McNamara et al., 1985 ²⁸	FR-2	+1.2 mm
Petrovic et al., 1994 ²⁹	Class II elastics	+1.0 mm
Tulloch et al., 1997 ³⁰	Bionator	+1.4 mm
Keeling et al., 1998 ³¹	Bionator	+0.4 mm
Baccetti et al., 2000 ²⁵	Twin-block	+1.8 mm
Baccetti and Franchi, 2001 ²⁶	FR-2	+1.0 mm
De Almeida et al., 2002 ³²	FR-2	+0.9 mm
Janson et al., 2003 ³³	FR-2	+0.5 mm
O'Brien et al., 2003 ³⁴	Twin-block	+1.6 mm
Faltin et al., 2003 ³⁵	Bionator	+0.8 mm
McNamara et al., 1985 ²⁸	FR-2	+3.6 mm
Petrovic et al., 1994 ²⁹	Class II elastics	+3.0 mm
Lund and Sandler, 1998 ³⁶	Twin-block	+2.4 mm
Franchi et al., 1999 ³⁷	Acrylic Herbst	+2.7 mm
Baccetti et al., 2000 ²⁵	Twin-block	+4.7 mm
Baccetti and Franchi, 2001 ²⁶	FR-2	+3.9 mm
Faltin et al., 2003 ³⁵	Bionator	+4.3 mm

*The appraisal of treatment timing in individual studies was based upon chronologic age, hand and wrist, or CVM method. All data are short-term and refer to controlled studies.

tient includes the CS3-CS4 interval (growth spurt), the net supplementary growth of the mandible in treated samples versus untreated controls ranges from 2.4 mm to 4.7 mm (Table 3). The data reported in Table 3 suggest also that in Class II patients, the timing of therapeutic intervention has a greater impact on supplementary elongation of the mandible than does the type of appliance used.

The only long-term study that deals with the evaluation of the role of treatment timing in Class II correction³⁵ revealed that the use of a Bionator followed by fixed appliances in contrast with untreated Class II controls is able to induce a supplementary elongation of the mandible of less than 2 mm when the functional appliance is used before the peak in mandibular growth, and of about 5 mm when the growth spurt is included in the treatment interval. These results possess significance not only at the statistical level, but also at the clinical level, as the correction of a full cusp Class II molar relationship to Class I represents a 5 to 6 mm sagittal correction at the level of the occlusal plane.

Treatment Timing for Class III Malocclusions

Early treatment of Class III disharmony has been advocated for a long time.³⁸ The clinical understanding that Class III malocclusion is established early in life and that it is not a self-correcting disharmony has led to the recommendation of intervention as early as in the deciduous dentition. Cephalometric and morphometric investigations using Class III untreated controls have demonstrated that treatment of Class III malocclusion by means of efficient protocols (eg, maxillary expansion and protraction) is more effective in the early than in the late mixed dentition.³⁹⁻⁴¹

Until recently, however, information about the possible role of treatment timing on long-term changes after active therapy for Class III malocclusion was not available in the literature.⁴² At a postpubertal observation (CS5 or CS6), when active growth of the craniofacial skeleton is completed for the most part, Class III subjects treated with a rapid maxillary expander and a facial mask well before the growth spurt (CS1) present with different long-term changes with respect to Class III subjects treated at a later stage, that is, at the peak in mandibular growth (CS3). Prepubertal orthopedic treatment of Class III malocclusion is effective both in the maxilla (which shows a supplementary growth of about 2 mm over Class III untreated controls) and in the mandible (restriction in growth of about 3.5 mm over controls), whereas treatment of Class III malocclusion at puberty is effective at the mandibular level only (restriction in growth of about 4.5 mm over controls).⁴²

The findings in the maxilla have a biological explanation in the physiology of the circummaxillary sutures, which are more amenable to orthopedic intervention during the early stages, whereas they become more heavily interdigitated around puberty.⁴³ On the other hand, the possibility of restricting mandibular growth both before and during puberty gives the clinician the chance of resuming facemask therapy at a later time when correction of Class III relationships is only partial after the prepubertal intervention.

Treatment Timing for Transverse Maxillary Deficiency

The issue of treatment timing for maxillary expansion aimed to correct transverse maxillary deficiency has been addressed in the past by Melsen⁴⁴ and by Wertz and Dreskin.⁴⁵ Melsen used autopsy material to examine histologically the maturation of the midpalatal suture at different developmental stages.⁴⁴ In the infantile stage (up to 10 years of age), the suture was broad and smooth, whereas in the juvenile stage (from 10 to 13 years) it had developed in a more typical squamous suture with overlapping sections. Finally, during the adolescent stage (13 and 14 years of age) the suture was wavier with increased interdigitation. From these histological data, the inference is that patients who show an advanced stage of skeletal maturation at the midpalatal suture may have difficulty in undergoing orthopedic maxillary expansion. Clinical support for the histologic findings by Melsen⁴⁴ is derived from the results of a study by Wertz and Dreskin⁴⁵ who noted greater and more stable orthopedic changes in young patients (under the age of 12 years). Either group of researchers, however, did not use any biological indicator of skeletal maturity to define "early" versus "late" treatment.

The use of the CVM method has been applied recently to the estimate of the effects of different treatment timing on the correction of transverse maxillary deficiency.⁴⁶ A sample of 42 patients was compared with a control sample of 20 subjects. Posteroanterior cephalograms were analyzed for each of the treated subjects at T₁ (pretreatment), T₂ (immediate post-expansion), and T₃ (long-term observation); films were available at T₁ and at T₃ for the controls. The mean age at T₁ was 11 years and 10 months for both the treated and the control groups. The mean ages at T₃ also were comparable (20 years 6 months for the treated group, and 17 years 8 months for the control group). Following rapid maxillary expansion and retention (2 months on average), fixed standard edgewise appliances were placed. The study included transverse measurements on dentoalveolar structures, maxillary and mandibular bases, and other craniofacial regions (nasal, zygomatic, orbital, and cranial).

Treated and control samples were divided into two groups according to individual skeletal maturation as evaluated by the CVM method. The early treated and early control groups consisted of subjects who had not reached the pubertal peak in skeletal growth velocity at T₁ (CS1 through CS3), whereas the late-treated and late control groups were comprised of subjects during or slightly after the peak at T₁ (CS4 through CS6). The group treated before the pubertal peak showed significantly greater short-term increases in the width of the nasal cavities. In the long-term, increments in maxillary skeletal width, maxillary intermolar width, lateronasal width, and latero-orbitale width were significantly greater in the early-treated group when compared with the corresponding control group. The late-treated group exhibited significant increases in lateronasal width and in maxillary and mandibular intermolar widths. The use of the CVM method demonstrated that rapid maxillary expansion before the peak in skeletal growth velocity is able to induce more pronounced transverse craniofacial changes at the skeletal level. Treat-

ment changes are more dentoalveolar in nature when expansion is performed during or after the peak.

Treatment Timing for Increased Vertical Dimension

The CVM method also has been applied to the appraisal of ideal treatment timing for a specific therapeutic protocol for the correction of vertical excess of the face by means of a bonded rapid maxillary expander in association with a vertical-pull chin cup. One of the goals of orthopedic treatment in subjects with increased vertical dimension is the control of the vertical growth of the mandibular ramus (expressed cephalometrically by the measure Co-Go). Available short-term data from our research group show that a significantly more favorable effect can be obtained when treatment is performed at CS3, that is, at the peak in mandibular growth, when compared with treatment performed at an earlier maturational stage (CS1). No significant increase in ramal height is observed in hyperdivergent subjects treated at CS1, whereas a significant increase of about 2 mm more than in untreated controls is recorded in hyperdivergent subjects who receive orthopedic treatment at CS3.

Final Remarks

The CVM method is comprised of six maturational stages (cervical stage 1 through cervical stage 6, CS1-CS6), with the peak in mandibular growth occurring between CS3 and CS4. The pubertal peak has not been reached without the attainment of both CS1 and CS2. In particular, the detection of CS2 indicates that the growth spurt is approaching, and it will start at CS3, which is approximately 1 year after CS2. Active growth is virtually completed when the CS6 is attained.

The method is particularly useful when skeletal maturity has to be appraised on a single cephalogram and only the cervical vertebrae from the second one through the fourth one are visible. The CVM method has the further advantage to be assessed on the lateral cephalogram, which is the radiographic record used routinely for orthodontic diagnosis and treatment planning.

The use of a reliable biological indicator of skeletal maturity such as the CVM method is highly recommended for a wide variety of research and clinical applications. In both prospective and retrospective controlled studies, CVM stages enable the researcher to categorize treated/untreated subjects for a biologically appropriate matching between experimental and control samples. Further, the appraisal of the CVM stage in the individual subject allows for a more precise definition of early and late samples in studies aimed to determine the role of treatment timing in the effectiveness of different treatment protocols for the correction of malocclusions. To date, the application of the method in investigations on treatment timing in orthodontics and dentofacial orthopedics has revealed that:

1. Class II treatment is most effective when it includes the peak in mandibular growth;
2. Class III treatment with maxillary expansion and protraction is effective in the maxilla only when it is performed before the peak (CS1 or CS2), whereas it is

effective in the mandible during both prepubertal and pubertal stages;

3. skeletal effects of rapid maxillary expansion for the correction of transverse maxillary deficiency are greater at prepubertal stages, while pubertal or postpubertal use of the rapid maxillary expander entails more dentoalveolar effects; and
4. deficiency of mandibular ramus height can be enhanced significantly in subjects with increased vertical facial dimension when orthopedic treatment is performed at the peak in mandibular growth (CS3).

To summarize, effects of therapies aimed to enhance/restrict mandibular growth appear to be of greater magnitude at the circumpubertal period during which the growth spurt occurs in comparison to earlier intervention, while effects of therapies aimed to alter the maxilla orthopedically (maxillary protraction/maxillary expansion) are greater at prepubertal stages.

The CVM method can be helpful for the assessment of completion of active growth in studies dealing with the long-term effects of orthodontic/orthopedic treatment strategies. Similarly, the method can be used to identify clinically the adequate time for intervention in subjects who need surgery for the late correction of facial disharmonies.

Due to its practical applications, the CVM method appears to be a powerful diagnostic tool. The implementation of the method in orthodontic decision making allows for an improvement of treatment outcomes by combining effective and efficient protocols with optimal treatment timing.

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