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REVIEW

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Systematic review: Validity, reliability, and diagnostic accuracy of the electrogustometer

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Abstract

Objective: What are the electrogustometer's (EGM) validity, reliability, and diagnostic accuracy in assessing taste sensation in adults compared to other taste tests?

Data Sources: PubMed Medline, Elseviers's Embase, and the six databases of Cochrane Library.

Methods: We conducted a systematic search on December 20, 2022, consisting of synonyms for EGM. We considered randomized controlled trials and observational studies with original data for inclusion if they included adults who underwent electrogustometry. Articles were excluded if no analysis regarding validity, reliability, or diagnostic accuracy had been performed or if these analyses could not be performed with the published data.

Results: Nineteen articles discussing 18 studies were included for data extraction. The included studies carry a high risk of bias. Overall, the association between a variety of reference taste tests and EGM was moderate or weak with correlation coefficients ranging from -0.51 to 0.40 with one outlier of -0.74 found in one study correlating EGM and taste solutions. Test-retest reliability was good with reported correlation coefficients between 0.78 and 1.0. The sensitivity, specificity, PPV, and NPV of EGM in identifying abnormal taste function varied widely between the four studies on diagnostic accuracy.

Conclusion: The included studies in this review lack the required standards regarding study design to draw firm conclusions about the validity, reliability, and diagnostic accuracy of the EGM. Future research is needed to assess these measurement properties. Based on the reported results, we would not recommend using the EGM as a screening test for taste disturbance in clinical practice.

Level of Evidence: NA.

KEYWORDS

diagnostic accuracy, electrogustometry, reliability, systematic review, taste sensation, validity

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1 | INTRODUCTION

Since 1958, the electrogustometer (EGM) has been used as a clinical tool for measuring electrical taste thresholds of the three gustatory nerves: the chorda tympani, glossopharyngeal and greater petrosal nerves.^{1,2} Over the years, several EGM models have been developed. All models produce microamp-level anodic currents to the tongue, causing sour or metallic taste sensations. There are two hypotheses regarding the mechanism of electrogustometry. Based on the electrolyte chemical hypothesis, the anodic current creates an acidic solution through electrolysis, which innervates the taste receptor cells. Based on the direct effect hypothesis, the electrical taste arises from the anodic current action on nerve fibers or taste receptor cells.³

The methods for assessing patients with taste disorders have not been fully standardized.⁴ Quantitative assessment of gustatory function with the taste strip test (TST) and filter paper disk (FPD) are established methods.⁵ The advantages of using an EGM in comparison to the TST and FPD are that only a short test time is required and its ability to detect subtle, subclinical taste disorders. In our center, the test time of the EGM is 6 min, while the TST takes 12 min to test both sides of the tongue.

Over the years, several studies have been published on the validity, reliability, and diagnostic accuracy of the EGM.^{3,4,6–22} Validity is the degree to which the EGM measures taste sensation. Reliability is the degree to which an electrical taste threshold is the same for patients who have not changed for repeated measurements under the same conditions.^{23,24} Diagnostic accuracy is the ability of the EGM to distinguish between patients with and without taste loss.²⁵ To date, no systematic review of the validity, reliability, or diagnostic accuracy of the EGM has been published. To create an up-to-date clear overview of the published literature, we aim to systematically assess the validity, reliability, and diagnostic accuracy of the EGM in assessing taste sensation in adults.

2 | METHODS

2.1 | Search strategy and study selection

We conducted a systematic search in PubMed Medline, Elseviers's Embase, and the six databases of Cochrane Library with the assistance of a clinical librarian on December 20, 2022. We used a broad search query consisting of synonyms for EGM and reported our findings according to the PRISMA guideline.²⁶ The review has not been registered and no protocol has been prepared. See Appendix S1 for the full search strategy. No MeSH terms exist for EGM. No restrictions on publication year and publication status were applied. Two independent authors (EB and MD) excluded duplicates and performed title and abstract screening. See Figure 1 for the selection criteria. We considered randomized controlled trials and observational studies (retrospective and prospective cohort studies and case series [$n \ge 10$]) for inclusion if they included adults (≥ 18 years old) who underwent electrogustometry. Articles with original data were included for full text

screening. Reviews, animal or laboratory studies, conference abstracts, poster presentations, letters, and case reports with <10 cases were excluded. Articles were also excluded if no analyses regarding validity, reliability, or diagnostic accuracy had been performed or if these analyses could not be performed with the published data. There were no restrictions regarding EGM model, stimulation method or study population. If no full text was available online or through our universities' libraries, the authors of the studies were contacted to retrieve the full text of the article. The full text screening was also performed by two independent authors (EB and DF) using the same selection criteria. Any differences were discussed until consensus was reached. Reference lists of included articles were used to identify possible additional relevant articles.

2.2 | Study outcome

The primary outcome was the construct validity of the EGM. Construct validity is the degree to which the scores of a measurement instrument are related to other measurement instruments that measure similar qualities. Therefore, the construct validity of the EGM is the degree to which the electrical taste threshold of the EGM is consistent with other taste instruments, such as TST and FPD.²⁴ The secondary outcomes were the reliability and diagnostic accuracy of the EGM. Reliability is the degree to which an electrical taste threshold is the same for patients who have not changed for repeated measurements under the same conditions.^{23,24} Diagnostic accuracy is the ability of the EGM to distinguish between patients with and without taste loss.²⁵

2.3 | Data extraction

Original data from included articles were extracted by two independent authors (EB and DF). The following data were extracted: country in which the study was performed, total number of participants, gender, age, study population, EGM model, stimulation method, stimulated nerve(s), duration of the stimulation, other taste instruments that were used and the measurement properties construct validity, reliability, and diagnostic accuracy. Construct validity and reliability are measured using correlation coefficients. A correlation coefficient between 0.1 and 0.3 was interpreted as weak, a correlation coefficient between 0.4 and 0.6 as moderate, a correlation between 0.7 and 0.9 as strong and a correlation of 1 as a perfect correlation.²⁷ Correlation coefficients were positive or negative. The lower the electrical taste threshold measured with the EGM, the better the taste sensation. If the reference test scale was the opposite of the EGM test scale (the higher the reference test score, the lower the electrical taste threshold), the correlation coefficient was negative. If the reference test scale was comparable to the EGM test scale (the lower the score of the reference test, the lower the electrical taste threshold), the correlation coefficient was positive. Diagnostic accuracy is measured using sensitivity, specificity, negative predictive value (NPV) and

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FIGURE 1 Prisma 2020 flow diagram.

positive predictive value (PPV). If data was not provided in the text of the article, it was extracted from figures if possible. If no measurement properties were reported, they were calculated.

2.4 | Quality assessment

Included studies were critically appraised by two independent authors (EB and DF). Predefined criteria from the COnsensus-based Standards for the selection of health status Measurements INstruments (COSMIN) tool were used to assess the quality of the studies on reliability (Appendix S2).^{28,29} Each box was scored on a four-point rating scale (i.e., "very good," "adequate," "doubtful," or "inadequate"). The overall score, the quality of the study on reliability, was the lowest score given for a box. This method is called the worst-score-counts method.^{28,29} The agreement on the scores was unanimous. The study designs and measurements used to quantify the validity and diagnostic accuracy of the EGM differed widely between the included studies. Therefore, we could not find one suitable tool to assess the quality of these studies. However, we assessed whether the studies formulated hypothesis beforehand, had an appropriate sample size (>100 patients) to assess validity and whether there was any review bias.

3 | RESULTS

3.1 | Search and study selection

The electronic search of articles yielded 4504 results. After removing duplicates. 3197 articles were screened on title and abstract. Title and abstract screening resulted in 56 articles. During full text screening, 37 articles were excluded: one article was a case report with <10 cases³⁰; one article only included children³¹; one article was a review³²; the language of 11 articles was Japanese (five articles), Polish (four articles), or Russian (two articles)³³⁻⁴³: 23 articles did not discuss validity, reliability, or diagnostic accuracy of the EGM or did not report values that allowed us to calculate correlation coefficients, sensitivity, specificity, PPV, or NPV^{44–66}; Nineteen articles were eligible for inclusion in the review.^{3,4,6–22} Two articles from the same medical center and the same author evaluated the same patient cohort, so the results of these articles were combined.^{14,15} We included the articles by Sakaguchi et al. and Saito et al., even though children were included in the study, because the mean age of the included patients was 39 and 42 years, respectively, and therefore we assumed only a small portion of the study population consisted of (young) children.^{10,17}

3.2 | Risk of bias assessment

3.2.1 | Reliability studies

Six studies were included that assessed test-retest reliability of the EGM.^{11,16–18,21,22} The majority of studies were not designed as a reliability study, which means that the methods used to assess test-retest reliability were poorly described (see Table 1).^{11,17,18,21,22} Only the study by Nicolaescu et al. was designed as a reliability study.¹⁶ Therefore, the overall quality of the included studies on reliability was rated as doubtful or inadequate, mostly due to the lack of information. All studies described the time interval between both tests, but none of the studies described if patients were clinically stable between the two measurements.^{11,16-18,21,22} Nicolaescu et al. and Nilsson described the measurement protocol in detail for the first and the second measurement and the measurement conditions were similar for the repeated

measurement.^{16,22} The other studies did not describe their protocol in detail, but we assume measurement conditions were similar for the repeated measures.^{11,17,18,21} None of the studies described whether professionals administered the measurements without knowledge of scores or values of the first measurement. Therefore, box four, administration of measurements, was rated as doubtful for all studies. Box five, assignment of the scores, does not apply to studies on reliability of the EGM, because no assignment of scores is necessary when using an EGM.^{11,16–18,21,22} The electrical taste threshold is determined during the measurement. Berling et al. used ANOVA's and Nilsson used t-tests, which are important flaws in the statistical method, as these statistical methods are insufficient to test the reliability of the EGM.^{11,22} The other four studies measured correlation coefficients, but none of the studies provided evidence that there was no systematic difference between measurements.^{16–18,21} Only the study by Saito et al. calculated a Kappa. but the weighting scheme was not described.¹⁷ Grant et al. did not report whether the correlation coefficient was statistically significant, which is also a major flaw in the statistical method.²¹

3.2.2 | Validity and diagnostic accuracy studies

Although we did not use a validated instrument to assess the risk of bias in the studies concerning validity and diagnostic accuracy, we think these studies carry a high risk of bias.^{3,4,6-15,17-20} First of all, only the study by Ellegård et al. was designed as a validity and diagnostic accuracy study.^{14,15} The other included studies retrospectively examined the validity and diagnostic accuracy of the EGM.^{3,4,6-13,17-20} No hypotheses were therefore formulated in advance about the expected relationships or differences between the EGM and the reference tests. Second, it is unknown for all studies whether the EGM assessors were blinded and whether they had access to the results of the reference test. Also, the number of assessors was not mentioned in any of the studies.^{3,4,6-15,17-20} Lastly, seven out of 12 studies on validity of the EGM did not have a sample size of more than 100 participants.^{4,6,9,11,18-20}

3.3 | Characteristics of included studies

Table 2 shows the characteristics of the included studies. The studies were very heterogeneous with regard to study population,

TABLE 1COSMIN risk of bias toolfor reliability.

Reference (year)	1	2	3	4	5	6	7	8	9	Overall score
Berling (2011) ¹⁹	D	V	A	D	NA		1	NA	NA	
Nicolaescu (2005) ⁷	D	V	V	D	NA	D	D	NA	NA	D
Saito (2001) ⁸	D	V	A	D	NA	D	D	A	NA	D
Murphy (1995) ⁹	D	V	A	D	NA	D	D	NA	NA	D
Grant (1987) ¹²	D	V	A	D	NA	l.	I.	NA	NA	l.
Nilsson (1977) ¹³	D	V	V	D	NA		I.	NA	NA	1

Abbreviations: V, very good; A, adequate; D, doubtful; I, inadequate; NA, not applicable.

Chara	cteristics of the in	Icluded studies.		Aco voar					Ctimulation		
Stud	y design	Country	N (male %)	Age, years (mean [range])	Study population	EGM model	Stimulation method	Stimulated o nerve(s) (sumulation duration (ms)	Reference tests	Measurement properties
Pro	spective ohort study	Greece	132 (58)	NR	Healthy	TR-06, Rion Co., Tokyo, Japan	Staircase protocol	CT, N IX, major petrosal nerve	500	TST	Validity (construct)
Rei	trospective case series	South Korea	46 (24)	53 (NR)	Taste disorder with and without burning mouth symptoms	NR	Ascending protocol	CT, N IX, major I petrosal nerve	NR	Subjective taste ability	Validity (construct)
Ϋ́.	ospective cohort study	South Korea	60 (50)	26 (NR)	Healthy	EGM-IIB, Nagashima Medical Instrument Co., Tokyo, Japan	Ascending protocol	CT, N IX, major I petrosal nerve	NR	TST, FDP, taste solutions	Validity (construct)
Ϋ́	ospective case series	Germany	180 (37)	59 (NR)	Healthy, taste disorder, olfactory disorder	TR-06, Rion Co., Tokyo, Japan	Ascending protocol	5	500	TST, taste solutions	Validity (construct)
ā	rospective case series	Germany	121 (47)	52 (21-82)	Healthy, taste disorder	TR-06, Rion Co., Tokyo, Japan	Staircase protocol	с С	500	TST	Validity (construct)
₽.	rospective cohort study	Japan	126 (53)	39 (7-70)	Middle ear surgery	TR-06, Rion Co., Tokyo, Japan	NR	ت ت	500	Subjective taste ability	Diagnostic accuracy
Ϋ́	ospective cohort study	Germany	18 (22)	45 (29-60)	Middle ear surgery	Halle II, Haberland, Halle, Germany	Ascending protocol	ct	500	Subjective taste ability	Validity (construct)
۵.	rospective cohort study	Sweden	39 (NR)	NR (27-62)	Healthy	TR-06, Rion Co., Tokyo, Japan	Ascending protocol	с 5	2000	FPD	Validity (construct), Reliability (test- retest)
Δ.	rospective cohort study	United Kingdom	143 (NR)	NR	Early clinical signs of parkinsonism	TR-06, Rion Co., Tokyo, Japan	Staircase protocol	CT, N IX	1500	Subjective taste ability	Validity (construct)
-	Prospective case series	Japan	24 (71)	62 (NR)	Stroke with dysesthesia in the face, oral cavity or tongue (with or without taste disorder)	TR-06, Rion Co., Tokyo, Japan	Ascending protocol	ь.	2000	FPD, subjective taste ability	Diagnostic accuracy

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	Measurement properties	Diagnostic accuracy, Validity (construct)	Reliability (test- retest)	Diagnostic accuracy, Reliability (test- retest)	Validity (construct), Reliability (test - retest)	Validity (construct)	Validity (construct)	Reliability (test- retest)	Reliability (test- retest)	
	Reference tests	Subjective taste ability	AN	Subjective taste ability	Taste solutions	Taste solutions	Taste solutions	A	AN 0	A, not applicable.
	duration (ms)	200	500	200	500	5000	N	5000	1000-1500	not reported; N
	Stimulated nerve(s)	Ъ	ن	Ե	Ь	Ъ	ن	Ъ	Ե	eal nerve; NR, 1
	Stimulation method	Staircase protocol	Staircase protocol	Ascending protocol	Ascending protocol	Ascending and descending protocol	Staircase protocol	Ascending protocol	Ascending and descending protocol	, glossopharyng
	EGM model	Automated EGM, computer- controlled	TR-06, Rion Co., Tokyo, Japan	EGM-IIB, Nagashima Medical Instrument Co, Tokyo, Japan	TR-06, Rion Co., Tokyo, Japan	Programmable current source and a digital volt meter (RS components 258-041)	NR	Programmable current source and a digital volt meter (RS components 258-041)	Madson GO 70 gustometer, Madsen Electronics, Copenhagen, Denmark	chorda tympani; N IX
	Study population	Healthy, head and neck cancer treated with radiotherapy, Sjögren's syndrome, burning mouth syndrome or oral dysesthesia	Healthy	Healthy, middle ear surgery	Healthy	Mastoidectomy, middle ear surgery and tympanoplasty	Healthy, diabetes mellitus	Я	Healthy), filter paper disk; CT,
A	Age, years (mean [range])	60 (NR)	24 (NR)	42 (3-78)	73 (64-85)	43 (18-74)	44 (NR)	R	25 (NR)	strip test; FPI
	N (male %)	114 (42)	16 (44)	391 (NR)	17 (NR)	20 (45)	95 (48)	20 (NR)	31 (61)	er; TST, taste
	Country	s Sweden	NSA	Japan	USA	 United Kingdom 	e France	 United Kingdom 	Sweden	ectrogustomet
	Study design	Prospective case series	Prospective cohort study	Retrospective cohort study	Prospective cohort study	Prospective case series	Prospective case series	Prospective case series	Prospective cohort study	iple size; EGM, ele
	Reference (year)	Ellegård (2007) ^{5,6}	Nicolaescu (2005) ⁷	Saito (2001) ⁸	Murphy (1995) ⁹	Grant (1989) ¹⁰	Le Floch (1989) ¹¹	Grant (1987) ¹²	Nilsson (1977) ¹³	Abbreviations: N, sam

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TABLE 2 (Continued)

used measurements instruments and reported outcomes.^{3,4,6–22} Major differences in type of patients and the reference tests that were used, precluded pooling of data.

3.3.1 | Study population

The mean age ranged from 24 to 73 years in 14 studies.^{4,6-11,13-20,22} Only four studies included patients who underwent middle ear surgery.^{9,10,17,19} The other studies included healthy participants and/or patients with a variety of diseases and syndromes.^{3,4,6-8,11-20,22}

3.3.2 | Measurement instruments

The used reference tests were the TST, FDP, taste solutions, and subjective taste ability. Subjective taste ability was measured with a yes/no question.^{10,13,17} visual analog scale (VAS).^{9,14,15} or numeric rating scale (NRS).^{4,12} The VAS used by Guder et al. ranged from 1 (no sense of taste) to 10 (excellent sense of taste).⁹ The VAS used by Ellegård et al. ranged from 0 (no sense of taste) to 100 (excellent sense of taste).^{14,15} Park et al. asked each patient to rate their sense of taste on a 3-point NRS from not at all recognized (0) to easily recognized (2).⁴ Deeb et al. asked each patient to rate their sense of taste on a 6-point NRS from absent (1) to excellent (6).¹² The TR-06 (Rion Co., Tokyo, Japan) was the most commonly used EGM model.^{3,7,8,10-13,16,18} Seven studies used a different EGM model and two studies did not report, which model was used.^{4,6,9,14,15,17,19-22} Fifteen studies used the ascending protocol or the staircase detection protocol as the stimulation method.^{3,4,6-9,11-18,20,21} Two studies used the ascending descending protocol and one study did not report the stimulation method.^{10,19,22} During the ascending threshold protocol, an electrical stimulus in ascending microamps was applied. The rater started by applying the lowest electric current and increased it to the highest electric current. When patients identified the electrical taste, the specific current was considered to be the electrical taste threshold.^{4,6,7,9,11,13,17} Park et al., Kang et al., and Han et al. performed the ascending threshold protocol three times and the mean value was considered to be the EGM threshold.^{4,6,7} A two-alternative forced choice paradigm was used during the staircase detection protocol. Two correct answers or one incorrect answer led to a decrease or increase in stimulus intensity, which was a reversal in the staircase. The average of the staircase reversals was the electrical taste threshold.^{3,8,12,14–16,20} Nilsson and Grant et al. used the descending protocol. After electric taste detection, the current was gradually reduced until the taste detection threshold was reached.^{19,22} All studies stimulated the chorda tympani.^{3,4,6-22} Pavlidis et al., Park et al., and Kang et al. also stimulated the glossopharyngeal nerve and the major petrosal nerve.^{3,4,6} Deeb et al. also stimulated the glossopharyngeal nerve.¹² The chorda tympani carries the taste afferents of the anterior two thirds of the tongue. The glossopharyngeal nerve carries the taste afferents of the posterior one third of the tongue (circumvallate papillae). The major petrosal nerve carries the taste afferents of the soft palatal area.³ Most studies stimulated for 500 ms.^{3,7-10,14-18} Nilsson stimulated for 1000-1500 ms, Deeb et al.

stimulated for 1500 ms, Berling et al. and Etoh et al. stimulated for 2000 ms and both studies by Grant et al. stimulated for 5000 ms.^{11-13,19,21,22} Three studies did not report the duration of stimulation.^{4,6,20}

3.4 | Primary outcome

Twelve studies evaluated construct validity by measuring correlation coefficients (Table 3). We reported the correlation coefficients between the EGM threshold in the chorda tympani area of the tongue and the reference test, because all studies stimulated this nerve.^{3,4,6-9,11,12,15,18-20} Two studies did not report a correlation coefficient. They only described whether the EGM and the reference test were significantly correlated.^{8,9} Ten studies showed a significant correlation between the EGM and a reference taste test. The level of statistical significance was set at a p-value of <0.05 for all studies.^{3,4,6-8,11,12,15,19,20} There is no unequivocal answer to the question of which reference test correlates best with the EGM. Overall, the association between EGM and a reference taste test was weak or moderate (correlation coefficients ranged from -0.51 to 0.40).^{3,4,6-9,11,12,15,18-20} Only Grant et al. found a strong correlation (correlation coefficient -0.74) between the EGM and taste solutions.¹⁹ Three out of five studies using taste solutions as a reference test showed a significant correlation between the EGM and taste solutions,^{7,19,20} whereas the other two did not find a significant correlation. Three out of four included studies using TST as a reference test showed a significant correlation between the EGM and TST. Pavlidis et al. and Walliczek-Dworschak et al. found a significant correlation between the EGM and all four taste strip scores (sweet score, salt score, sour score, and bitter score). However, Walliczek-Dworschak et al. did not report the values and they found negative correlations.^{3,7,8} The correlations between the EGM and the FPD and between the EGM and subjective taste ability were generally weak with correlation coefficients around (-)0.3 and most of these were not statistically significant.^{4,6,9,11,12,14}

Pavlidis et al. also found significant correlations, with similar values, between the electrical taste threshold of the glossopharyngeal nerve and the major petrosal nerve and the sweet, salty, sour and bitter scores of the TST. Park et al. found a significant correlation, with a similar value, between the electrical taste threshold of the major petrosal nerve and the sum score of the subjective taste ability in both the healthy and burning mouth study group. The electrical taste threshold of the major petrosal nerve was only associated with the sweet score in patients with burning mouth symptoms. Kang et al. and Deeb et al. found no significant correlations between the electrical taste threshold of the other nerves and the reference tests.^{6,12}

3.5 | Secondary outcomes

3.5.1 | Test-retest reliability

Six studies evaluated test-retest reliability. The time between the first and the second measurement ranged from 1 day to 16 days.^{11,16-18,21,22} Four studies calculated a correlation coefficient

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		Correlations						
Test	Reference (year)	Sweet score	Salt score	Sour score	Bitter score	Umami score	Sum score	
TST	Pavlidis (2021) ²	0.40	0.40	0.25	0.40	-	-	
	Kang (2020) ¹⁴	NC	NC	NC	NC	-	NC	
	Han (2018) ¹⁵							
	Healthy	NC	NC	NC	NC	NC	NC	
	Taste disorder	0.29	NC	NC	NC	NC	NC	
	Olfactory disorder	NC	NC	NC	NC	NC	NC	
	Walliczek-Dworschak (2017) ¹⁶	Negative correlation	Negative correlation	Negative correlation	Negative correlation	-	Negative correlation	
FPD	Kang (2020) ¹⁴	NC	NC	NC	NC	-	0.38	
	Berling (2011) ¹⁹	0.29	0.27	0.31	0.24	-	-	
Taste solutions	Kang (2020) ¹⁴	NC	NC	NC	NC	-	NC	
	Han (2018) ¹⁵							
	Healthy	NC	NC	- 0.34	- 0.27	NC	NC	
	Taste disorder	0.29	NC	NC	NC	-0.35	-0.38	
	Olfactory disorder	NC	NC	NC	NC	NC	NC	
	Murphy (1995) ⁹	-	-0.22	0.28	-	-	-	
	Grant (1989) ¹⁰	-	-	-	-	-	- 0.74 ª	
	Le Floch (1989) ¹¹	-	-	-	-	-	-0.51	
Subjective	Park (2021) ³							
taste ability	Burning mouth (right)	-0.38	-0.30	-0.39	-0.19	-	-0.48	
	Burning mouth (left)	-0.15	-0.14	-0.27	0.00	-	-0.28	
	No burning mouth (right)	-0.30	-0.39	-0.31	-0.25	-	-0.32	
	No burning mouth (left)	-0.37	- 0.41	- 0.40	-0.39	-	-0.44	
	Guder (2012) ¹⁷	-	-	-	-	-	NC	
	Deeb (2010) ²⁰	-	-	-	-	-	-0.35	
	Ellegård (2007) ⁶	-0.08	-0.19	-0.15	-0.12	-	-	

Note: p-value <0.05 is statistically significant and displayed in bold.

Abbreviations: -, not reported; FPD, filter paper disk; NC, no correlation; TST, taste strip test.

^aCalculated.

TABLE 4 Results—Diagnostic accuracy.

Reference (year)	Prevalence (%)	Sensitivity (%)	Specificity (%)	PPV (%)	NPV (%)
Sakaguchi (2013) ¹⁸					
Subjective taste ability	33 ^a	97 ^a	68 ^a	60 ^a	98 ^a
Etoh (2008) ²¹					
FPD	75 ^a	39 ^a	83 ^a	88 ^a	32 ^a
Subjective taste ability	46 ^a	36 ^a	69 ^a	50 ^a	56ª
Ellegård. (2007) ⁵					
Subjective taste ability		33 (4-71)	80 (71-87)	9 (1-28)	95 (88–99)
Sweet		0 (0-37)	77 (68-85)	0 (0-15)	91 (82-96)
Salt		40 (12-74)	81 (72-88)	17 (5–39)	93 (85–98)
Sour		30 (7–65)	80 (71-87)	13 (3–34)	92 (84–97)
Bitter					
Saito (2001) ⁸					
Subjective taste ability	7 ^a	80 ^a	91 ^a	40 ^a	98 ^a

Abbreviations: NPV, negative predictive value; PPV, positive predictive value. ^aCalculated.

between the first and second measurement. The first and second measurement were strongly associated (correlation coefficient ranged from 0.78 to 1.00).^{16–18,21} The other two included studies did not report correlation coefficients, but found no significant threshold shift between the first and second measurement.^{11,22} The level of statistical significance was also set at a *p*-value of <0.05 for all studies.^{11,16–18,21}

3.5.2 | Diagnostic accuracy

Table 4 shows the prevalence, sensitivity, specificity, PPV, and NPV of the four included studies. The study population varied (see Table 2) and as a result the prevalence of taste loss ranged from 8% to 33%.^{10,13,14,17} All studies used subjective taste ability as the reference test. Sensitivity ranged from 33% to 97%. Specificity was generally high, ranging between 68% and 91%. The PPV ranged from 0 to 88% with higher PPVs in study populations with a higher prevalence of taste disturbances. The NPVs were high in three studies with values ranging between 91% and 98%.^{10,14,17} Etoh et al. reported lower NPVs, but also reported a higher prevalence of taste disturbances.

Etoh et al. also used the FPD as a reference test. Diagnostic measurement properties for the EGM were generally higher when the FPD was used as the reference test.

4 | DISCUSSION

4.1 | Summary of main results

The objective of this systematic review was to evaluate the validity, reliability, and the diagnostic accuracy of the EGM in assessing taste sensation. Nineteen articles discussing 18 studies were included for data extraction. The included studies on validity and diagnostic accuracy carry a high risk of bias and the overall quality of the studies on reliability were rated as doubtful or inadequate.^{34,6–22} Overall, the association between a variety of reference taste tests and EGM was moderate or weak with correlation coefficients ranging from -0.51 to $0.40^{3,4,6-9,11,12,15,18-20}$ with one outlier of -0.74 found in one study correlating EGM and taste solutions.¹⁹ Test–retest reliability was good with reported correlation coefficients between 0.78 and $1.00.^{16-18,21}$ The sensitivity, specificity, PPV, and NPV of EGM in identifying abnormal taste function varied widely between the four studies on diagnostic accuracy. Only the specificity was above 68% in all four studies.^{10,13,14,17}

4.2 | Quality of evidence

The included studies on validity carry a high risk of bias. Eleven out of 12 included studies on validity retrospectively examined the validity of the EGM. Therefore, none of the included studies formulated hypothesis beforehand.^{3,4,6-9,11,12,18-20} To investigate construct validity it is important to test predefined hypotheses, as it allows the authors to draw

unbiased conclusions after data collection and analyses.²⁹ Furthermore, it is unknown for all studies whether the EGM assessors were blinded.^{3,4,6-22} If the EGM assessors were not blinded, they were able to adjust their interpretation of the electric threshold to match the results of the reference test, which is called review bias. Review bias can falsely increase validity and diagnostic accuracy of the EGM.⁶⁷ Finally, a sample size of more than 100 patients is an appropriate number to test validity and reliability, but only seven of 14 studies evaluating validity and/or reliability had a sample size of more than 100 participants.²⁴

4.3 | Factors that influence electrical taste threshold

Older age, smoking, a longer duration of electrical stimuli, a smaller electrode and stimulation of a tongue region less densely populated by taste buds are associated with a higher electrical taste threshold. Electrical taste thresholds are higher in patients older than 60 years compared to younger patients and in smokers compared to non-smokers.⁵⁸ Pavlidis et al. increased the stimulus duration from 500 to 2000 ms and found that the electrical taste threshold increased with longer stimulus duration.³ Nicolaescu et al. found that a 25 mm² electrode is associated with a higher electrical taste threshold compared to a 125 mm² electrode.¹⁶ Their hypothesis is that a smaller electrode results in more reliable threshold values.^{16,68} Taste thresholds are higher in tongue regions that are less densely populated by taste buds, such as the medial tongue region, than tongue regions that are more densely populated, such as the tongue tip.¹⁶

The method of detecting the electrical taste threshold varies. Most studies used the ascending protocol or the staircase detection protocol.^{3,4,6-9,11-19} None of the included studies compared different stimulation methods.^{3,4,6-22} Staircase procedures have been widely used in measuring thresholds of sensory systems. Therefore, this is currently the recommended stimulation method when using the EGM.^{69,70} The recommended procedure during the staircase detection protocol is to continue testing until at least six reversals are obtained.^{69,70} Ellegård et al. obtained eight reversals per patient, following the protocol of Stillman et al., and Nicolaescu et al. obtained seven reversals per patients.^{14–16,52} The other four studies did not document the number of reversals or obtained fewer than six reversals.^{38,12,20}

Nine out of the 18 included studies reported that the ion current during electrical stimulation was anode (+).^{10,14–22} The other nine studies did not report whether the anode or cathode (–) touched the tongue during EGM testing.^{3,4,6–9,11–13} Anodic stimulation results in a sour taste, while cathodic stimulation leads to a bitter/soapy taste. The advantage of using the anode is that the risk of trigeminal stimulation being perceived by the subject while measuring electrical taste thresholds is lower. Trigeminal stimulation is sensed when using higher current levels, causing a tingling or burning stimulation.^{3,51,63} Cathodic stimulation only evokes taste at higher currents, so using a cathode ion current leads to higher risk of trigeminal nerve stimulation perceived by the subjects when measuring electrical taste

thresholds.⁷¹ To avoid trigeminal involvement, participants should be instructed to distinguish between the sensation experienced by electrical stimulation of the gustatory nerve and the sensation experienced by electrical stimulation of the trigeminal nerve.³

4.4 | Strengths and weaknesses of review process

To our knowledge, this is the first systematic review on this topic. The search strategy was transparent and developed with the help of a clinical librarian. The title abstract screening, full text screening, quality assessment, and data extraction of eligible articles was performed independently by two authors. The evaluation of validity, reliability, and diagnostic accuracy was based on predetermined criteria, which were described in the methods section.

Several limitations may have biased the outcomes of our systematic review. First of all, we did not use a risk of bias tool to assess the quality of the studies on validity and diagnostic accuracy. We evaluated the quality of the includes studies, but a risk of bias tool helps to transparently evaluate the quality of the included studies by the authors. However, we would have had to use numerous different tools to assess the risk of bias in these studies, because the study design and measurements used to quantify the diagnostic accuracy or validity of the EGM differed widely between the studies. Second, study populations and experimental procedures differed between studies. As a result, data could not be pooled. Lastly, articles written in languages other than English, Dutch, French, and German were excluded during title abstract and full text screening. Therefore, we excluded articles written in Japanese, Russian, and Polish. There is a possibility that these articles were relevant to our review.

5 | CONCLUSION

The included studies in this review lack the required standard of the study design to draw firm conclusions about the validity, reliability and diagnostic accuracy of the EGM. Future research is needed to assess the validity, reliability, and diagnostic accuracy of the EGM. However, based on the reported results, we would not recommend using the EGM as a screening test for taste disturbance in clinical practice.

CONFLICT OF INTEREST STATEMENT

The authors declare no conflicts of interest.

DATA AVAILABILITY STATEMENT

Data, code, and other materials will be available upon reasonable request.

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Additional supporting information can be found online in the Supporting Information section at the end of this article.

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