Acupuncture inhibits vagal gastric acid secretion stimulated by sham feeding in healthy subjects

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Abstract
In a prospective randomised study, the effect of acupuncture on sham feeding stimulated gastric acid secretion was investigated. In eight healthy volunteers (five men, three women, mean (SEM) age 26·3 (4·7) years) various methods of acupuncture were performed. Apart from the sham procedure, the acupuncture was performed at the classic acupuncture points. Electroacupuncture reduced gastric acid secretion expressed as median (range) significantly during the first 30 minute period to 1·6 (0·5–5·2) mmol compared with 3·8 (2·3–14·5) mmol (p<0·05) during control period (sham feeding without acupuncture). Inhibition of gastric acid secretion by electro-acupuncture was also significant during the second 30 minute period (0·2 (0·5–6) v 3·6 (0·3–9·1) mmol; p<0·05) and for peak acid output (0·8 (0·2–5·1) v 7·6 (3·4–12·1) mmol; p<0·05). Transcutaneous electrical nerve stimulation also resulted in significant reduction of gastric acid secretion during the first 30 minute period (1·0 (0·3–6·6) mmol v 3·8 (2·3–14·5) mmol; p<0·05), and peak acid output (3·6 (1·2–12·0) v 7·6 (3·4–12·1) mmol; p<0·05). The classic needle acupuncture, laser acupuncture, and sham acupuncture had no significant effect on gastric acid secretion. This study shows firstly that in healthy volunteers, only the versions of acupuncture using more pronounced stimulation (electroacupuncture, transcutaneous electrical nerve stimulation), but not those with only mild stimulation of the nerves (classic needle acupuncture, laser acupuncture), and secondly only acupuncture performed at defined points lead to significant reduction in gastric acid secretion.

Methods
Eight healthy subjects (five men, three women, mean (SEM) age 26·3 (4·7) years) took part in the study. As one participant failed to complete the study, the data of seven volunteers were considered for evaluation. Different methods of acupuncture used included needle acupuncture, electroacupuncture, laser acupuncture, and transcutaneous electric nerve stimulation. Apart from sham acupuncture the acupuncture was performed at the classic acupuncture points. For the control period sham feeding without acupuncture was given in all subjects in the study.

The subjects gave their written informed consent, and the study protocol was accepted by the ethics commission of the Medical University Hospital of Erlangen.

Analysis of gastric acid secretion
Gastric acid secretion analysis was started after an overnight fasting period of at least 12 hours. The placing of the gastric tube (outside diameter (OD) 5 mm) was verified by a radiological check on the correct position of its tip in the

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lower pole of the stomach. Mechanical aspiration of the gastric acid secretion using an intermittent vacuum of 120 mbar (Hico-Gastrovac 271, Hirtz & Co, Cologne, Germany) was started after an adaptation phase. The gastric juice was collected in 15 minutes periods.

Titration of the gastric juice was performed in 10 ml samples with one tenth normal solution NaOH until the indicator (phenol red) changed colour at a pH of 7-4.

CORRECTION FOR LOSS OF VOLUME
Phenol red was infused continuously into the stomach through a second tube (OD 2 mm). The opening of the second tube was located 2 cm proximal to the most proximal aspiration opening of the gastric tube. Phenol red solution (40 mg/100 ml) was instilled at 30 ml/h by an automatic infusion device (Perfusor V, Braun Melsungen, Germany).

The extinction was determined in the aspirate (0-5 ml aspirate to which 5-5 ml double distilled water was added), and 1 ml borate buffer (61:83 boric acid in 1000 ml double distilled water, adjustment of the pH to 9-2 with normal solution NaOH) at wavelengths of 520 nm, 560 nm, and 600 nm. The total extinction of the aspirated secretion (E-pra) was calculated using the formula:

\[ E_{-pra} = E_{-520} - \frac{(E_{-520} + E_{-600})}{2} \]

The correction factor \( F \) is calculated from:

\[ F = \frac{V_{-inst} \times E_{-pri}}{(V_{-asp} \times E_{-pra})} \]

Where \( V_{-inst} \) = instilled volume; \( E_{-pri} \) = extinction of instilled volume; \( V_{-asp} \) = aspirated volume; \( E_{-pra} \) = extinction of aspirated volume.

By multiplying the titrated acid value by the correction factor, \( F \), the loss corrected acid secretion is obtained.

STIMULATION OF GASTRIC ACID SECRETION BY SHAM FEEDING
Modified sham feeding was carried out using the 'chew and spit' technique. For stimulation of gastric acid secretion, the test subjects were given white bread with boiled ham to eat; after thorough chewing, the contents of the mouth were spat out and the mouth rinsed with water. Before the study was started, this procedure was used at least five times by each test subject, and the reproducibility of the acid figures obtained was established. During the test period, all subjects had reproducible figures, with a scatter of <10 per cent.

ACUPUNCTURE
Acupuncture was performed in each case by one of the authors (RV), who was an instructor with the German Medical Society for Acupuncture. The acupuncture points were used that in the Chinese acupuncture scheme are stimulated to treat diseases of the stomach,13,15, bladder 21 (lateral to the lower edge of the spiny process of the 12th thoracic vertebra); stomach 36 (lateral to the edge of the tibia below the knee joint gap); and knee joint 12 (unpaired, in the middle of the line joining the tip of the sternum and the umbilicus).

In each case, the acupuncture was maintained for 30 minutes.

In the case of classic acupuncture and acupuncture with stimulation by electric current, Japanese disposable acupuncture needles (diameter 0-3 mm) were inserted into the skin to varying depths (stomach 36 3-4 cm deep, bladder 21 1-3 cm deep, knee joint 12 to a depth of 1 cm, then 2-3 cm caudal).

For classic acupuncture, the needles are rotated or wiggled every 10 minutes for 10 seconds. For electrostimulation, the needles were connected to an electrostimulation device (AKU-IMPULS, Pitterling) and stimulated at a period relaxation oscillation (5 Hz) at a current intensity that produced a 'De-Qi' sensation at the acupuncture point (soreness, numbness or distension) showing correct acupuncture technique. To compensate for adaptation, the intensity was increased every 10 minutes.

For laser acupuncture, a laser acupuncture device (AKUPLAS MFL, MBB, Munich) operating at a frequency of 20 Hz was used; depth of penetration of the laser beam: 0-8-1.5 cm.

Transcutaneous electrical nerve stimulation was performed with the aid of adhesive electrodes (surface area about 4 cm²), and the stimulation device (Tensi Pitterling) with bipolar impulses (3 Hz), the intensity being increased until a distinctly discernible tingling sensation was produced.

Sham acupuncture was carried out at points for which no acupuncture effect has been described (medial to the insertion of the deltoid muscle, caudal to the iliac crest in the glutaeus maximus muscle), and electrical stimulation was applied.

STUDY PROTOCOL
The test periods for needle acupuncture, electroacupuncture, transcutaneous electrical nerve stimulation, laser acupuncture, sham acupuncture, and the control period were performed in randomised sequence. The test subjects were not informed about the nature of the specific examination.

In each case, the experiments were started at 7.15 am, so that the procedures, including acupuncture, always took place in the same period of time. After placement of the gastric tube, the contents of the stomach were aspirated with the patient in a semireclining position, in seven 15 minute periods, that is, over a total period of 105 minutes. The first 15 minute sample was discarded. After another 30 minutes, stimulation of the gastric acid secretion by sham feeding for a 15 minute period was performed, aspiration for the stimulation period started with sham feeding. Stimulated
acid output included the two 30 minute periods after sham feeding began, peak acid output represented the peak in any 30 minute period from 45–105 minutes.16. Acupuncture began with insertion of the needles five minutes before the start of the sham feeding, and was performed for a total of 30 minutes.

STATISTICS
As basal and sham feeding acid output rates show a notoriously skewed distribution Wilcoxon’s matched pairs signed rank test was used for comparison. The results are given as medians and their ranges. Calculation was based on 30 minute samples. Statistical significance was assumed at the 5% value.

Results
Gastric acid output stimulated by sham feeding was 3·8 (2·3–14·5) mmol during the first 30 minute period of control period where no acupuncture was performed. This was not statistically different from gastric acid output during sham acupuncture (7·4 (3·1–20·8) mmol), laser acupuncture (6·0 (0–9·6) mmol), and needle acupuncture (8·2 (0·8–28·9) mmol). A significant reduction was seen during electroacupuncture (1·6 (0·5–2·2) mmol) (p<0·05) and during transcutaneous nerve stimulation (1·0 (0–3·6) mmol) (p<0·05) (Figure A).

During the second 30 minute period acid output was significantly (p<0·05) inhibited only by electroacupuncture (0·2 (0–5·6) mmol) compared with the control period (3·6 (0·3–9·6) mmol). There was no significant influence on gastric acid secretion effected by sham acupuncture (4·8 (2·2–23·5) mmol), laser acupuncture (2·2 (0–13·4) mmol), needle acupuncture (3·5 (1·6–14·7) mmol) or by transcutaneous electric nerve stimulation (6·0 (0–25·7) mmol) (Figure B).

Peak acid output was significantly reduced by electroacupuncture (0·8 (0·2–5·1) mmol/h; p<0·05) and transcutaneous nerve stimulation (3·6 (1·2–12·0) mmol/h; p<0·05). Peak acid output during the control period was 7·6 (3·4–12·1) mmol/h, during sham acupuncture 9·1 (5·5–12·9) mmol/h, during laser acupuncture 5·5 (0·7–9·0) mmol/h, and during needle acupuncture 7·3 (2·4–25·3) mmol/h (Figure C).

The gastric acid secretion stimulated by sham feeding was significantly reduced by two forms of acupuncture – electroacupuncture and transcutaneous electric nerve stimulation. The inhibitory effect was more pronounced during the first 30 minute period, where median values of gastric acid secretion obtained during control period were reduced to 42·1% by electroacupuncture and to 26·3% by transcutaneous electric nerve stimulation.

The effect of electroacupuncture lasted longer than transcutaneous nerve stimulation and was significant also during the second 30 minute period. Two further forms of acupuncture – classic needle acupuncture and laser acupuncture – did not prove effective in the acute experiment with healthy test subjects. Also, sham acupuncture entailing the insertion of needles at sites not defined as effective acupuncture points did not have any influence on gastric acid secretion stimulated by sham feeding.

Discussion
In this study, sham feeding stimulated gastric acid secretion was significantly reduced by electroacupuncture and transcutaneous electric nerve stimulation, the two forms of acupuncture that entail the most potent stimulation. Classic needle acupuncture, laser acupuncture, and sham acupuncture had no influence on gastric acid secretion. In previous studies effects were achieved by electroacupuncture12 and classic needle acupuncture,10 and in animal experiments by electroacupuncture.13 The same acupuncture points were used as in this study.10 12 13 17

In this study the influence of electroacupuncture on gastric acid secretion is not explained merely by inserting the needles or by the electrostimulatory effect, as these procedures applied at other sites (sham acupuncture) led to no reduction in acid output. Although gastric acid secretion can be significantly inhibited, for example by hypnosis,18 in this study, ‘suggestion’ can be ruled out as an important factor, as in the
remaining acupuncture procedures with more mild stimulation, no significant action on acid secretion was found.

Li et al.\textsuperscript{12} stimulated gastric acid secretion by sham feeding in healthy subjects and reported that electroacupuncture reduced basal acid output significantly by 23.8 per cent and maximal acid output by 71 per cent.

Also in duodenal ulcer patients, classic needle acupuncture significantly reduced gastric acid secretion with a reduction rate of about 45–75\%.\textsuperscript{10} In earlier experiments,\textsuperscript{13} it was found that with classic needle acupuncture, and more noticeably with electroacupuncture, a significant influence on gastric acid secretion in the dog was obtained: in addition to the reduction in acid secretion, an increase in bicarbonate secretion was also found. Anti-muscarinic agents, but also local anaesthetics applied to the insertion points of the acupuncture needles abrogated this effect, so that the authors concluded that the effect of acupuncture was mediated by somatic afferent visceral fibres.

Acupuncture is considered to have some value in the treatment of ulcers. In four patients with duodenal ulcers resistant to medical treatment healing was seen after six to seven acupuncture sessions.\textsuperscript{11} Duodenal ulcer patients became complaint free after acupuncture.\textsuperscript{10} As the number of patients in both studies is limited, the effect of acupuncture on ulcer healing has yet to be proved.

The stimulation of gastric acid secretion by sham feeding entails mainly the central vagal component of gastric acid stimulation.\textsuperscript{19} Acupuncture seems to reduce gastric acid secretion by afferent visceral fibres.\textsuperscript{13} At present there can only be speculation about the mechanisms produced by acupuncture, as regulation of gastric acid secretion is complex including neural, hormonal, and paracrine mechanisms.\textsuperscript{20} It cannot be excluded that at least part of the effect of acupuncture on gastric secretion also entails stimulation of bicarbonate secretion.\textsuperscript{13}

At present, the treatment available for reducing gastric acid secretion is based on pharmacological blocking of parietal cell receptors for histamine, muscarine, gastrin, and inhibition of H+/K+-ATPase.\textsuperscript{20} The possibility of inhibiting gastric acid secretion by electroacupuncture, as shown in this study, offers an alternative to drug related reduction of gastric acid secretion. Our results could give a rational basis for peptic ulcer treatment only if the effect of electroacupuncture would last for a period longer than two hours. Furthermore, the effect of electroacupuncture on ulcer healing has yet to be proved in controlled trials.

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