

## *Local and Systemic Influences in Periodontal Disease:*

### II. EFFECT OF PROPHYLAXIS AND NATURAL VERSUS SYNTHETIC VITAMIN C UPON GINGIVITIS\*

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THERE is growing evidence that periodontal disease is the result of the interplay of the local environment and the host.<sup>1-6</sup> The purpose of this study is to analyze the relative effects of natural versus synthetic vitamin C with and without bioflavonoids and with and without scaling upon the gingival state.

#### REVIEW OF THE LITERATURE

For this study the review will be limited to one oral factor (calculus) since it was the local variable which was altered and three systemic factors (natural vitamin C concentrate versus synthetic vitamin C with and without bioflavonoids) since these were varied.

Of all of the known local factors, calculus is considered the most important of the gingival irritants.<sup>7-11</sup> Whether calculus actually initiates the inflammation or whether gingival inflammation caused by other irritants is necessary before calculus can be formed remains to be established.<sup>9</sup> Whatever the sequence of events, the irritating effect of calculus as a result of the combined mechanical, chemical and/or microbial factors, contributes to the development of gingival inflammation. As the calculus increases in size, the inflammatory reaction spreads and gingival recession eventually occurs. Progressively the deeper periodontal tissues become involved with pocket formation.

Generally speaking, lower animals subjected to vitamin C deficient diets show detrimental changes in the gingiva, periodontal membrane and alveolar bone.<sup>12-16</sup>

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Human studies have likewise revealed that there is a correlation between ascorbic acid status and the appearance of the gingiva.<sup>17-21</sup> The manifestations of vitamin C deprivation are held to proceed in a definite order affecting in sequence the interdental papilla, marginal and the alveolar gingiva.<sup>22-23</sup>

Increasing attention is being paid to capillary integrity. The bioflavonoids have been found to be one of the essential factors in the maintenance of capillary resistance. Some few investigators have reported upon the successful use of the water-soluble bioflavonoid preparations in the treatment of the periodontal diseases.<sup>24-26</sup>

Very limited reports indicate the possible superiority of natural versus synthetic vitamin C.<sup>27</sup> No known stomatologic studies are available.

#### METHOD OF INVESTIGATION

One hundred and two subjects participated in this study. The selection was as random as possible from volunteers at the University of Alabama Medical and Extension Centers and from the Birmingham Fire Department. Table 1 shows the patient distribution in terms of age and sex.

All subjects reported to the Department of Oral Medicine following a 12-hour fast. Upon admission, a fasting venous blood sample was drawn. The oral examination was conducted with the patient comfortably seated in a dental chair. The mouth was illuminated with a dental spotlight. In all cases, the examination started in the maxillary right canine region and progressed across to the left cuspid. The same pattern was followed in the mandible. One

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TABLE 1  
Patient Age and Sex Distribution

Age Groups	Male Group		Female Group		Total Group	
	Number	Percentage	Number	Percentage	Number	Percentage
20-29	25	25.5	27	26.5	52	51.0
30-39	17	16.7	9	8.8	26	25.5
40-49	12	11.8	6	5.9	18	17.7
50-59	2	1.9	4	3.9	6	5.8
Total	56	54.9	46	45.1	102	100.0

examiner (G. M. A.) made all of the observations.

**Calculus Scores:** Although the quantity of calculus undoubtedly has a bearing upon the degree of pathosis, it was felt that the location of these deposits was of major importance. Hence, the evaluation was based upon whether there was an absence of calculus or whether supragingival, subgingival, or both supra- and subgingival calculus was present on each tooth examined. The inspection was done with an explorer and mirror. Compressed air was used to retract the gingiva for better detection of subgingival deposits. The legend for scoring calculus is shown (Table 2). In the individual with all 12 anterior teeth, 12 calculus scores were recorded. The individual scores were then added and divided by the number of teeth to obtain a mean value for each side.

**Gingival Scores:** The soft tissues surrounding the 12 anterior teeth were evaluated by scoring the labial and lingual mar-

ginal gingiva on each tooth and the labial and lingual aspects of the gingival papillae between each tooth. Thus, in a subject with 12 anterior teeth, 48 measurements were obtained. Table 3 shows the gingival scoring system. From the individual measurements, mean right and left values were derived. Finally, a color photograph of the anterior teeth and gingiva was taken.

**Measurement of Vitamin C State:** The method of Mindlin and Butler<sup>28</sup> was utilized for the determination of plasma ascorbic acid levels. The test was always done twice from equal portions of the same blood sample. These duplicate determinations served as a check on the accuracy of the procedure ( $r = +0.988$ ,  $P < 0.001$ ).

For the determination of tissue ascorbic acid status, the lingual vitamin C test was employed.<sup>29-30</sup> This is a simple procedure which involves the timing (in seconds) of the decolorization of one minim of N/300 dichlorophenolindolphenol dye deposited upon the dorsum of the dried tongue. In order to evaluate the reproducibility of the test, the procedure was always performed twice. The correlation between

TABLE 2  
Calculus Scoring System

0=no calculus present
1=supragingival calculus covering not more than one-third of the exposed tooth surface
2=supragingival calculus covering more than one-third but not more than two-thirds of the exposed tooth surface or the presence of individual flecks of subgingival calculus around the cervical portion of the tooth or both
3=supragingival calculus covering more than two-thirds of the exposed tooth surface or a continuous heavy band of subgingival calculus around the cervical portion of the tooth or both

TABLE 3  
Gingival Evaluation

0=no gingivitis present
1=slight hyperemia, swelling and loss of stippling, patient unaware of the condition
2=moderate hyperemia, swelling and loss of stippling, tendency to bleed upon pressure and may be tender or painful
3=marked hyperemia, swelling and loss of tissue tone, bleeds so-called spontaneously, may be ulcerated, tender and painful

the first and second lingual time readings was found to be highly significant ( $r = +0.989$ ,  $P < 0.001$ ).

Bioflavonoid state was not ascertained since there are no known laboratory tests.

By means of a table of random numbers the patients were arranged into two groups: (1) those to be scaled on the right side, and (2) those to receive prophylaxis on the left. Also, the patients were randomized to receive one of the systemic regimes. Thus, in terms of systemic therapy, four groups were developed. Table 4 shows the composition of the capsules which were administered. All capsules looked exactly alike. Thus, neither the patient nor the examiner were aware at any time during the experiment which preparation was employed.

At the first visit, each patient received thorough scaling of the teeth on one side of the mouth. Jaquette and McCall scalers were employed to remove the calcareous deposits. Moistened flower of pumice was applied with a revolving rubber cup in a contra-angle handpiece to polish the accessible crown and root surfaces. Finally, dental tape and pumice were utilized to polish the interproximal surfaces of the teeth.

At the completion of the first visit each patient was instructed to take by mouth three capsules per day for three weeks. Thus, 25 subjects received a placebo, 25 were given 300 mgm. synthetic vitamin C daily, 25 were administered 300 mgm. synthetic vitamin C plus 300 mgm. citrus bioflavonoids per day, and the remaining 27 were supplemented with 300 mgm. of a natural vitamin C concentrate with 300 mgm. bioflavonoids each day.

Approximately 21 days later each of the patients returned. All participants were re-examined clinically and biochemically without reference to the earlier records or the nature of the supplementation.

#### RESULTS

*Pretberapy Findings:* Table 5 summar-

TABLE 4  
Codes and Composition of the Capsules

#1	100 mg. milk sugar per capsule
#2	100 mg. synthetic vitamin C
#3	100 mg. synthetic vitamin C plus 100 mg. citrus bioflavonoids
#4	100 mg. natural vitamin C concentrate plus 100 mg. citrus bioflavonoids

izes the initial calculus scores. It is clear that about one-half of the examined areas are without calculus. The initial gingival scores are also shown (Table 6). About 13 per cent of the gingival areas showed no signs of pathosis. The remaining 87 per cent ranged from slight pathosis (57.4 per cent) to severe gingivitis (7.9 per cent). Table 7 outlines the patient distribution in terms of the initial plasma ascorbic acid levels. It was found that the scores ranged from 0 to 1.38 mgm. per cent. The distribution of the initial lingual times is included (Table 8). The values ranged from 12 to 60 seconds.

The relationship between calculus and gingivitis is pictorially portrayed (Figure 1). It will be noted that the subjects with low calculus scores also have low mean gingivitis values. Conversely, those individuals with high calculus ratings are associated with high mean gingival readings. A significant correlation was found between calculus and gingivitis ( $r = +0.568$ ,  $P < 0.001$ ).

Figure 2 illustrates graphically the correlation of plasma ascorbic acid to gingivitis. The chart shows that those subjects

TABLE 5  
Calculus Distribution  
(Initial Scores)

Calculus Scores	Number of Examinations	Percentage of Examinations
0	585	49.3
1	466	39.2
2	112	9.4
3	25	2.1
Total	1188	100.0

TABLE 6  
Distribution of Initial Gingival Scores

Gingival Scores	Number of Examinations	Percentage of Examinations
0	619	13.0
1	2734	57.4
2	1032	21.7
3	378	7.9
Total	4763	100.0

with the highest mean gingivitis scores are associated with the lowest vitamin C levels. On the other hand, those persons with the lowest mean gingival values are in parallel with the highest plasma vitamin C levels. The relationship between plasma ascorbic acid and gingivitis was found significant ( $r = -0.218$ ,  $P < 0.005$ ). The correlation between the lingual times and gingivitis is also shown (Figure 3). Subjects with high mean gingivitis ratings are associated with long lingual times; individuals with low gingival values have short lingual times. A statistically significant correlation was obtained ( $r = +0.200$ ,  $P < 0.005$ ).

The relationship of both calculus and vitamin C state (plasma vitamin C levels) is included (Table 9). The participants with the lower calculus (0.0-0.5) and higher vitamin C levels (0.60-1.40) have the lowest gingival scores ( $1.0 \pm 0.2$ ). Those individuals with the higher calculus values ( $> 0.5$ ) and poorer vitamin C levels (0.00-0.59) are associated with the greatest gingivitis ratings ( $1.8 \pm 0.6$ ). As one proceeds from left to right (irrespective of plasma ascorbic acid levels) the gin-

TABLE 7  
Patient Distribution in Terms of Initial Plasma Ascorbic Acid

Plasma Ascorbic Acid Groups	Number of Patients	Percentage of Patients
0.00-0.59	36	35.3
0.60-0.99	45	44.1
1.00-1.40	21	20.6
Total	102	100.0

TABLE 8  
Patient Distribution in Terms of Initial Lingual Time

Lingual Time Groups (Seconds)	Number of Patients	Percentage of Patients
Less than 20	28	27.5
20-30	50	49.0
More than 30	24	23.5
Total	102	100.0

givitis increases with increase in calculus values. As one moves from top to bottom in either column (irrespective of calculus state) the gingivitis increases with a decrease in the vitamin C level.

Though interesting, the preceding relationships do not necessarily demonstrate cause-and-effect. These correlations simply indicate that variables co-exist. To try and pinpoint possible cause and effect, it was felt advisable to study the gingival pattern after both local (prophylaxis) and systemic therapy (synthetic and natural vitamin C concentrate with and without bioflavonoids).

*Posttherapy Findings:* The nature of the experimental design provided the opportunity to study the changes in the gingivae after both local (prophylaxis) and/or systemic (synthetic and natural vitamin C concentrate with and without bioflavonoids) therapy.

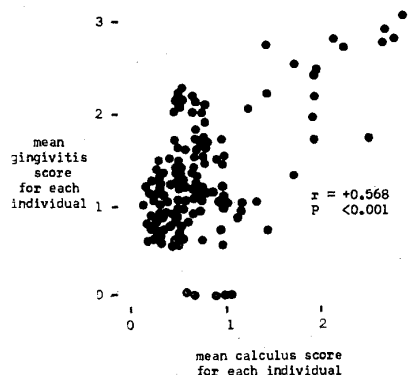


Fig. 1. Pretherapy relationship of a local factor (calculus) to periodontal pathosis (gingivitis).

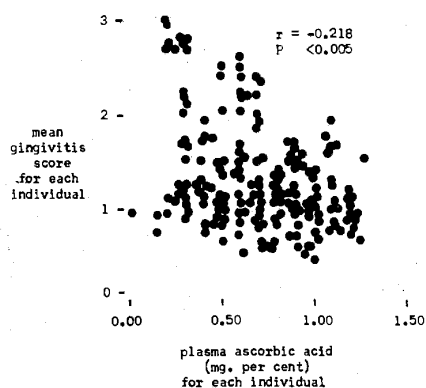


Fig. 2. Pretherapy relationship of a systemic factor (plasma ascorbic acid) to periodontal pathosis (gingivitis).

**Effect of Scaling:** In each subject one half of the teeth were scaled at the first visit. Thus, in the placebo group, it was possible to examine the effect of the removal of calculus upon the periodontium on one side versus the untouched (nonscaled) other side of the gingiva. Table 10 shows the original and mean initial and final gingival scores for the scaled and nonscaled sides in those persons without systemic treatment (placebo). There appears to be a 30 per cent and significant ( $P < 0.050$ ) improvement in mean gingivitis score with scaling. The nonscaled side showed an insignificant ( $P > 0.200$ ) slight reduction (10 per cent).

**Effect of Systemic Therapy:** It was also possible to evaluate the effect of altering the host without local treatment (no scaling). Table 11 includes the initial and final original and mean gingival ratings of those subjects receiving systemic therapy versus placebo supplementation. There appears to be a 57 per cent improvement in the mean gingivitis scores ( $P < 0.001$ ) in both the groups subjected to natural vitamin C concentrate and synthetic vitamin C with bioflavonoids. The synthetic vitamin C group without bioflavonoids followed with 45 per cent significant reduction ( $P < 0.001$ ). Finally, the 10 per cent decrease in the placebo group is not significant ( $P > 0.200$ ).

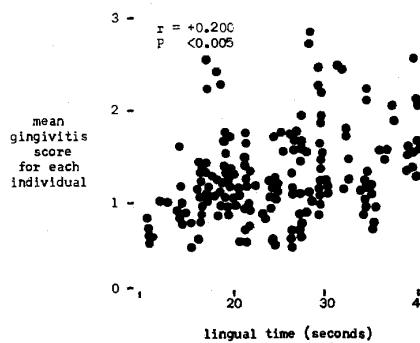


Fig. 3. Pretherapy relationship of a systemic factor (lingual time) to periodontal pathosis (gingivitis).

**Effect of Local and Systemic Therapy:** Table 12 illustrates that the greatest improvement on a mean basis (71 per cent) occurred in those persons subjected to both prophylaxis and natural vitamin C concentrate with bioflavonoids. In order followed those with scaling and synthetic vitamin C with bioflavonoids (69 per cent). The next greatest improvement was shown with prophylaxis and synthetic vitamin C without bioflavonoids. Finally, the patients with only local therapy (placebo group) had a decrease of 30 per cent. On a statistical basis, all groups given both local and systemic treatment had more significant percentage improvement ( $P < 0.001$ ) than those given only local therapy ( $P < 0.050$ ).

#### DISCUSSION

It should be recalled that two possible factors were studied in terms of their relationship to gingival pathosis. These variables were calculus, representing the local prob-

TABLE 9  
Pretherapy Relationship of the Mean Calculus Scores and Ascorbic Acid Levels (mg. per cent) to the Subjects' Mean Gingivitis Scores

Plasma Ascorbic Acid Groups	Calculus Groups	
	0.0-0.5	>0.5
0.60-1.40	1.0±0.2*	1.4±0.4*
0.00-0.59	1.2±0.3*	1.8±0.6*

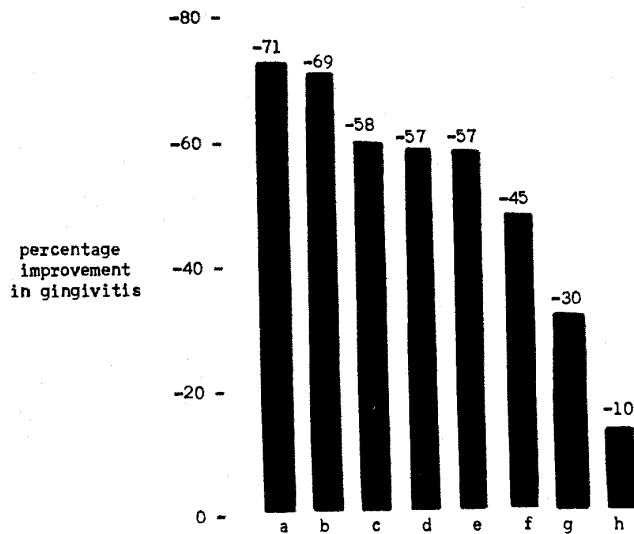
\*Mean and standard deviation gingivitis scores.

TABLE 10  
Gingivitis Change after Prophylaxis  
(Placebo Group)

Gingival Grades	Unscaled Side		Scaled Side	
	Initial Scores	Final Scores	Initial Scores	Final Scores
0	120	196	116	243
1	336	268	360	258
2	80	75	67	57
3	24	21	29	14
Total	560	560	572	572
Mean	1.0	0.9	1.0	0.7
S.D.	0.4	0.5	0.4	0.5
Percentage Change		-10		-30
P		>0.200		<0.050

lem, and vitamin C state on the systemic side. The higher correlation ( $r = +0.568$ ,  $P < 0.001$ ) was found between calculus and gingivitis. Then followed the correlation of plasma ascorbic acid to gingivitis ( $r = -0.218$ ,  $P < 0.005$ ). An equally significant relationship was obtained between the lingual time and gingivitis ( $r = +0.200$ ,  $P < 0.005$ ). It is well to underscore the fact that, at the initial visit, the correlation of calculus and gingivitis is greater than that of vitamin C and gingival states.

Figure 4 graphically depicts the percentage change in gingival state with the eight different forms of therapy. Table 13



- a = prophylaxis with natural vitamin C concentrate and bioflavonoids
- b = prophylaxis with synthetic vitamin C and bioflavonoids
- c = prophylaxis with synthetic vitamin C
- d = no prophylaxis with natural vitamin C concentrate and bioflavonoids
- e = no prophylaxis with synthetic vitamin C and bioflavonoids
- f = no prophylaxis with synthetic vitamin C
- g = prophylaxis with placebo
- h = no prophylaxis with placebo

Fig. 4. Posttherapy percentage gingivitis change.

TABLE 11  
Gingivitis Change after Systemic Therapy  
(Unscaled Side)

Gingival Grades	Natural Vitamin C Concentrate Plus Bioflavonoids		Synthetic Vitamin C Plus Bioflavonoids		Synthetic Vitamin C		Placebo	
	Initial	Final	Initial	Final	Initial	Final	Initial	Final
0	57	316	70	282	83	299	120	196
1	359	273	289	257	388	259	336	268
2	142	42	190	59	105	33	80	75
3	74	1	52	3	18	3	24	21
Total	632	632	601	601	594	594	560	560
Mean	1.4	0.6	1.4	0.6	1.1	0.6	1.0	0.9
S.D.	0.6	0.3	0.5	0.3	0.4	0.3	0.4	0.5
Percentage Change	-57		-57		-45		-10	
P	<0.001		<0.001		<0.001		>0.200	

summarizes the intergroup statistical significance. It is noteworthy that all non-scaled groups were statistically significantly different except for the nonscaled natural versus synthetic vitamin C with bioflavonoids (57 per cent). Table 13 also illustrates that all scaled groups were significantly different except for the gingivitis reductions (71 and 69 per cent) in those subjects treated with natural versus synthetic vitamin C with bioflavonoids (Table 12). It is also clearly shown that only two of the four groups were significantly different (Tables 11 and 12) in terms of scaling versus non-scaling. Specifically, natural vitamin C concentrate with bioflavonoids with (71 per cent) and without scaling (57 per cent) were significantly different. Also, the placebo groups with (30 per cent) and without prophylaxis (10 per cent) were statistically different. It appears that both synthetic and natural vitamin C concentrate exerts a beneficial effect in reducing gingivitis and particularly so along with bioflavonoids and prophylaxis.

It should be recalled that vitamin therapy was administered on a random basis. Obviously, by this technique, some of the vitamin-supplemented subjects actually had satisfactory vitamin C levels at the start and therefore did not require vitamin ther-

apy. Hence, it was thought proper to re-study the gingival patterns in the subjects with the lowest plasma ascorbic acid levels and highest lingual times. For this phase, the ten patients in each group with the poorest scores were selected. Figure 5 outlines the findings for the entire groups (white columns) and those with the poorer vitamin C findings (black columns).

Three points deserve particular attention. Firstly, the overall pattern for the 10 subjects follows the pattern for the entire group. Secondly, in each group both locally and systemically treated, greater improvement occurred in the selected group than in the entire sample. Finally, those 10 persons given only local therapy showed less change than the entire group. This may possibly be due to the fact that these relatively sicker persons required more than prophylaxis. Also, the relatively sicker (placebo treated with no scaling) showed zero change in contrast to the 10 per cent mean improvement for the entire group.

The question arises as to whether the biochemical patterns followed the clinical change. No significant change in plasma ascorbic acid levels and lingual times followed placebo therapy ( $P = 0.500$  and

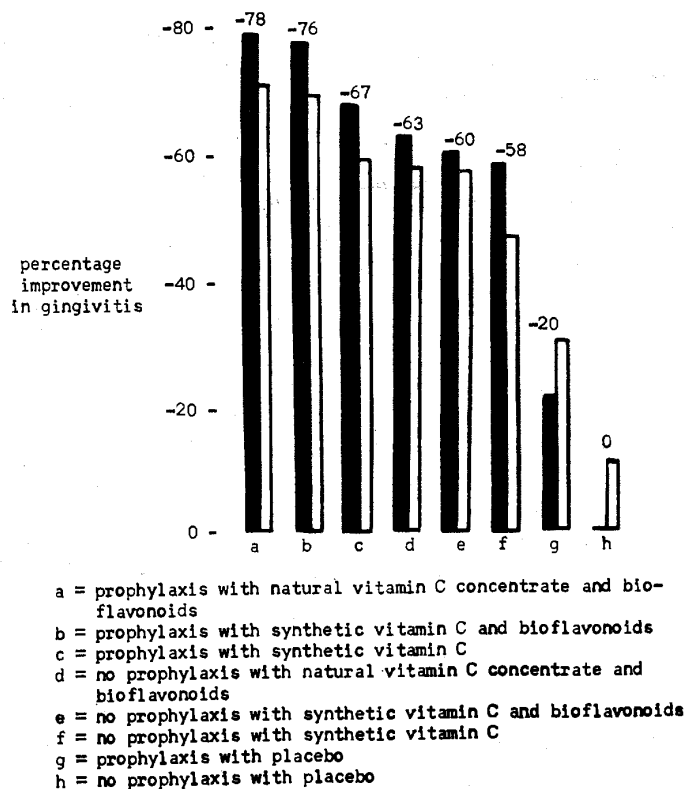


Fig. 5. Comparison of the percentage improvement in gingivitis of each group listed below (white column) with the ten subjects from each group with the poorest vitamin C levels (black columns).

> 0.100 respectively). On the other hand, there were significant increases in the plasma vitamin C levels following natural vitamin C concentrate with bioflavonoids ( $P < 0.005$ ), and synthetic vitamin C with and without bioflavonoids ( $P < 0.001$ ). Also, all vitamin-treated groups showed significant reductions in lingual time ( $P < 0.001$ ). Finally, all three supplemented groups significantly differed from the placebo-treated subjects. All of these findings parallel the clinical patterns. However, there were no other intergroup biochemical differences. This is at variance with the clinical changes. This difference is probably due to the fact that neither plasma ascorbic acid nor lingual vitamin C testing reflects bioflavonoid change.

#### SUMMARY AND CONCLUSIONS

This study was intended to evaluate the relative effects of correction of a local factor (calculus) and systemic factors (natural vitamin C concentrate versus synthetic vitamin C with and without bioflavonoids) upon periodontal pathosis (gingivitis).

An attempt was first made to correlate gingival state with calculus and ascorbic acid status. The greater correlation was found between calcareous deposits and gingivitis. Though these relationships were all statistically significant, one cannot ascribe from these data any cause and effect.

To attempt possible cause-and-effect relationships, the latter phase of the study involved the appraisal of gingival state



TABLE 12  
Gingivitis Change after Prophylaxis and Systemic Therapy

Gingi- val Grades	Natural Vitamin C Concentrates Plus Bioflavonoids		Synthetic Vitamin C Plus Bioflavonoids		Synthetic Vitamin C		Placebo	
	Initial	Final	Initial	Final	Initial	Final	Initial	Final
0	41	397	66	362	66	316	116	243
1	372	234	293	200	339	225	360	258
2	143	17	178	24	126	28	67	57
3	92	0	50	1	39	1	29	14
Total	648	648	587	587	570	570	572	572
Mean	1.4	0.4	1.3	0.4	1.2	0.5	1.0	0.7
S.D.	0.6	0.2	0.4	0.2	0.4	0.3	0.4	0.5
Percentage Change		-71		-69		-58		-30
P		<0.001		<0.001		<0.001		<0.050

after the subjects received one of eight different therapeutic regimes. The treatments involved placebo versus synthetic vitamin C with and without bioflavonoids versus natural vitamin C concentrate with

bioflavonoids with and without scaling of one half of the mouth.

Re-examination three weeks later showed a significant reduction in gingivitis in those groups given local and/or systemic ther-

TABLE 13  
Comparison of the Statistical Significance of the Therapeutic Effectiveness on  
Gingivitis of the Various Subgroups

		Placebo		Synthetic Vitamin C		Synthetic Vitamin C Plus Bioflavonoids		Natural Vitamin C Concentrate Plus Bioflavonoids	
		uns	sc	uns	sc	uns	sc	uns	sc
Placebo Group	uns	—	s	s	s	s	s	s	s
	sc	—	—	s	s	s	s	s	s
Synthetic Vitamin C Group	uns			—	ns	s	s	s	s
	sc				—	ns	s	ns	s
Synthetic Vitamin C Plus Bioflavonoids Group	uns					—	ns	ns	s
	sc						—	ns	ns
Natural Vitamin C Concentrate Plus Bioflavonoids Group	uns							—	s
	sc								—

uns = unscaled    s = significant  
sc = scaled       ns = not significant

apy. It appears, within the limits of this study, that the combined treatments yielded the best results.

We wish to take this opportunity to thank Dr. Michael J. Walsh for supplying the placebo and vitamin C and bioflavonoid preparations.

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