

Inhibitory Effect of White Wine on Gastric Myoelectrical Activity and the Role of Vagal Tone

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Although extensively investigated throughout the gastrointestinal tract, the influence of alcohol on gastric motility is still unclear. Our aim was to investigate the effect of wine on gastric myoelectrical activity and vagal activity. Ten healthy subjects were studied in two sessions with the electrogastrogram (EGG) for 30 min at baseline, 30 min after ingesting the test liquid [white wine (12.5% alcohol) or matched juice], and 60 min after a standard test meal. Spectral analysis was performed to compute EGG parameters and their postprandial changes. The vagal activity was assessed based on spectral analysis of the heart rate variability (HRV) signal derived from the ECG recording. White wine preload significantly diminished the postprandial increase in EGG dominant power compared to juice preload (1.16 ± 1.57 vs 5.48 ± 1.01 dB, $P < 0.001$). A significant decrease in vagal activity was observed after wine (23.40 ± 4.30 vs $17.43 \pm 3.40\%$, $P < 0.005$), which remained unchanged after the test meal (23.40 ± 4.30 vs $16.77 \pm 4.40\%$, $P < 0.05$). This decrease was not noted in the juice session. A correlation was established between changes after wine consumption in EGG dominant power and in the percentage of the vagal activity ($r = 0.89$, $P < 0.05$). In conclusion, white wine preload inhibits the postprandial EGG dominant power, suggesting a possible inhibition of postprandial gastric contractions. This effect may be associated with diminished vagal activity.

KEY WORDS: alcohol; electrogastrography; gastrointestinal motility; gastric slow wave; stomach; heart rate variability.

Controversial reports concerning the role of alcohol in gastric motility have been presented by numerous investigators: accelerated gastric emptying (1, 2), delayed gastric emptying (3–6), or no motility changes at all (7, 8). All studies have been performed using different methods of gastric emptying. Most of them were done invasively by inserting a tube into the stomach. It is also of note that in many of these studies, a liquid meal was used, which evaluates only

one component of the process of gastric emptying. Reasons accounting for the wide range of the results may be attributed to the diversity of variables among the studies. Different alcohol beverages were used (2, 9), with different percentages and doses (10), routes of administration (11), control groups, etc. The number of explanations matches the diversity of results, implicating neural, humoral, and local mediators. The effect of alcohol on gastric myoelectrical activity that regulates gastric motility has not been studied.

Gastric motility is regulated by gastric myoelectrical activity that is composed of slow waves and spikes. While the slow waves are associated with rhythm and regularity, the superimposed spikes are associated with antral contractions (12). The electrogastrogram

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(EGG) is a measurement of this activity from the abdominal surface. Its simultaneous recording from the gastric serosa has shown that the dominant frequency of the EGG reflects the frequency of the gastric slow waves, and its relative change in dominant power/amplitude is associated with gastric contractility (13–15). Further studies have emphasized the correlation between the EGG and impaired motility. In one study, 74% of patients with documented gastroparesis (delayed gastric emptying measured by scintigraphy) exhibited a range of abnormalities in the EGG (16). In another, all patients with abnormalities in both EGG rhythmicity and power had delayed gastric emptying (17).

The effect of alcohol on the autonomic nervous system has also been investigated using spectral analysis of heart rate variability (HRV) (18–20). Similar to its influence on the gastrointestinal tract, alcohol was shown to exert inconsistent effects on the autonomic system.

The aim of the current study was to investigate the effect of white wine on gastric myoelectrical activity and its possible correlation with cardiac vagal tone using noninvasive electrogastrography and electrocardiography. We created an almost natural setting utilizing a noninvasive method, a socially acceptable amount of wine, and a practical standard test meal. The effect of white wine was evaluated by comparison with a corresponding matched beverage, in order to eliminate as many variables as possible and get closer to the “net” effect of the alcoholic beverage chosen, the white wine. The possible mechanism for the influence of white wine on the gastrointestinal tract has been further investigated by adding a simultaneous heart rate variability measurement. Thus, the autonomic nervous system has been reflected in both myoelectrical activity and heart rate variability

MATERIALS AND METHODS

Subjects. Ten healthy volunteers (all European Americans) were studied. Asians were excluded to avoid possible different responses to wine. None had any gastrointestinal symptomatology, disease, or surgery. Average age was 29.7 ± 3 years, ranging from 23 to 56 (8 men, 2 women). The mean body mass index of the subjects was 25.3 kg/m^2 with a range from 21.3 kg/m^2 to 29.2 kg/m^2 . One woman was menopausal and the other women were studied in the follicular phase of menses to avoid the influence of hormonal changes. All volunteers consumed wine occasionally, but not more than twice a week. No subject was under

any medications during the study period. The Institutional Review Board approved the protocol and each participant signed a consent form prior to the study.

Study Design. The study was performed in two separate sessions after a fast of at least 6 hr before the recording. In one session, wine (120 ml of white wine, 12.5% by volume; Lindshein, Australia) was ingested within 3 min and in the other, the same amount of juice matched with wine in volume, pH, sodium concentration, sugar, and caloric content. The drinks were at room temperature. The two sessions were randomized and were done on two separate days within one week. The sessions were in the same period of the day. The EGG and ECG were performed for 30 min before and 30 min after the ingestion of wine or juice, and a consecutive hour following a standard meal (475 kcal, protein 21%, fat 17%, carbohydrate 61% and fiber 2 gms) (21, 22) consumed within 10 min. The measurement of the EGG took place in a quiet room where the subjects could watch regular TV programs in a supine position. They were asked to minimize their movements, and talking or reading was not allowed during the recording.

EGG Measurement and Analysis. Gastric myoelectrical activity was recorded noninvasively using surface electrogastrography. The abdominal skin was prepared with skin-prep jelly to reduce the impedance. Then, three silver–silver chloride ECG electrodes were placed on the prepared abdominal surface according to information published in previous studies, where the stomach was localized by ultrasonography (21, 22). The first electrode was set above the antrum (located 1–3 cm right of the midline, between the xiphoid process and the umbilicus), the second, 45 degrees and 5 cm above and left to the first electrode, and the third (reference) electrode at the left costal margin, horizontal to the first electrode. The electrodes were connected to a portable ambulatory device with a recording frequency of 1–18 cpm, (Digitrapper EGG, Synectics, Medical Inc., Shoreview, Minnesota, USA). The EGG signal was sampled at a rate of 1 Hz.

Motion artifacts or profound breathing movements were deleted by visual inspection before computerized spectral analysis (23). EGG data were assessed using the following parameters: (1) the dominant frequency—the frequency at which the EGG power spectrum has a peak power in the range of the 2–4 cpm; and (2) the dominant power—the power of the dominant frequency is associated with the amplitude and regularity of the EGG. The dominant frequency and power of the EGG were computed using

smoothed (or overall) power spectral analysis method (16).

ECG Measurement. The ECG was recorded using three bipolar electrodes connected to an Alice 3 Polysomnographic System (Healthdyne, Inc., Marietta, Georgia, USA). The R-R intervals were derived from the ECG. The signal of R-R intervals was interpreted and sampled at 1 Hz using MATLAB (The MATH Works, Inc., Natick, Massachusetts, USA). The spectral analysis method was used to compute percentage of low frequency (LF) component at a frequency of 0.04–0.15 Hz and the percentage of high frequency (HF) component at a frequency of 0.15–0.5 Hz (24). The percentage was defined as the ratio of the area under the curve in the power spectrum in the specified frequency band and that in the whole spectral band (0.0–0.5 Hz). The ratio (LF/HF) reflects the balance between the sympathetic activity and the parasympathetic activity.

Statistical Analysis. All data were expressed as mean \pm SEM, and $P < 0.05$ was considered as statistically significant. Student's *t* test was performed to assess the differences in EGG and HRV parameters between wine and juice. The Pearson test was used to study the correlation between gastric myoelectrical activity and heart rate variability.

RESULTS

Immediate Effect of Wine or Juice on Dominant Power of EGG. Regular gastric slow waves, which were recorded in the baseline EGG, and recordings obtained in one subject before and after wine are presented in Figure 1. The immediate effects of juice and wine ingestion were measured in the 30-min period following their ingestion, and the results were compared with the 30-min baseline recording in the fasting state. As shown in Figure 2, the dominant EGG power was increased by 3.01 ± 1.15 dB ($P < 0.03$) after juice ingestion, while after wine it showed a decrease of 0.73 ± 0.90 dB (NS). The difference between these two responses was statistically significant ($P < 0.04$).

Effect of Wine or Juice Preload on Postprandial Dominant Power of EGG. Effects of juice or wine preload on the postprandial EGG were evaluated by comparing the relative changes of the dominant power in the postprandial state versus the fasting baseline state. A significant increase (5.48 ± 1.01 dB, $P < 0.0007$) in the dominant power of the postprandial EGG was observed with juice preload. This increase was, however, not noted in the study session

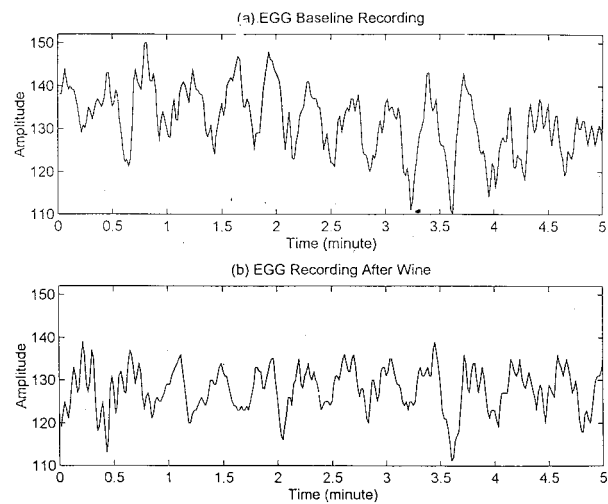


Fig 1. EGG recordings in one subject before (a) and after (b) ingestion of wine. A decrease in the amplitude of the EGG after wine is noted.

with wine preload (1.15 ± 1.57 dB, $P > 0.05$). The difference in the postprandial change in the dominant power between the two sessions was statistically significant ($P < 0.001$, see Figure 3).

Effect of Wine on Dominant Frequency of EGG. The dominant frequency was not changed after ingestion of wine (2.88 ± 0.08 vs 2.96 ± 0.12 cpm, $P > 0.6$), but increased significantly postprandially (2.88 ± 0.08 vs 3.1 ± 0.05 cpm, $P < 0.009$) similar to the expected response to food with no intervention.

Immediate and Preload Effects of Wine on High Frequency (HF) Band of HRV. A significant decrease in the HF component of the HRV was observed immediately after wine ingestion (23.40 ± 4.3 vs

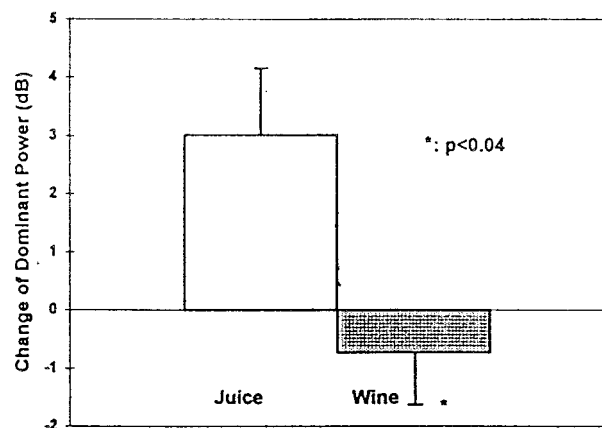


Fig 2. Immediate effect of wine and juice ingestion on the dominant power of the EGG in the fasting state. The difference between the increase in the dominant power after juice ingestion and the decrease after wine ingestion was statistically significant.

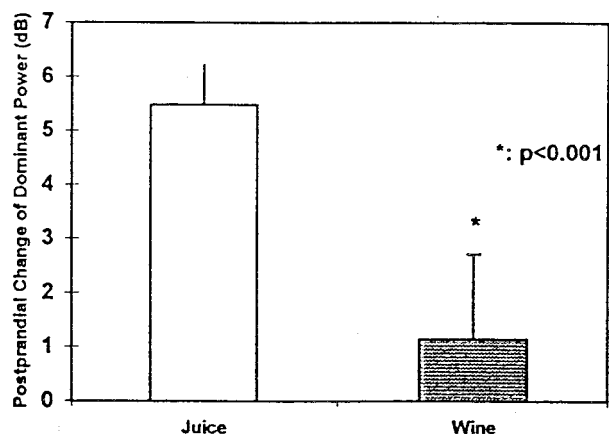


Fig 3. Postprandial changes of the dominant power of the EGG with wine and juice preload. The postprandial increase of the dominant power was significantly diminished with wine preload, in comparison with juice preload.

17.4 ± 3.40%, *P* < 0.005). A significant decrease in the HF was also noticed following the test meal, with wine preload (23.40 ± 4.30 vs 16.80 ± 4.40%, *P* < 0.05) (Figure 4).

Immediate and Preload Effects of Wine on Low Frequency/High Frequency (LF/HF) Ratio of HRV. Wine increased the LF/HF ratio, both immediately after ingestion (1.92 ± 0.51 vs 2.40 ± 0.65, *P* < 0.05) and following the test meal (1.92 ± 0.51 vs 2.85 ± 0.77, *P* < 0.04) (Figure 5).

Correlation Between EGG and HRV. A correlation was established between the postprandial changes of the EGG dominant power and the postprandial changes of the percentage of the HF component in the power spectrum of the HRV (*r* = 0.89, *P* < 0.05).

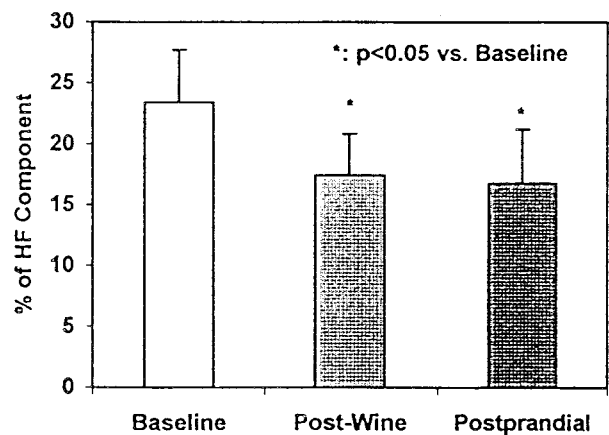


Fig 4. Immediate and preload effects of wine on the high frequency (HF) component of the HRV. Wine influences this component by a significant decrease in both instances.

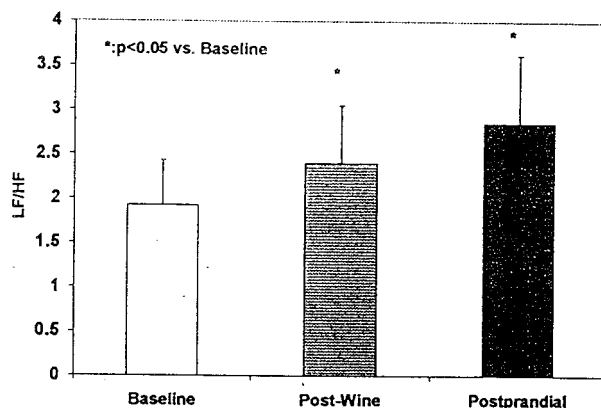


Fig 5. Immediate and preload effects of wine on the low frequency/high frequency (LF/HF) ratio of the HRV. Wine decreases both significantly.

DISCUSSION

Effect of Wine on Gastric Myoelectrical Activity.

Unlike most studies investigating the influence of alcohol on gastric function by measuring gastric emptying, we assessed gastric myoelectrical activity using the EGG. Standardization of information collected in those studies was difficult, owing to the lack of common denominators (type, percentage, and quantity of alcohol used, test meals, controls, etc.). We tried to eliminate some of the confounding variables by using a closely matched beverage and a solid test meal. In our study, the acute ingestion of white wine in healthy volunteers resulted in a characteristic pattern that was confined to a single EGG parameter: the dominant power. White wine reduced gastric myoelectrical activity, and this might be translated into the inhibition of the forceful propulsive movements of the stomach. No changes were observed in the dominant frequency or the percentage of normal slow waves, suggesting that wine does not affect the basic rhythm and regularity of the gastric slow wave.

Review of the literature classifies the acute effect of alcohol on gastric emptying according to its kinetic effect; in some, certain types of alcohol or doses are considered. Pure ethanol and whisky caused a delayed gastric emptying of both liquid and solid meals in several studies in humans (2-5, 7), as well as in animals (6, 11), while others noted an increased rate of emptying (1). White wine produced varying results ranging from no change (8), to delay (2) and even acceleration of gastric emptying (9). Beer was reported to increase the rate of emptying (9). In terms of quantity, larger amounts of alcohol were shown to cause a significant prolongation of gastric emptying

(10, 11). The diversity of observations produced numerous theories and additional studies, most targeting hormonal changes. In one study, alcohol beverages were reported to decrease in pancreatic enzyme secretion (25). The role of gastrin, a potent inhibitor of gastric contractile activity, was explored by several groups (26, 27). One of them documented a high production of gastrin following wine and beer. Another study demonstrated a delay in the release of ACh, which is known to stimulate gastric contractions and emptying (27).

Effect of Wine on Heart Rate Variability. In this study we found that both immediate and preload ingestion of white wine resulted in a significant decrease of the percentage of the high frequency components of the HRV, with a simultaneous increase of the LF/HF ratio. While the HF is believed to reflect the parasympathetic or vagal activity, the LF represents mostly the sympathetic activity (28, 29).

Several previous studies investigated the acute effect of alcohol on the autonomic nervous system, using a similar methodology, and the data were in an agreement with this current study. Koskinen et al reported a significant decrease in the HF component with a moderate amount of alcohol in healthy volunteers (18). Murata et al found an increase in the heart rate and a negative correlation with the HF change (19). The study by Van Der Borne et al showed an increase in LF/HF, attributed to an increase in the LF component (20). The chronic use of alcohol was shown to provide a more uniform pattern of reduced vagal activity (30, 31). The total damage to the autonomic nervous system in general and the parasympathetic activity in particular was found to be dose dependent (32).

Correlation Between EGG and HRV. Simultaneous recordings of the EGG and the ECG were performed in two previous studies. Watanabe et al investigated the effect of water ingestion on the EGG dominant power and the HF component of the HRV (33). Although the HF component showed no significant changes, it was positively correlated with the increase of the EGG dominant power. Kaneko et al performed a similar study and reported an increase in the EGG dominant power, a transient increase in the HF, and no changes in the LF/HF ratio (34); the correlation between the EGG power change and the change of the HF component was not reported. It is of note that both studies used a low noncaloric liquid meal and a short period of recording, which might have led to results different from the current study.

One of the most interesting results in our study is the correlation between the gastric myoelectrical activity and the cardiac rhythm. White wine induced comparable changes in the dominant power of the EGG and the HF component of the HRV. The common mechanism may involve the vagal tone that affects both organs. The postprandial increase in the dominant power of the EGG has been established in numerous studies (21, 22, 35). The role of the vagus as a mediator of this increase has also been shown in several studies using sham feeding or demonstrating the lack of this phenomenon in patients who underwent vagotomy (36). The correlation of the EGG and the HF component of the HRV may suggest that the reduction of the EGG power was mediated via the vagal mechanism.

In summary, white wine diminishes the amplitude of gastric myoelectrical activity and cardiac vagal activity. The change of gastric myoelectrical activity may be mediated via vagal mechanism.

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REFERENCES

1. Kaufman SE, Kaye MD: Effect of ethanol upon gastric emptying. *Gut* 20:688-692, 1979
2. Lenz HJ, Ferrari-Taylor J, Isenberg JJ: Wine and five percent ethanol are potent stimulants of gastric acid secretion in humans. *Gastroenterology* 85:1082-1087, 1983
3. Barboriak JJ, Meade RC: Effect of alcohol on gastric emptying in man. *Am J Clin Nutr* 23:1151-1153, 1970
4. Jian R, Corot A, Ducrot F, Jobin G, Chayvialle JA, Modigliani R: Effect of ethanol ingestion on postprandial gastric emptying and secretion, biliopancreatic secretions, and duodenal absorption in man. *Dig Dis Sci* 31:604-614, 1986
5. Horowitz M, Maddox A, Bochner M, Wishart J, Bratasiuk R, Collins P, Shearman D: Relationships between gastric emptying of solid and caloric liquid meals and alcohol absorption. *Am J Physiol* 20:G291-G298, 1989
6. Wilson CA, Bushnell D, Keshavarzian A: The effect of acute and chronic ethanol administration on gastric emptying in cats. *Dig Dis Sci* 35:444-448, 1990
7. Cooke AR: The simultaneous emptying and absorption of ethanol from the human stomach. *Am J Dig Dis* 15:449-454, 1970
8. Moore JG, Christian PE, Datz FL, Coleman RE: Effect of wine on gastric emptying in humans. *Gastroenterology* 81:1072-1075, 1981
9. Pfeiffer A, Hogel B, Kaess H: Effect of ethanol and commonly ingested alcoholic beverages on gastric emptying and gastrointestinal transit. *Clin Invest* 70:487-491, 1992

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10. Mushambi MC, Bailey SM, Trotter TN, Chadd GD, Rowbotham DJ: Effect of alcohol on gastric emptying in volunteers. *Br J Anaesth* 71:674–676, 1993
11. Knight LC, Maurer AH, Wikander R, Kreysky B, Malmud LS, Fisher RS: Effect of ethyl alcohol on motor function in canine stomach. *Am J Physiol* 262:G223–G230, 1992
12. Hasler WH: The physiology of gastric motility and gastric emptying. *In* Textbook of Gastroenterology, 2nd ed. T Yamada (ed). JB Philadelphia, Lippincott Company, 1995.
13. Smout AJPM, van der Schee EJ, Garshuis JL: What is measured in electrogastrography? *Dig Dis Sci* 25:179–187, 1980
14. Brown BH, Smallwood RH, Duthie HL, Stoddard CJ: Intestinal smooth muscle electrical potentials recorded from the surface electrodes. *Med Biol Eng* 13:97–102, 1975
15. Hamilton JW, Bellhasene B, Reichelderfer M, Webster JG, Bass P: Human electrogastrograms: Comparison of surface and mucosal recordings. *Dig Dis Sci* 31:33–39, 1986
16. Chen JDZ, McCallum RW: Gastric slow wave abnormalities in patients with gastroparesis. *Am J Gastroenterol.* 87(4):477–482, 1992
17. Chen JDZ, Lin Z, Pan J, McCallum RW: Abnormal gastric myoelectrical activity and delayed gastric emptying in patients with symptoms suggestive of gastroparesis. *Dig Dis Sci* 41(8):1538–1545, 1996
18. Koskinen P, Virolainen J, Kupari M: Acute alcohol intake decreases short-term heart rate variability in healthy subjects. *Clin Sci* 87:225–230, 1994
19. Murata K, Araki S, Yokoyama K, Sata F, Yamashita K, Ono Y: Autonomic neurotoxicity of alcohol assessed by heart rate variability. *J Auton Nerv Syst* 48:105–111, 1994
20. Van Der Borne P, Mark AL, Montano N, Mion N, Somers VK: Effects of alcohol on sympathetic activity, hemodynamics, and chemoreflex sensitivity. *Hypertension* 29(6):1278–1283, 1997
21. Levanon D, Zhang M, Orr WC, Chen JDZ: Effects of meal volume and composition on gastric myoelectrical activity. *Am J Physiol* 274:G430–G434, 1998
22. Levanon D, Zhang M, Chen JDZ: Efficiency and efficacy of the electrogastrogram. *Dig Dis Sci* 43:1023–1030, 1998
23. Liang J, Cheung JY, Chen JDZ: Detection and elimination of motion artifacts in electrogastrogram using feature analysis and neural networks. *Ann Biomed Eng* 25:850–857, 1997
24. Lu CL, Zou XP, Orr WC, Chen JDZ: Postprandial changes of sympathovagal balance measured by heart rate variability. *Dig Dis Sci* 44:857–861, 1999
25. Hajnal F, Flores MC, Radly S, Valensuala JE: Effect of alcohol and alcoholic beverages on meal-stimulated pancreatic secretion in humans. *Gastroenterology* 98(1):191–196, 1990
26. MacGregor IL, Zealous DW, Martin PM: Effect of pentagastrin infusion on gastric emptying rate of solid food in man. *Am J Dig Dis* 23:72–75, 1978
27. Rivera-Calimlin L, Hartley D, Osterhout D: The effects of ethanol and pantothenic acid on brain acetylcholine synthesis. *Br J Pharmacol* 95:77–82, 1988
28. Pomeranz B, Macaulay RJB: Caudill MA, Kutz I, Adam D, Gordon D, Kilborn KM, Barger AC, Shannon DC, Cohen RJ, Benson H: Assessment of autonomic function in humans by heart rate spectral analysis. *Am J Physiol* 248:H151–H153, 1985
29. Ewing DJ: Analysis of heart rate variability and other non-invasive tests with special reference to diabetes melitus. *In* Autonomic Failure: A Textbook of Clinical Disorders of the Autonomic Nervous System, 3rd ed. R Bannister, CJ Mathias (eds). Oxford, Oxford University Press, 1992, pp 312–333
30. Duncan G, Johnson RH, Lambie DG, Whiteside EA: Evidence of vagal neuropathy in chronic alcoholics. *Lancet* 2:1053–1057, 1980
31. Barter F, Tanner AR: Autonomic neuropathy in an alcoholic population. *Postgrad Med J* 63:1033–1036, 1987
32. Monforte R, Estruch R, Valls-Sole J, Nicolas J, Villalta J, Urbano-Marquez A: Autonomic and peripheral neuropathies in patients with chronic alcoholism. A dose-related toxic effect of alcohol. *Arch Neurol* 52(1):45–51, 1995
33. Watanabe M, Shimada Y, Sakai S, Shibahara N, Matsuda H, Umeno K, Asanoi H, Terasawa K: Effects of water ingestion on gastric electrical activity and heart rate variability in healthy human subjects. *J Auton Nerv Syst* 58(1–2):44–50, 1996
34. Kaneko H, Sakakibara M, Mitsuma T, Morise K: Possibility of postprandial electrogastrography for evaluating vagal/nonvagal cholinergic activity in humans, through simultaneous analysis of postprandial heart rate variability and serum immunoreactive hormone levels. *Am J Gastroenterol* 90(4):603–609, 1995
35. Koch KL, Stern RM: The relationship between the cutaneously recorded electrogastrogram and antral contractions in man. *In* Electro-gastrography: Methodology, Validation and Applications. RM Stern, KL Koch, (eds). New York, Praeger. 1985, pp 116–131
36. Stern RM, Crawford HE, Stewart WR, Vasey MW, Koch KL: Sham feeding: Cephalic–vagal influences on gastric myoelectrical activity. *Dig Dis Sci* 34:521–527, 1989