ClinicalEvidence

Halitosis

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ABSTRACT

INTRODUCTION: Halitosis can be caused by oral disease or by respiratory tract conditions such as sinusitis, tonsillitis, and bronchiectasis, but an estimated 40% of affected individuals have no underlying organic disease. METHODS AND OUTCOMES: We conducted a systematic review and aimed to answer the following clinical question: What are the effects of treatments in people with physiological halitosis? We searched: Medline, Embase, The Cochrane Library, and other important databases up to July 2013 (Clinical evidence reviews are updated periodically; please check our website for the most up-to-date version of this review). We included harms alerts from relevant organisations such as the US Food and Drug Administration (FDA) and the UK Medicines and Healthcare products Regulatory Agency (MHRA). RESULTS: We found 11 studies that met our inclusion criteria. We performed a GRADE evaluation of the quality of evidence for interventions. CON-CLUSIONS: In this systematic review, we present information relating to the effectiveness and safety of the following interventions: artificial saliva; cleaning, brushing, or scraping the tongue; regular use of mouthwash; sugar-free chewing gums; and zinc toothpastes.

QUESTIONS

What are the effects of treatments in people with physiological halitosis?						
INTERV	ENTIONS					
TREATMENTS FOR PHYSIOLOGICAL HALITOSIS	OO Unknown effectiveness					
Likely to be beneficial	Artificial saliva					
Regular-use mouthwash (containing chlorhexidine, zinc,	Sugar-free chewing gum					
hydrogen peroxide, or other antimicrobial agents) 2	Tongue cleaning, brushing, or scraping 11					
Zinc toothpastes						

Key points

- Halitosis can be caused by oral disease or by respiratory tract conditions such as sinusitis, tonsillitis, and bronchiectasis, but an estimated 40% of affected people have no underlying organic disease.
 - The main chemicals causing the odour seem to be volatile sulfur compounds, but little is known about the cause of physiological halitosis.
- Regular use of a mouthwash may reduce breath odour compared with placebo.
- Zinc toothpastes seem to reduce breath odour compared with placebo for people with halitosis.
- · We don't know whether tongue cleaning, sugar-free chewing gums, or artificial saliva reduce halitosis, as no studies of adequate quality have been found.

DEFINITION

Halitosis is an unpleasant odour emitted from the mouth. It may be caused by oral conditions, including poor oral hygiene and periodontal disease [1] [2] or by respiratory tract conditions, such as chronic sinusitis, tonsillitis, and bronchiectasis. In this review, we deal only with physiological halitosis (i.e., confirmed persistent bad breath in the absence of systemic, oral, or periodontal disease). We have excluded halitosis caused by underlying systemic disease that would require disease-specific treatment, pseudo-halitosis (in people who believe they have bad breath but whose breath is not considered malodorous by others), and artificially induced halitosis (e.g., in studies requiring people to stop brushing their teeth). This review is only applicable, therefore, to people in whom such underlying causes have been ruled out, and in whom pseudo-halitosis has been excluded. There is no consensus regarding duration of bad breath for the diagnosis of halitosis. although the standard organoleptic test for bad breath involves smelling the breath on at least two or three different days. [1] Professional tooth cleaning may be of value where periodontal disease or poor oral hygiene contribute to malodour.

INCIDENCE/ **PREVALENCE**

We found no reliable estimate of prevalence, although several studies report the population prevalence of halitosis (physiological or because of underlying disease) to be about 50%. [1] [3] ^[5] One cross-sectional study of 491 people found that about 5% of people with halitosis have pseudo-halitosis and about 40% have physiological bad breath not caused by underlying disease.

[6] We found no reliable data about 7. We found no reliable data about age or sex distribution of physiological halitosis.

AETIOLOGY/

We found no reliable data about risk factors for physiological bad breath. Mass spectrometric and RISK FACTORS gas chromatographic analysis of expelled air from the mouths of people with any type of halitosis have shown that the principal malodorants are volatile sulfur compounds, including hydrogen sulfide, methyl mercaptan, and dimethyl sulfide. [7] [8]

PROGNOSIS We found no evidence on the prognosis of halitosis.

AIMS OF To improve social functioning; to reduce embarrassment; to reduce odour, with minimum adverse **INTERVENTION** effects.

OUTCOMES

Breath odour, measured by organoleptic test scores or other odour scales; **quality of life**, including embarrassment and social functioning; **adverse effects**. We excluded non-clinical outcomes such as gas chromatography and spectroscopy results, and concentrations of compounds in exhaled air.

METHODS

Clinical Evidence search and appraisal July 2013. The following databases were used to identify studies for this systematic review: Medline 1966 to July 2013, Embase 1980 to July 2013, and The Cochrane Database of Systematic Reviews 2013, issue 7 (1966 to date of issue). Additional searches were carried out in the Database of Abstracts of Reviews of Effects (DARE) and the Health Technology Assessment (HTA) database. We also searched for retractions of studies included in the review. Titles and abstracts identified by the initial search, run by an information specialist, were first assessed against predefined criteria by an evidence scanner. Full texts for potentially relevant studies were then assessed against predefined criteria by an evidence analyst. Studies selected for inclusion were discussed with an expert contributor. All data relevant to the review were then extracted by an evidence analyst. Study design criteria for inclusion in this review were: published RCTs and systematic reviews of RCTs in the English language, at least singleblinded, and containing at least 20 individuals (at least 10 per arm) of whom at least 80% were followed up. There was no minimum length of follow-up. We excluded all studies described as 'open', 'open label', or not blinded unless blinding was impossible. We included RCTs and systematic reviews of RCTs where harms of an included intervention were studied, applying the same study design criteria for inclusion as we did for benefits. In addition, we use a regular surveillance protocol to capture harms alerts from organisations such as the FDA and the MHRA, which are added to the reviews as required. To aid readability of the numerical data in our reviews, we round many percentages to the nearest whole number. Readers should be aware of this when relating percentages to summary statistics such as relative risks (RRs) and odds ratios (ORs). We have performed a GRADE evaluation of the quality of evidence for interventions included in this review (see table, p 15). The categorisation of the quality of the evidence (high, moderate, low, or very low) reflects the quality of evidence available for our chosen outcomes in our defined populations of interest. These categorisations are not necessarily a reflection of the overall methodological quality of any individual study, because the Clinical Evidence population and outcome of choice may represent only a small subset of the total outcomes reported, and population included, in any individual trial. For further details of how we perform the GRADE evaluation and the scoring system we use, please see our website (www.clinicalevidence.com).

QUESTION

What are the effects of treatments in people with physiological halitosis?

OPTION

REGULAR-USE MOUTHWASH (CONTAINING CHLORHEXIDINE, ZINC, HYDROGEN PEROXIDE, OR OTHER ANTIMICROBIAL AGENTS)

- For GRADE evaluation of interventions for Halitosis, see table, p 15.
- Regular use of a mouthwash may reduce breath odour compared with placebo.

Benefits and harms

Regular-use mouthwash versus placebo:

We found two systematic reviews (search date 2008; [9] and 2012 [10]), which identified four RCTs of sufficient quality [11] [12] [13] [14] and one subsequent RCT. [15] The reviews did not pool data. We have reported the five RCTs from their original reports. [11] [12] [13] [14] [15] The first RCT compared an active-treatment mouthwash (containing chlorhexidine plus cetylpyridinium chloride plus zinc lactate) versus a placebo mouthwash. The mouthwashes were used twice-daily for 2 weeks. [11] The second RCT compared four mouthwashes, used twice-daily for 4 weeks (one containing essential oils; one containing cetylpyridinium chloride; one containing chlorine dioxide plus zinc; and a placebo mouthwash [composition not reported]]. [12] The third RCT compared three interventions over 4 weeks (mouthwash containing zinc chloride plus sodium chlorite; mouthwash containing zinc chloride alone; and placebo mouthwash). All participants were instructed to use mouthwash for 30 seconds twice-daily. [13] The fourth RCT compared three mouthwashes with placebo (one containing amine fluoride/stannous fluoride, zinc lactate, and oral malodour counteractives; one containing chlorhexidine, cetylpyridinium chloride, and zinc lactate; and one containing chlorhexidine). [14] The fifth RCT compared mouthwash with placebo mouthwash and mouthwash plus tongue scraping with placebo mouthwash plus tongue scraping. The mouthwash contained zinc acetate, chlorhexidine diacetate, and sodium fluoride. [15]

Breath odour

Regular-use mouthwash compared with placebo Regular use of a mouthwash containing chlorhexidine plus cetylpyridinium chloride plus zinc lactate; cetylpyridinium chloride alone; zinc chloride plus sodium chlorite; amine fluoride/stannous fluoride, zinc lactate, and oral malodour counteractives; or chlorhexidine alone may be more effective than placebo at reducing breath odour at 2 to 4 weeks. However, regular use of mouthwash containing essential oil or chlorine dioxide plus zinc may be no more effective at reducing breath odour at 2 weeks. We don't know if regular use of mouthwash containing zinc acetate, chlorhexidine diacetate, and sodium fluoride (with or without tongue scraping) is more effective than placebo mouthwash (with or without tongue scraping) at reducing breath odour at 2 weeks (low-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Breath o	dour				·
[11] RCT	40 people	Mean odour score change from baseline (scale from 0 [no halitosis]—5 [offensive halitosis]), at 2 weeks -1.3 with mouthwash containing chlorhexidine plus cetylpyridinium	P <0.005		regular-use mouth- wash containing
		chloride plus zinc lactate (used twice-daily) -0.2 with placebo mouthwash (used twice-daily) Breath odour was assessed by 1 trained examiner		000	chlorhexidine plus cetylpyridinium chloride plus zinc lactate
[12] RCT	99 people	Mean odour score change from baseline (scale from 0 [no halitosis]—5 [offensive halitosis])	P <0.05 for cetylpyridinium chloride mouthwash <i>v</i> placebo		
4-armed trial		, at 4 weeks	4 people were excluded or with- drew after randomisation		
uiai		-0.41 with mouthwash containing cetylpyridinium chloride (used twice-daily)	Analysis not by intention-to-treat		regular-use mouth-
		+0.16 with placebo mouthwash (used twice-daily)		000	wash containing cetylpyridinium chloride
		The remaining arms evaluated chlorine dioxide plus zinc mouthwash and essential oil mouthwash			
		Breath odour was assessed by 2 experienced examiners			
RCT 4-armed	99 people	Mean odour score change from baseline (scale from 0 [no hali- tosis]—5 [offensive halitosis]) , at 4 weeks	P value reported as not significant for chlorine dioxide plus zinc mouthwash <i>v</i> placebo 4 people were excluded or with-		
trial		+0.06 with chlorine dioxide plus zinc mouthwash (used twice-dai- ly)	drew after randomisation Analysis not by intention-to-treat		
		+0.16 with placebo mouthwash (used twice-daily)		\longleftrightarrow	Not significant
		The remaining arms evaluated mouthwash containing cetylpyridinium chloride and essential oil mouthwash			
		Breath odour was assessed by 2 experienced examiners			
[12] PCT	99 people	Mean odour score change from baseline (scale from 0 [no hali-	P value reported as not significant for essential oil mouthwash		
RCT 4-armed		tosis]—5 [offensive halitosis]) , at 4 weeks	v placebo 4 people were excluded or with-	\longleftrightarrow	Not significant
trial		0 with essential oil mouthwash (used twice-daily)	drew after randomisation		
			Analysis not by intention-to-treat		

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
		+0.16 with placebo mouthwash (used twice-daily) The remaining arms evaluated mouthwash containing cetylpyridinium chloride and chlorine dioxide plus zinc mouthwash Breath odour was assessed by 2 experienced examiners			
RCT 3-armed trial	48 people	Organoleptic breath scores , 4 weeks with regular-use mouthwash containing zinc chloride plus sodium chlorite with placebo mouthwash Absolute results reported graphically The remaining arm evaluated regular-use mouthwash containing zinc chloride only	Reported as significant for zinc chloride plus sodium chlorite mouthwash <i>v</i> placebo mouthwash P value not reported	000	regular-use mouth- wash containing zinc chloride plus sodium chlorite
RCT 4-armed trial	174 people	Mean odour score change from baseline, 21 days with amine fluoride/stannous fluoride, zinc lactate, oral malodour counteractives mouthwash with placebo mouthwash (tap water) Absolute results reported graphically The remaining arms evaluated mouthwashes containing chlorhexidine, cetylpyridinium chloride, and zinc lactate, or chlorhexidine	P <0.05 Statistically significant improvement in odour scores in the mouthwash group ν placebo group were observed additionally at days 1 and 7, but not day 14	000	regular-use mouth- wash containing amine fluoride/stan- nous fluoride, zinc lactate, and oral malodour counter- actives
RCT 4-armed trial	174 people	Mean odour score change from baseline , 21 days with chlorhexidine, cetylpyridinium chloride, and zinc lactate mouthwash with placebo mouthwash (tap water) Absolute results reported graphically The remaining arms evaluated mouthwashes containing amine fluoride/stannous fluoride, zinc lactate, and oral malodour counteractives, or chlorhexidine	P <0.05 Statistically significant improvement in odour scores in the mouthwash group ν placebo group were observed additionally at days 7 and 14, but not day 1	000	regular-use mouth- wash containing chlorhexidine, cetylpyridium chlo- ride, and zinc lac- tate
RCT 4-armed trial	174 people	Mean odour score change from baseline , 21 days with chlorhexidine mouthwash with placebo mouthwash (tap water) Absolute results reported graphically The remaining arms evaluated mouthwashes containing amine fluoride/stannous fluoride, zinc lactate, and oral malodour counteractives, or chlorhexidine,	P <0.05 Statistically significant improvement in odour scores in the mouthwash group ν placebo group were observed additionally at days 1 and 7, but not day 14	000	regular-use mouth- wash containing chlorhexidine

				H	łalitosis
Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
		cetylpyridinium chloride, and zinc lactate			
[15] RCT Crossover design 4-armed trial	21 people	Proportion of people reporting reduction in organoleptic breath scores from baseline , 14 days 38% with zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash 24% with placebo mouthwash Absolute numbers not reported The remaining arms of this 2 × 2 factorial trial evaluated combination of zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash plus tongue scraping and combination of placebo mouthwash plus tongue scraping	Not reported		
RCT Crossover design 4-armed trial	21 people	Proportion of people reporting reduction in organoleptic breath scores from baseline, 14 days 67% with combination of zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash plus tongue scraping 33% with combination of placebo mouthwash plus tongue scraping Absolute numbers not reported The remaining arms of this 2 × 2 factorial trial evaluated zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash and placebo mouthwash	Not reported		

Quality of life

No data from the following reference on this outcome. $^{[11]}$ $^{[12]}$ $^{[13]}$ $^{[14]}$ $^{[15]}$

Adverse effects

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Adverse 6	effects				
RCT	40 people	Mean change in tongue discoloration score from baseline (assessed using the Winkel tongue discoloration index [measured in 6 tongue areas; range 0 = no discoloration to 12 = severe discoloration]), 2 weeks +2.8 with mouthwash containing chlorhexidine plus cetylpyridinium chloride plus zinc lactate +0.3 with placebo mouthwash	P <0.002	000	placebo mouth- wash

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
RCT	40 people	Tooth staining, 2 weeks with mouthwash containing chlorhexidine plus cetylpyridinium chloride plus zinc lactate with placebo mouthwash Absolute results not reported	Reported as non-significant P value not reported	\longleftrightarrow	Not significant
RCT 4-armed trial	99 people	Adverse effects , 4 weeks with mouthwash containing cetylpyridinium chloride with chlorine dioxide plus zinc mouthwash with essential oil mouthwash with placebo mouthwash Absolute results not reported 13 people reported adverse effects such as lip blisters, localised gingival oedema, and sores (figures not reported by treatment group); the RCT reported, "it was determined that these adverse events were unlikely to be related to the product usage"	4 people were excluded or with- drew after randomisation Analysis not by intention-to-treat		
RCT 4-armed trial	174 people	Discoloration of at least 1 upper anterior or incisor tooth, day 21 8/44 (18%) with amine fluoride/stannous fluoride, zinc lactate, oral malodour counteractives mouthwash 10/44 (23%) with chlorhexidine, cetylpyridinium chloride, and zinc lactate mouthwash 13/44 (30%) with chlorhexidine mouthwash 5/44 (11%) with placebo mouthwash (tap water)	Reported as not significant	\longleftrightarrow	Not significant

No data from the following reference on this outcome. [13] [15]

Regular-use mouthwashes versus each other:

We found two systematic reviews (search date 2008; ^[9] and 2012 ^[10]), which identified three RCTs ^[12] ^[13] ^[14] and one subsequent RCT. ^[16] We have reported all four RCTs from their original reports. ^[12] ^[13] ^[14] ^[16] The first RCT compared four mouthwashes, used twice-daily, for 4 weeks (one containing essential oils; one containing cetylpyridinium chloride; one containing chlorine dioxide plus zinc; and a placebo mouthwash [composition not reported]). ^[12] The second RCT compared three interventions over 4 weeks (mouthwash containing zinc chloride plus sodium chlorite; mouthwash containing zinc chloride alone; and placebo mouthwash). All participants were instructed to use mouthwash for 30 seconds, twice-daily. ^[13] The third RCT compared four mouthwashes (one containing amine fluoride/stannous fluoride, zinc lactate, and oral malodour counteractives; one containing chlorhexidine, cetylpyridinium chloride, and zinc lactate; one containing chlorhexidine; and a placebo mouthwash [tap water]). ^[14] The fourth RCT compared a chlorine dioxide mouthwash with a chlorhexidine gluconate mouthwash. All participants also had tongue scraping. ^[16]

Breath odou

Regular-use mouthwashes compared with each other Regular use of a mouthwash containing zinc chloride plus sodium chlorite may be more effective than mouthwash containing zinc chloride alone at reducing breath odour at

4 weeks. We don't know how other regular-use mouthwashes with different ingredients compare with each other. (low-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Breath oc	lour	*	·		
[12] RCT 4-armed	99 people	Mean odour score change from baseline (assessed on a scale from 0 [no halitosis] to 5 [offen- sive halitosis]) , at 4 weeks	P value (among the 3 treatment groups) reported as not significant 4 people were excluded or with-		
trial		-0.41 with mouthwash containing cetylpyridinium chloride (used twice-daily) +0.06 with chlorine dioxide plus zinc mouthwash (used twice-daily) 0 with essential oil mouthwash The remaining arm evaluated placebo mouthwash Breath odour was assessed by 2 experienced examiners	drew after randomisation Analysis not by intention-to-treat	\longleftrightarrow	Not significant
RCT 3-armed trial	48 people	Organoleptic breath scores 4 weeks with regular-use mouthwash containing zinc chloride plus sodium chlorite with regular-use mouthwash containing zinc chloride only Absolute results reported graphically The remaining arm evaluated regular-use placebo mouthwash	Reported as significant for zinc chloride plus sodium chlorite mouthwash v zinc chloride alone mouthwash P value not reported	000	regular-use mouth- wash containing zinc chloride plus sodium chlorite
[14] RCT 4-armed trial	174 people	Mean odour score change from baseline, day 21 with amine fluoride/stannous fluoride, zinc lactate, oral malodour counteractives mouthwash with chlorhexidine, cetylpyridinium chloride, and zinc lactate mouthwash with chlorhexidine mouthwash Absolute results reported graphically The remaining arm evaluated placebo mouthwash (tap water) No statistically significant differences in odour scores among the 3 mouthwashes were additionally observed at days 1, 7, and 14	Reported as not significant	\longleftrightarrow	Not significant
[16] RCT	22 people	Proportion of individuals with organoleptic scores 1 or less, 1 week 64% with chlorine dioxide mouthwash plus tongue scraping 81% with chlorhexidine gluconate mouthwash plus tongue scraping Absolute numbers not reported	Reported as not significant	\longleftrightarrow	Not significant

Quality of life

No data from the following reference on this outcome. $^{[12]}$ $^{[13]}$ $^{[14]}$ $^{[16]}$

Adverse effects

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Adverse e			•		
[12]	99 people	Adverse effects , 4 weeks	4 people were excluded or with-		
RCT		with mouthwash containing	drew after randomisation		
4-armed		cetylpyridinium chloride	Analysis not by intention-to-treat		
trial		with chlorine dioxide plus zinc mouthwash			
		with essential oil mouthwash			
		with placebo mouthwash			
		Absolute results not reported			
		13 people reported adverse effects such as lip blisters, localised gingival oedema, and sores (figures not reported by treatment group); the RCT reported, "it was determined that these adverse events were unlikely to be related to the product usage"			
[14]	174 people	Discoloration of at least 1 up-	Reported as not significant		
RCT		per anterior or incisor tooth , day 21			
4-armed trial		8/44 (18%) with amine fluo- ride/stannous fluoride, zinc lac- tate, oral malodour counterac- tives mouthwash 10/44 (23%) with chlorhexidine, cetylpyridinium chloride, and zinc lactate mouthwash		\longleftrightarrow	Not significant
		13/44 (30%) with chlorhexidine mouthwash			
		5/44 (11%) with placebo mouthwash (tap water)			
[16]	22 people	Adverse effects , 1 week	Not reported		
RCT		with chlorine dioxide mouthwash plus tongue scraping			
		with chlorhexidine gluconate mouthwash plus tongue scraping			
		Chlorine dioxide group reported altered taste sensation, which resolved upon mouthwash discontinuation (n = 1)			
		No other adverse effects includ- ing soft tissue lesions or effect on teeth or restorations were report- ed			

No data from the following reference on this outcome. [13]

Regular-use mouthwash versus artificial saliva, sugar-free gum, tongue cleaning, or zinc toothpastes:

We found one RCT comparing regular-use mouthwash with tongue scraping. [15]

Breath odour

Regular-use mouthwash compared with placebo mouthwash plus tongue scraping We don't know how effective regular-use mouthwash is compared with placebo mouthwash plus tongue scraping in reducing halitosis (very low-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Breath od	lour			*	
RCT Crossover design 4-armed trial	21 people	Proportion of people reporting reduction in organoleptic breath scores from baseline, day 14 38% with zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash 33% with combination of placebo mouthwash plus tongue scraping Absolute numbers not reported The remaining arms of this 2 × 2 factorial trial evaluated combination of zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash plus tongue scraping and placebo mouthwash	Not reported		

Quality of life

No data from the following reference on this outcome. [15]

Adverse effects

No data from the following reference on this outcome. [15]

Mouthwash plus tongue scraping versus placebo:

We found one RCT comparing regular-use mouthwash plus tongue scraping with placebo mouthwash alone. [15]

Breath odour

Regular-use mouthwash plus tongue scraping compared with placebo mouthwash We don't know how effective regular-use mouthwash plus tongue scraping is compared with placebo mouthwash alone (very-low-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Breath od	lour				
[15]	21 people	Proportion of people reporting	Not reported		
RCT		reduction in organoleptic breath scores from baseline .			
Crossover		day 14			
design		67% with combination of zinc ac-			
4-armed trial		etate, chlorhexidine diacetate, and sodium fluoride mouthwash			
liidi		plus tongue scraping			
		24% with placebo mouthwash			

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
		Absolute numbers not reported			
		The remaining arms of this 2 x 2 factorial trial evaluated zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash and combination of placebo mouthwash plus tongue scraping			

Quality of life

No data from the following reference on this outcome. [15]

Adverse effects

No data from the following reference on this outcome. [15]

Comment: None.

OPTION ARTIFICIAL SALIVA

- For GRADE evaluation of interventions for Halitosis, see table, p 15 .
- We don't know whether artificial saliva reduces halitosis, as no trials of adequate quality have been found.

Benefits and harms

Artificial saliva:

We found no systematic review or RCTs comparing artificial saliva versus placebo or versus the other interventions covered by this review.

Comment:

Although we searched for artificial saliva, it is mainly given to people with dry mouth syndrome, which lies outside of the remit of this review.

OPTION SUGAR-FREE CHEWING GUM

- For GRADE evaluation of interventions for Halitosis, see table, p 15.
- We don't know whether sugar-free chewing gums reduce halitosis, as no trials of adequate quality have been found.

Benefits and harms

Sugar-free chewing gum:

We found no systematic review or RCTs comparing sugar-free chewing gum versus placebo or versus the other interventions covered by this review.

Comment: None.

OPTION TONGUE CLEANING, BRUSHING, OR SCRAPING

- For GRADE evaluation of interventions for Halitosis, see table, p 15 .
- We don't know whether tongue cleaning reduces halitosis.

Benefits and harms

Tongue scraping versus no tongue scraping:

We found two systematic reviews (search date 2005; [17] and 2009 [18]), which identified no RCTs of sufficient quality. We found one subsequent RCT comparing tongue scraping with no tongue scraping. [15]

Breath odour

Tongue scraping compared with no tongue scraping We don't know how effective tongue scraping with or without mouthwash is compared with mouthwash or placebo in reducing halitosis (very-low-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours				
Breath od	Breath odour								
RCT Crossover design 4-armed trial	21 people	Proportion of people reporting reduction in organoleptic breath scores from baseline, day 14 67% with combination of zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash plus tongue scraping 38% with zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash Absolute numbers not reported The remaining arms of this 2 × 2 factorial trial evaluated combination of placebo mouthwash plus tongue scraping, and placebo mouthwash	Not reported						
RCT Crossover design 4-armed trial	21 people	Proportion of people reporting reduction in organoleptic breath scores from baseline, day 14 33% with combination of placebo mouthwash plus tongue scraping 24% with placebo mouthwash Absolute numbers not reported The remaining arms of this 2 × 2 factorial cross-over trial evaluated combination of zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash plus tongue scraping and zinc acetate, chlorhexidine diacetate, and sodium fluoride mouthwash	Not reported						

Quality of life

No data from the following reference on this outcome. [15]

Adverse effects

No data from the following reference on this outcome. [15]

Further information on studies

The mouthwash used in the RCT contained 0.3% zinc acetate, 0.025% chlorhexidine diacetate, and 0.05% sodium fluoride.

Comment: None.

OPTION ZINC TOOTHPASTES

- For GRADE evaluation of interventions for Halitosis, see table, p 15.
- Zinc toothpastes seem to reduce breath odour compared with placebo for people with halitosis.

Benefits and harms

Zinc toothpastes versus placebo:

We found one RCT comparing zinc toothpaste versus placebo. [19]

Breath odour

Zinc toothpastes compared with placebo Use of zinc toothpastes may be more effective than placebo at reducing breath odour at 4 weeks (moderate-quality evidence).

Ref (type)	Population	Outcome, Interventions	Results and statistical analysis	Effect size	Favours
Breath oc	lour				
[19] RCT	187 people	Mean organoleptic breath scores before brushing , 4 weeks 2.15 with toothpaste containing zinc sulfate and fluoride 2.80 with placebo toothpaste containing fluoride	P = 0.0001	000	zinc sulfate tooth- paste
[19] RCT	187 people	Mean organoleptic breath scores 2 hours after brushing , 4 weeks 1.54 with toothpaste containing zinc sulfate and fluoride 2.85 with placebo toothpaste containing fluoride	P <0.0001	000	zinc sulfate tooth- paste

Quality of life

No data from the following reference on this outcome. [19]

Adverse effects

No data from the following reference on this outcome. [19]

Comment: None.

GLOSSARY

Low-quality evidence Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Moderate-quality evidence Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

Organoleptic test scores These are assigned by one or more examiners who sniff the person's exhaled breath on two or three different days. People having this examination should not have had antibiotics in the previous 3 weeks, and should have refrained from eating garlic, onions, and spicy foods for 48 hours, and should have refrained from usual oral hygiene and smoking for the previous 12 hours. [1] Scoring systems vary among studies.

Very low-quality evidence Any estimate of effect is very uncertain.

SUBSTANTIVE CHANGES

Regular-use mouthwash (containing chlorhexidine, zinc, hydrogen peroxide, or other antimicrobial agents) Two systematic reviews added, [9] [10] and three RCTs. [14] [15] [16] Categorisation unchanged (likely to be beneficial).

Tongue cleaning, brushing, or scraping Two systematic reviews added, ^{[17] [18]} and one subsequent RCT. ^[15] Categorisation unchanged (unknown effectiveness).

Zinc toothpastes One RCT added. [19] Categorisation changed from unknown effectiveness to likely to be beneficial.

REFERENCES

- Yaegaki K, Coil JM. Examination, classification, and treatment of halitosis; clinical perspectives. J Can Dent Assoc 2000;66:257–261.[PubMed]
- Frascella J, Gilbert RD, Fernandez P, et al. Efficacy of a chlorine dioxide-containing mouthrinse in oral malodor. Compend Contin Educ Dent 2000;21:241–254.[PubMed]
- Meningaud JP, Bado F, Favre E, et al. Halitosis in 1999. Rev Stomatol Chir Maxillofac 1999;100:240–244. [In French][PubMed]
- Bollen CM, Rompen EH, Demanez JP. Halitosis: a multidisciplinary problem. Rev Med Liege 1999;54:32–36. [In French][PubMed]
- Tomas Carmona I, Limeres Posse J, Diz Dios P, et al. Extraoral etiology of halitosis. Med Oral 2001;6:40–47.[PubMed]
- Delanghe G, Bollen C, van Steenberghe D, et al. Halitosis, foetor ex ore [in Dutch]. Ned Tijdschr Tandheelkd 1998;105:314–317.[PubMed]
- Tonzetich J. Direct gas chromatographic analysis of sulphur compounds in mouth air in man. Arch Oral Biol 1971;16:587–597. [PubMed]
- Kleinberg I, Westbay G. Oral malodor. Crit Rev Oral Biol Med
 1990;1:247–259.[PubMed]
 Federovica Z. Allufairi H. Necces M. et al. Mouth/inces for the treatment.
- Fedorowicz Z, Aljufairi H, Nasser M, et al. Mouthrinses for the treatment of halitosis. In: The Cochrane Library. Issue 7, 2013. Search date 2008. PubMedl
- Blom T, Slot DE, Quirynen M, et al. The effect of mouthrinses on oral malodor: a systematic review. Int J Dent Hyg. 2012;10:209–222.[PubMed]
- Winkel EG, Roldan S, Van Winkelhoff AJ, et al. Clinical effects of a new mouthrinse containing chlorhexidine, cetylpyridinium chloride and zinc-lactate

- on oral halitosis. A dual-center, double-blind placebo-controlled study. J Clin Periodontol 2003;30:300–306.[PubMed]
- Borden LC, Chaves ES, Bowman JP, et al. The effect of four mouthrinses on oral malodor. Compend Contin Educ Dent 2002;23:531–546.[PubMed]
- Codipilly DP, Kaufman HW, Kleinberg I. Use of a novel group of oraLmalodor measurements to evaluate an anti-oral malodor mouthrinse (TriOral M) in humans. J Clin Dent 2004;15:98–104.[PubMed]
- Wigger-Alberti W, Gysen K, Axmann EM, et al. Efficacy of a new mouthrinse formulation on the reduction of oral malodour in vivo. A randomized, double-blind, placebo-controlled, 3 week clinical study. J Breath Res 2010;4:017102.[PubMed]
- Erovic Ademovski S, Lingström P, Winkel E, et al. Comparison of different treatment modalities for oral halitosis. Acta Odontol Scand 2012;70:224–233.[PubMed]
- Wirthlin MR, Im T, Ellis RR, et al. Effect of tongue scraper and rinses on bad breath, a double-blind, randomized, parallel group clinical trial. J West Soc Periodontol Periodontal Abstr 2011;59:67–73.[PubMed]
- Outhouse TL, Al-Alawi R, Fedorowicz Z, et al. Tongue scraping for treating halitosis. In: The Cochrane Library, Issue 7, 2013. Search date 2005.[PubMed]
- Van der Sleen MI, Slot DE, Van Trijffel E, et al. Effectiveness of mechanical tongue cleaning on breath odour and tongue coating: a systematic review. Int J Dent Hyg 2010;8: 258–268.[PubMed]
- Navada R, Kumari H, Le S, et al. Oral malodor reduction from a zinc-containing toothpaste. J Clin Dent 2008;19:69–73. [PubMed]

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Evaluation of interventions for Halitosis.

Important out- comes	Breath odour, Quality of life									
Studies (Partici- pants)	Outcome	Comparison	Type of evi- dence	Quality	Consisten- cy	Directness	Effect size	GRADE	Comment	
What are the effects of treatments in people with physiological halitosis?										
5 (382) ^[11] ^[12] ^[13] ^[14] ^[15]	Breath odour	Regular-use mouthwash versus placebo	4	-2	0	0	0	Low	Quality points deducted for methodological flaws and incomplete reporting of results	
4 (less than 343 peo- ple) [12] [13] [14] [16]	Breath odour	Regular-use mouthwashes versus each other	4	-2	0	0	0	Low	Quality points deducted for methodological flaws and incomplete reporting of results	
1 (21) ^[15]	Breath odour	Regular-use mouthwash versus artificial saliva, sugar- free gum, tongue cleaning, or zinc toothpastes	4	-2	0	– 1	0	Very low	Quality points deducted for sparse data and incomplete reporting of results; di- rectness point deducted for uncertainty about the definition of the outcome	
1 (21) ^[15]	Breath odour	Mouthwash plus tongue scraping versus placebo	4	-2	0	– 1	0	Very low	Quality points deducted for sparse data and incomplete reporting of results; di- rectness point deducted for uncertainty about the definition of the outcome	
1 (21) ^[15]	Breath odour	Tongue scraping versus no tongue scraping	4	-2	0	– 1	0	Very low	Quality points deducted for sparse data and incomplete reporting of results; di- rectness point deducted for uncertainty about the definition of the outcome	
1 (187) ^[19]	Breath odour	Zinc toothpastes versus placebo	4	-1	0	0	0	Moderate	Quality point deducted for sparse data	

We initially allocate 4 points to evidence from RCTs, and 2 points to evidence from observational studies. To attain the final GRADE score for a given comparison, points are deducted or added from this initial score based on preset criteria relating to the categories of quality, directness, consistency, and effect size. Quality: based on issues affecting methodological rigour (e.g., incomplete reporting of results, quasi-randomisation, sparse data [<200 people in the analysis]). Consistency: based on similarity of results across studies. Directness: based on generalisability of population or outcomes. Effect size: based on magnitude of effect as measured by statistics such as relative risk, odds ratio, or hazard ratio.

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