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## Restriction of salt, caffeine and alcohol intake for the treatment of Ménière's disease or syndrome (Review)

Hussain K, Murdin L, Schilder AGM

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[Intervention Review]

# Restriction of salt, caffeine and alcohol intake for the treatment of Ménière's disease or syndrome

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## ABSTRACT

### Background

Ménière's disease or syndrome is a chronic inner ear disorder that results in sporadic attacks of vertigo, sensorineural hearing loss, aural fullness and tinnitus.

There is no definitive treatment for Ménière's disease and treatment options range from dietary modification through medication to surgery.

Modification of diet, including restriction of salt, caffeine and alcohol intake, is a management option that is widely recommended to patients with Ménière's as a first-line treatment. There has not previously been a systematic review of this intervention.

### Objectives

To assess the effects of dietary restriction of salt, caffeine and alcohol intake in patients with Ménière's disease or syndrome.

### Search methods

The Cochrane ENT Information Specialist searched the Cochrane ENT Trials Register; Central Register of Controlled Trials (CENTRAL); PubMed; Ovid Embase; CINAHL; Web of Science; ClinicalTrials.gov; ICTRP and additional sources for published and unpublished trials. The date of the search was 28 March 2018.

### Selection criteria

Randomised controlled trials of dietary modification, specifically salt, caffeine and alcohol restriction or substitution (or both), compared to no modification of these agents or a placebo intervention, in adult patients with Ménière's disease or syndrome.

### Data collection and analysis

We used the standard methodological procedures expected by Cochrane. Our primary outcomes were control of vertigo or decrease in vertigo attacks, and adverse effects. Secondary outcomes included hearing (change in hearing loss or its progression), tinnitus (severity), perception of aural fullness, well-being and quality of life (overall changes), and other adverse effects. We planned to use GRADE to assess the quality of the evidence for each outcome.

### Main results

We did not identify any studies that met the inclusion criteria for the review.

## Authors' conclusions

There is no evidence from randomised controlled trials to support or refute the restriction of salt, caffeine or alcohol intake in patients with Ménière's disease or syndrome.

High-quality research in this field is warranted. The best evidence may come from a randomised controlled trial comparing dietary interventions (e.g. low salt versus general healthy diet advice), using rigorous methodology for patient selection, randomisation and blinding, and strictly adhering to the Bárány Society diagnostic criteria. However, this research question might be more pragmatically addressed by using information from carefully constructed patient registries that include information on dietary intake of substances of interest such as salt, caffeine and alcohol. It will be important to address the question of any possible harms or unwanted effects of dietary modification.

## PLAIN LANGUAGE SUMMARY

### Dietary changes in the treatment of Ménière's disease or syndrome

#### Review question

What is the effect of changing diet by restricting salt, caffeine and alcohol, alone or together with each other, on the symptoms of people with Ménière's disease?

#### Background

Approximately 200 per 100,000 people suffer from Ménière's. The condition is called Ménière's disease if no cause can be identified and Ménière's syndrome if there is a known reason for its development. Patients who suffer from it experience vertigo (dizzy episodes), hearing loss, a sensation of pressure in their ears and tinnitus (ringing or buzzing in the ears).

At present there is no standard treatment for Ménière's disease and options can range from dietary changes, to medicines and in some cases surgery. Ménière's disease is thought to be caused by disturbance of the volume or composition of the fluid in the inner ear (called endolymph). Dietary intake of salt can affect the concentrations of electrolytes (salts and minerals that can conduct electrical impulses in the body) in the blood, which in turn may affect the composition of the endolymph. Salt intake may therefore contribute to attacks and so restriction in the diet could be used to control both the volume and composition of the endolymph. Caffeine and alcohol intake can result in constriction of blood vessels (vasoconstriction) and could result in a reduction in the blood supply to the inner ear, which may make patients' symptoms worse. Many doctors advise dietary changes as a first-line treatment as it is thought to be a relatively simple and inexpensive option. We wanted to find out whether dietary changes are effective to ensure that patients are receiving the correct advice about treatment options, and to ensure that potentially more appropriate treatment is not delayed by spending time on ineffective interventions, resulting in unpleasant symptoms and disease progression.

#### Study characteristics

We searched for high-quality studies (randomised controlled trials) of dietary changes (salt, caffeine and alcohol restriction or substitution, or both) compared to no restriction in adult patients with Ménière's disease or syndrome. Our search is up to date to March 2018.

#### Key results

We did not identify any randomised controlled trials that met the inclusion criteria for the review.

#### Quality of evidence and conclusions

There is no evidence from randomised controlled trials about the restriction of salt, caffeine or alcohol intake in patients with Ménière's disease or syndrome. High-quality research in this field is needed if this question is to be answered, in the form of a study that uses rigorous methods (for example, randomisation and blinding, or careful use of patient registries) and carefully recruits only patients that meet accepted Ménière's disease diagnostic criteria. It will be important to address the question of any possible harms or unwanted effects of dietary changes.

## SUMMARY OF FINDINGS

### Summary of findings for the main comparison.

#### Dietary modifications compared with no modification/placebo for Ménière's disease or syndrome

**Patient or population:** adults with Ménière's disease or syndrome

**Settings:** hospital and community

**Intervention:** dietary modification

**Comparison:** no modification/placebo

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	No of participants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk				
	No modification/placebo	Dietary modification				
Proportion of patients with control of vertigo or decrease in vertigo attacks	No data	No data	No data	No data	—	—
Proportion of patients with adverse effects (e.g. hyponatraemia, mood disturbance and headaches)	No data	No data	No data	No data	—	—
Proportion of patients with loss or gain of hearing/reduction in progression of hearing loss	No data	No data	No data	No data	—	—
Proportion of patients with a reduction in the severity of tinnitus	No data	No data	No data	No data	—	—
Proportion of patients with a reduction in the perception of aural fullness	No data	No data	No data	No data	—	—
Overall changes in well-being and quality of life	No data	No data	No data	No data	—	—
Any other adverse effects	No data	No data	No data	No data	—	—

\*The basis for the **assumed risk** (e.g. the median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).

**CI:** confidence interval

---

GRADE Working Group grades of evidence

**High quality:** Further research is very unlikely to change our confidence in the estimate of effect.

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

**Low quality:** 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.

**Very low quality:** We are very uncertain about the estimate.

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## BACKGROUND

### Description of the condition

Ménière's disease or syndrome is a chronic inner ear disorder that results in sporadic attacks of vertigo, sensorineural hearing loss, aural fullness and tinnitus. The term 'Ménière's disease' refers to an idiopathic disorder, but a clinically identical presentation can occur secondary to other conditions, such as infections, genetic disorders or trauma, and this is referred to as 'Ménière's syndrome'.

In a large US study the prevalence of Ménière's disease was estimated at 200 per 100,000 people (Alexander 2010). It most commonly affects people between the ages of 40 and 60 years (Harcourt 2014), with a slight female preponderance (da Costa 2002). Acute attacks usually occur in clusters, with those affected often being symptom-free for months in between. There is a higher frequency of attacks in the initial period after presentation with an eventual reduction but sustained deterioration in hearing (Moffat 1997). The vertiginous episodes cease eventually (Silverstein 1989). However, there can be considerable persistent disability and the economic costs of the condition are estimated to be significant (Salvador-Carulla 2010; Tyrrell 2016).

At present no 'gold standard' diagnostic test for Ménière's disease exists and the diagnosis is frequently based on the practitioner's assessment of the patient's history and neurotologic evaluation. The American Academy of Otolaryngology – Head and Neck Surgery (AAO-HNS) has produced diagnostic guidelines (Alford 1972), which have since been revised twice (Ménière's Guide 1995; Pearson 1985). The guidelines state that a diagnosis can be made if the following criteria are met:

- at least two spontaneous episodes of rotational vertigo lasting at least 20 minutes;
- audiometric confirmation of a sensorineural hearing loss;
- tinnitus and/or a perception of aural fullness.

When patients meet the AAO-HNS criteria and the symptoms are attributed to a specific cause they are classified as having Ménière's syndrome.

Although the pathophysiology of Ménière's disease is unknown, there is thought to be an association with endolymphatic hydrops, which is distortion of the membranous labyrinth due to the over accumulation of endolymph (Hallpike 1938).

At present there is no definitive treatment for Ménière's disease and treatment options range from dietary modification through medication (for example, betahistine (James 2001) or diuretics (Burgess 2006)) to intratympanic injections (Pullens 2011; Phillips 2011) or surgery (Pullens 2013).

### Description of the intervention

Modification of diet can include the restriction of salt, caffeine and alcohol intake.

Dietary salt restriction is a potentially simple and relatively inexpensive option that is widely recommended to patients with Ménière's as a first-line treatment. The level and duration of salt intake restriction that is advised can vary.

Other additional dietary modifications include limiting the intake of caffeine (Luxford 2013) and alcohol (Ménière's Society). Caffeine is a commonly ingested substance found in liquid form in beverages such as tea and coffee, and it is also found in food such as chocolate. Alcohol is widely consumed, with reported figures of 7.8 litres per person per year on average throughout the adult population in 2015 in the UK (Office of National Statistics). As with dietary salt restriction the level and duration of caffeine and or alcohol restriction advised may vary.

### How the intervention might work

Disturbance of the volume and/or electrolyte composition of the endolymph is considered to be the cause of the symptoms experienced by patients with Ménière's disease. High dietary intake of salt can affect the concentrations of electrolytes in the blood, which in turn affects the composition of the endolymph. It has been reported that high salt intake can contribute to attacks and it therefore follows that dietary modification can be used to control both the volume and composition of the endolymph (Stahle 1984). Fluctuation in the composition and volume of the endolymph is considered to contribute to the fluctuating nature of the symptoms experienced by sufferers of Ménière's. The observation that water retention can exacerbate the symptoms of Ménière's disease was first documented in 1929 (Dederding 1929). Subsequent uncontrolled studies suggested that the manipulation of salt intake influences the symptoms experienced by those suffering with Ménière's (Furstenberg 1934; Furstenberg 1941). More recent studies of restricted salt intake, usually together with other treatment modalities, such as the use of diuretics, have also suggested better symptom control in patients with Ménière's disease (Klockhoff 1974; Santos 1993). Excessive reduction in salt intake may, however, in extreme and rare cases result in hyponatraemia, although this is more commonly due to specific diseases. Hyponatraemia is associated with conditions ranging from mood disturbance to cerebral oedema and possible death in extreme cases (Thompson 2010).

Alcohol and caffeine, in high concentrations, can both result in vasoconstriction and a reduction in the blood supply to the inner ear, which can exacerbate the symptoms of sufferers. Dietary restriction of these substances may therefore be beneficial in Ménière's patients. Caffeine, however, is a recognised ergogenic aid even at physiological levels and enhances concentration and alertness whilst reducing fatigue. A reduction in the intake of caffeine may therefore lead to withdrawal effects in individuals who are accustomed to its effects, which can result in symptoms ranging from mood disturbance to headaches (Pesta 2013). It is well recognised to have clinical effects on balance and known to produce symptoms of vertigo and vomiting when taken in sufficient doses, reminiscent of an episode of Ménière's. Alcohol also causes vasoconstriction and therefore acts in a similar fashion to caffeine. However, to date no studies have been published that specifically address the effect of reducing/omitting alcohol in the treatment of Ménière's disease, as is routinely recommended. Although no recognised adverse effects of reducing alcohol consumption in individuals with consumption within recommended limits have been documented, a reduction in people who have a dependency on alcohol can result in withdrawal symptoms such as psychotic episodes and delirium tremens (Stern 2010).

## Why it is important to do this review

Dietary modification, including the restriction of salt, caffeine and alcohol, is widely recommended to those suffering with Ménière's and generally considered the first-line treatment. There has been no previous systematic review of the evidence for this advice, which is routinely given to patients. This may delay the use of more appropriate and effective treatments. Restricting the intake of salt, caffeine and alcohol, substances which people may find palatable and that potentially provide a degree of pleasure to their daily lives, may also have an effect on mood or quality of life, with no potential benefit. Dietary modification could also have potentially negative implications for social, work and family life. It is therefore important to conduct a systematic review of randomised controlled trials of these dietary modifications in patients with Ménière's disease or syndrome.

## OBJECTIVES

To assess the effects of dietary restriction of salt, caffeine and alcohol intake in patients with Ménière's disease or syndrome.

## METHODS

### Criteria for considering studies for this review

#### Types of studies

Randomised and quasi-randomised controlled trials. We planned to include cluster-randomised and cross-over trials if identified.

#### Types of participants

Adult patients, aged 18 and over, with a diagnosis of Ménière's disease or syndrome.

We planned to classify studies according to the diagnostic criteria used to diagnose Ménière's disease or syndrome, grading those using the AAO-HNS criteria, or equivalent, to define probable, definite or certain Ménière's as grade 'I' studies and the remaining studies as grade 'II'.

Settings included both the community and hospital.

#### Types of interventions

The intervention of interest was dietary modification: specifically, salt, caffeine and/or alcohol restriction or substitution (or both). The control intervention was no modification.

The main comparison pairs were:

- dietary restriction of salt *versus* no restriction;
- dietary restriction of caffeine *versus* no restriction;
- dietary restriction of alcohol *versus* no restriction.

Other possible comparison pairs included:

- dietary restriction of salt *versus* dietary restriction of caffeine;
- dietary restriction of salt *versus* dietary restriction of alcohol;
- dietary restriction of alcohol *versus* dietary restriction of caffeine;
- dietary restriction of salt + caffeine *versus* no restriction;
- dietary restriction of salt + alcohol *versus* no restriction;
- dietary restriction of alcohol + caffeine *versus* no restriction.

We planned to include studies where multiple dietary restrictions were used in conjunction but to note this in the analysis. We expected that there would be variability in the level and duration of dietary modification, but we planned to deal with this in subgroup analysis.

We excluded studies where dietary modification plus another treatment modality (e.g. a pharmacological agent) was used due to the potential for interactive effects.

### Types of outcome measures

We analysed the following outcomes in the review, but we did not use them as a basis for including or excluding studies.

#### Primary outcomes

- Control of vertigo or decrease in the intensity, frequency and duration of vertigo attacks, as suggested by the AAO-HNS\* (*Ménière's Guide 1995*), using the results of the various questionnaire-based assessment tools including the Vertigo Symptom Scale, the Vertigo Dizziness Imbalance Questionnaire and the Dizziness Handicap Inventory amongst others.
- Adverse effects; any reported adverse effects such as hyponatraemia, mood disturbance and headaches would be noted.

#### Secondary outcomes

- Hearing: change in hearing loss or its progression (or both) as measured by a pure-tone audiogram. The proportion of patients with progression of hearing loss (more than 15 dB), based on the four-tone average of thresholds at 0.5 kHz, 1 kHz, 2 kHz and 3 kHz, as measured by a pure-tone audiogram.
- Tinnitus: the proportion of patients with a reduction in tinnitus as measured by using patient-reported questionnaire scores (the Tinnitus Handicap Index (THI), the Tinnitus Functional Index, the Tinnitus Handicap Questionnaire, the Tinnitus Questionnaire, the Tinnitus Reaction Questionnaire and the Tinnitus Severity Scale).
- Perception of aural fullness: the proportion of patients with reduction of aural fullness as measured by using patient-reported questionnaire scores such as the Vertigo Dizziness Imbalance Questionnaire or a visual analogue scale.
- Well-being and quality of life: overall changes, as reported in the Dizziness Handicap Inventory questionnaire and any other appropriate scale.
- Any other adverse effects; these would be noted based on patient-reported symptoms of potential unpleasant effects, as well as the results of specific questionnaires including the Vertigo Symptom Scale, the Vertigo Dizziness Imbalance Questionnaire and the Dizziness Handicap Inventory amongst others.

\*The AAO-HNS Committee on Hearing and Equilibrium proposed the "control of vertigo" as a main objective outcome measure when assessing therapy in Ménière's disease. The number of attacks six months prior to treatment is compared to the number in the period between 18 and 24 months following treatment. The resulting number indicates the extent of "control of vertigo". The AAO-HNS further divides the control of vertigo into classes, where Class A (CoV = 0) is complete control and class B (CoV 1 to 40) is substantial control. They recommend a period of at least two years of follow-



up in order to assess fully the effect of the intervention. We also considered studies with shorter periods of follow-up for this review.

We also anticipated various questionnaire-based assessment tools being used in the different studies, including the Vertigo Symptom Scale (VSS), the Vertigo Dizziness Imbalance Questionnaire and the Dizziness Handicap Inventory amongst others. We included all forms of questionnaire that address the patient's perception of their symptoms, if used consistently. These questionnaires enable us to assess the impact on the patients' quality of life, functional impairment and disability.

### Search methods for identification of studies

The Cochrane ENT Information Specialist conducted systematic searches for randomised controlled trials and controlled clinical trials. There were no language, publication year or publication status restrictions. The date of the search was 28 March 2018.

#### Electronic searches

Published, unpublished and ongoing studies were identified by searching the following databases from their inception:

- Cochrane ENT Trials Register (searched via the Cochrane Register of Studies 28 March 2018);
- Cochrane Central Register of Controlled Trials (CENTRAL 2018, Issue 4);
- PubMed (1946 to 29 March 2018);
- Ovid EMBASE (1974 to 29 March 2018);
- EBSCO CINAHL (1982 to 29 March 2018);
- Ovid AMED (1985 to 29 March 2018);
- Ovid CAB abstracts (1910 to 29 March 2018);
- LILACS (searched 29 March 2018);
- KoreaMed (searched via Google Scholar 29 March 2018);
- IndMed (searched 29 March 2018);
- PakMediNet (searched 29 March 2018);
- Web of Knowledge, Web of Science (1945 to 29 March 2018);
- CNKI (searched via Google Scholar 29 March 2018);
- ClinicalTrials.gov ([www.clinicaltrials.gov](http://www.clinicaltrials.gov)) (searched 29 March 2018);
- World Health Organization (WHO) International Clinical Trials Registry Platform (ICTRP), [www.who.int/ictip](http://www.who.int/ictip) (searched 29 March 2018);
- ISRCTN, [www.isrctn.com](http://www.isrctn.com) (searched 29 March 2018).

The Information Specialist modelled subject strategies for databases on the search strategy designed for CENTRAL. Where appropriate, they were combined with subject strategy adaptations of the highly sensitive search strategy designed by Cochrane for identifying randomised controlled trials and controlled clinical trials (as described in the *Cochrane Handbook for Systematic Reviews of Interventions* Version 5.1.0, Box 6.4.b. ([Handbook 2011](#))). Search strategies for major databases including CENTRAL are provided in [Appendix 1](#).

#### Searching other resources

We scanned the reference lists of identified publications for additional trials and contacted trial authors where necessary. In addition, the Information Specialist searched PubMed to retrieve existing systematic reviews relevant to this systematic review, so

that we could scan their reference lists for additional trials. The Information Specialist also ran non-systematic searches of Google Scholar to retrieve grey literature and other sources of potential trials.

### Data collection and analysis

#### Selection of studies

Two authors (KH and LM) independently scanned the initial search results to identify studies that appeared to meet the inclusion criteria. We used abstract review to eliminate any studies that were clearly ineligible. If either author identified a paper as potentially suitable, we reviewed the full text of the article. We resolved disagreements by discussion or, failing that, with the input of the third author (AS).

#### Data extraction and management

Two authors (KH and LM) extracted data independently. We used standardised data extraction forms. There was no blinding of journal, author names or affiliations.

For each study, we planned to extract the following information:

- study design;
- duration of study;
- randomisation;
- allocation concealment;
- number of participants;
- setting of study;
- diagnostic criteria;
- exclusion criteria;
- age and sex distribution of participants;
- country of recruitment;
- co-morbidity;
- date of study;
- number of intervention groups;
- type of dietary modification;
- outcomes measured and definition of outcomes;
- missing data and final sample size.

#### Assessment of risk of bias in included studies

KH and LM planned to undertake assessment of the risk of bias of the included studies independently, with the following taken into consideration, as guided by the *Cochrane Handbook for Systematic Reviews of Interventions* ([Handbook 2011](#)):

- sequence generation;
- allocation concealment;
- blinding;
- incomplete outcome data;
- selective outcome reporting; and
- other sources of bias.

We planned to use the Cochrane 'Risk of bias' tool in RevMan 5.3 ([RevMan 2014](#)), which involves describing each of these domains as reported in the study and then assigning a judgement about the adequacy of each entry: 'low', 'high' or 'unclear' risk of bias.

We also intended to judge two extra domains: the certainty of the diagnosis of Ménière's (see also [Types of participants](#)) and the quality of outcome assessment (see [Types of outcome measures](#)). However, we would have reported and addressed these domains in the 'Characteristics of included studies' table rather than consider them as a risk of bias domain.

### Measures of treatment effect

We planned to use appropriate statistical tests based on the data. For dichotomous data we planned to calculate an odds ratio (OR), risk ratio (RR) and risk difference (RD, also called absolute risk reduction), as well as the number of participants needed to treat to avoid a case of the disease (number needed to treat to benefit - NNTB) from the pooled results, based on the median risk in the control groups.

For the intervention effect measures of continuous data we planned to calculate the difference in means (mean difference, MD) between the groups, provided that different studies were using the same scale of measurement. We planned to calculate the standardised mean difference (SMD) if different scales were used.

For ordinal data we planned to check to see whether the scale used had been validated. Depending on the number points in these scales (and how the data were reported), we planned either to dichotomise these or analyse them as continuous outcomes.

### Unit of analysis issues

#### Cluster-randomised trials

Cluster-randomised trials allocate groups instead of individuals. The participants in each group may be related in some way, therefore this needs to be taken into account in the analysis otherwise there is a unit of analysis error, which would produce an artificially smaller P value and a risk of false positive results. For this purpose we planned to use a special statistical method, as detailed in chapter 16.3 of the *Cochrane Handbook for Systematic Reviews of Interventions* ([Handbook 2011](#)), with the appropriate statistical advice.

#### Cross-over trials

Cross-over trials may have a carry-over effect. For this reason we planned to use data from cross-over trials only if data from before the cross-over could be obtained.

#### Multi-arm studies

If we had found studies with more than two groups (e.g. two or more active treatments being tested against placebo), we would have established which of the comparisons were relevant to the systematic review and relevant to each of the meta-analyses that we may implement. If the study design used independent groups, we would have treated the study as having independent comparisons. However, if we had encountered participants that had been included in several groups, there would be a risk of unit of analysis error and we planned to ensure that participants were only included once per meta-analysis as per chapter 16.5 of the *Cochrane Handbook for Systematic Reviews of Interventions* ([Handbook 2011](#)).

### Repeated observations on participants

In longer studies, results may be recorded at more than one time interval. In order to avoid unit of analysis error when combining these results in a single meta-analysis (and therefore counting the same participants in more than one comparison), we aimed to retrieve individual patient data and perform a time-to-event analysis using the whole follow-up for each participant.

We also aimed to establish short-term (three months), medium-term (12 months) and long-term (over 24 months) effects.

### Dealing with missing data

When the required data were not available in published accounts, we planned to contact the principal investigator to request the data. If no useful response could be obtained, we planned to treat missing data differently if they were judged to be 'missing at random', in which case the effect may not be important, or 'not missing at random', where missing data may affect the overall result. In the first case, the data can be ignored. If large numbers of dropouts were found, we planned to conduct sensitivity analysis with different assumptions.

We were alert to potential mislabelling or non-identification of standard errors and standard deviations. Unless missing standard deviations could be derived from confidence intervals we would not have assumed values for analysis purposes.

### Assessment of heterogeneity

We planned to assess studies for clinical, statistical and methodological heterogeneity. If sufficient non-heterogeneous studies were found, we planned to subject the data to a meta-analysis with a fixed-effect model where applicable. If there was statistical heterogeneity, we planned to use random-effects modelling. If the level of heterogeneity and the appropriate model that should be used was unclear, we planned to take statistical advice.

### Assessment of reporting biases

If an individual meta-analysis contained at least 10 studies, we would have assessed publication bias using funnel plots and Egger's test.

### Data synthesis

Analysis was to be on an intention-to-treat basis. If the data were compatible we planned to combine data to give summary measures of effect. If data were missing we planned to use available case analysis – using all data (as reported) for all randomised patients available at the end of the trial/time point of interest, regardless of actual treatment received.

### Subgroup analysis and investigation of heterogeneity

We planned to undertake subgroup analysis based on:

- patients meeting *versus* not meeting the AAO-HNS criteria for the diagnosis of Ménière's disease or syndrome;
- the treatment protocol; the subgroups would be based on the specific duration:
  - \* complete *versus* partial restriction of the dietary modification of interest *versus* modification of multiple simultaneous

dietary modifications. This latter group may also be further subdivided based on the level of restriction.

### Sensitivity analysis

We planned to conduct a sensitivity analysis by comparing the effect of the inclusion and exclusion of studies with different risk of bias. If we deemed studies to have a high risk of bias, we would have excluded them from the analysis.

### GRADE and 'Summary of findings' table

We planned to use the GRADE approach to rate the overall quality of evidence. The quality of evidence reflects the extent to which we are confident that an estimate of effect is correct and we planned to apply this in the interpretation of results. There are four possible ratings: high, moderate, low and very low. A rating of high quality of evidence implies that we are confident in our estimate of effect and that further research is very unlikely to change our confidence in the estimate of effect. A rating of very low quality implies that any estimate of effect obtained is very uncertain.

The GRADE approach rates evidence from RCTs that do not have serious limitations as high quality. However, several factors can lead to the downgrading of the evidence to moderate, low or very low. The degree of downgrading is determined by the seriousness of these factors:

- study limitations (risk of bias);
- inconsistency;
- indirectness of evidence;
- imprecision; and
- publication bias.

We planned to include a 'Summary of findings' table, constructed according to the recommendations described in Chapter 10 of

the *Cochrane Handbook for Systematic Reviews of Interventions* ([Handbook 2011](#)). We would have included the following outcomes in our 'Summary of findings' table(s):

- proportion of patients with control of vertigo or decrease in vertigo attacks (as suggested by the AAO-HNS);
- proportion of patients with adverse effects (e.g. hyponatraemia, mood disturbance and headaches);
- proportion of patients with loss or gain of hearing/reduction in progression of hearing loss;
- proportion of patients with a reduction in the severity of tinnitus;
- proportion of patients with a reduction in the perception of aural fullness;
- overall changes in well-being and quality of life;
- any other adverse effects.

## RESULTS

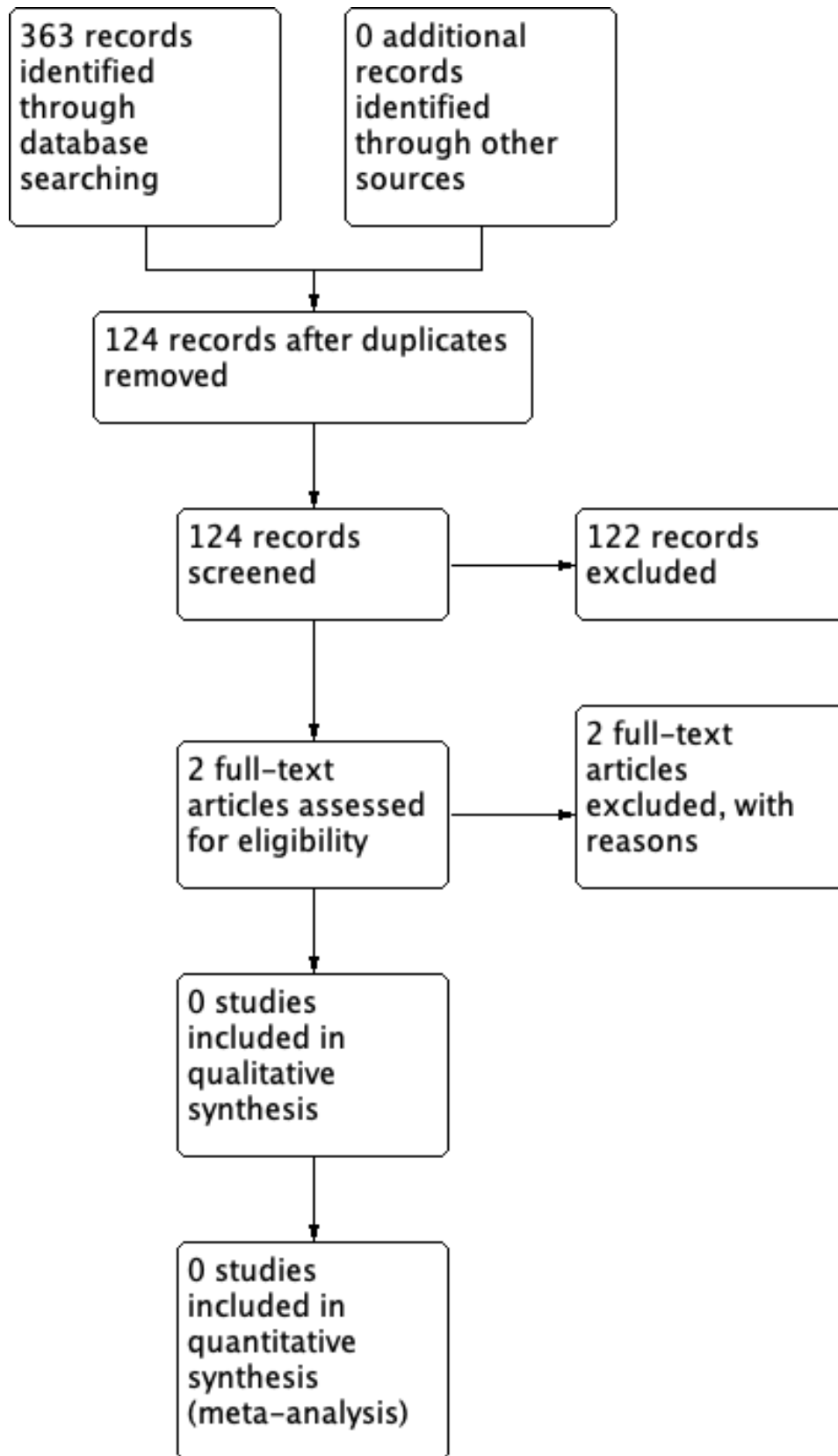
### Description of studies

#### Results of the search

The searches retrieved 363 records including papers, reviews, conference abstracts and registered clinical trials, of which 124 remained after removing duplicates. Review of the titles and abstracts identified two potential studies for full-text review. Subsequently, we formally excluded these two studies for the reasons given in the [Characteristics of excluded studies](#) table. We identified no ongoing studies and there are no studies awaiting assessment. No studies met the inclusion criteria for the review.

Full details of the search and study selection process are shown in a PRISMA flowchart ([Figure 1](#)).

**Figure 1. Process for sifting search results and selecting studies for inclusion.**



## Included studies

We did not identify any studies that met the inclusion criteria for the review.

## Excluded studies

We excluded two studies after a full-text review (see [Characteristics of excluded studies](#)).

[Acharya 2017](#) was a double-blind randomised controlled trial in the outpatient department of an ENT unit in a teaching hospital in Nepal. Patients who were diagnosed with Ménière's disease were randomised into three groups: dietary salt restriction, diuretics or a vasodilator. Outcome measures included the number and severity of vertigo attacks, tinnitus and hearing loss. We excluded this study because the control groups were both active interventions. The study did not have a no treatment (no dietary restriction) or placebo control group.

[Bittar 2016](#) was a randomised controlled trial evaluating the effect of a fractionated diet without glucose as the treatment for labyrinthine disorders associated with the glucose insulin index. We excluded this study because it excluded patients with Ménière's disease and did not assess the restriction of salt, caffeine or alcohol.

## Risk of bias in included studies

We did not identify any studies that met the inclusion criteria.

## Effects of interventions

See: [Summary of findings for the main comparison](#)

There were no studies meeting our inclusion criteria. See [Summary of findings for the main comparison](#).

## DISCUSSION

### Summary of main results

We were unable to identify any randomised controlled trials or quasi-randomised controlled trials investigating the role of dietary modification (restriction of salt, caffeine and/or alcohol) in the treatment of Ménière's disease or syndrome.

Ménière's disease is a debilitating and progressive condition that can result in the affected person being unable to carry out acts of daily living and can lead to the development of irreversible hearing loss. Modification of diet, including restriction of salt, caffeine and alcohol intake, is an option that is widely recommended to patients with Ménière's as a first-line treatment, and is in fact featured on both the UK National Health Service (NHS) and Ménière's Society websites ([Ménière's Society](#); [NHS 2017](#)). However, it is evident that there is no high-level evidence on which to base these recommendations. The question was prioritised by a James Lind Alliance patient and public involvement (PPI) initiative in 2011 ([JLA 2011](#)), but still cannot be answered.

### Overall completeness and applicability of evidence

There are no studies to include in this review.

### Quality of the evidence

There is no evidence meeting our inclusion criteria to assess.

## Potential biases in the review process

We conducted this review according to a published protocol ([Hussain 2016](#)). The search is comprehensive and up to date as of March 2018. We have not identified any specific biases in our review process.

## Agreements and disagreements with other studies or reviews

This is the first systematic review that sought to identify randomised controlled trials investigating the role of dietary modification of salt, caffeine and alcohol in the treatment of Ménière's disease. However, other studies have been conducted in an attempt to investigate the effect on the disease. Most recently, [Acharya 2017](#) addressed the role of diet in the control of symptoms in Ménière's disease. Patients were required to meet the American Academy of Otolaryngology – Head and Neck Surgery (AAO-HNS) diagnostic criteria and they were randomly allocated into the trial. The treatment arms were dietary salt restriction (plus vitamin B complex daily as a placebo) versus 5 mg of amiloride and 40 mg of furosemide, versus 24 mg of betahistine. There was no advice about intake of salt for participants in these latter two arms. The study also did not include a no treatment (no dietary restriction) or placebo control group. [Bittar 2016](#) addressed the impact on vestibular disorders of a fractionated diet with glucose restriction versus no restriction at all. This study, however, excluded patients with Ménière's disease/syndrome and did not look specifically at the impact of restricting salt, caffeine or alcohol intake.

We found one review article that looked at the role of caffeine in the causation or aggravation of various otolaryngology conditions, including Ménière's. The authors were unable to identify any evidence to support or refute this association ([Trinidad 2014](#)).

## AUTHORS' CONCLUSIONS

### Implications for practice

There is no evidence from randomised controlled trials (RCTs) to support or refute the restriction of salt intake or other dietary modifications for patients with Ménière's disease or syndrome. There are therefore no implications for the alteration of current practice.

### Implications for research

High-quality research into the restriction of salt and other dietary modifications for the treatment of Ménière's disease or syndrome is warranted. This intervention is widely recommended to patients without any proven benefit or clear understanding of any potential harms. This may delay the use of more effective treatment options resulting in disease progression and patient suffering or adverse effects. If shown to be effective, however, this is a cost-neutral treatment option for the medical treatment provider. There is also undoubtedly interest from patients in the potential effectiveness of dietary modifications. The James Lind Alliance patient-professional priority-setting group identified the efficacy of dietary intervention for Ménière's disease as one of the top 10 priorities for research into balance disorders ([JLA 2011](#)).

It must be acknowledged there has been a move away from the American Academy of Otolaryngology – Head and Neck Surgery (AAO-HNS) diagnostic criteria within the Ménière's

disease community since the start of this review process. The Bárány Society represents an international collaboration of basic scientists, otolaryngologists, oto-neurologists, physical therapists and other allied health professionals considered experts in the field. A classification committee for an International Classification for Vestibular Disorders (ICVD) was established by the society and developed international consensus diagnostic criteria for Ménière's disease. They worked with the Equilibrium Committee of the AAO-HNS as part of their process and defined two sets of diagnostic criteria: definite and probable Ménière's disease (Lopez-Escamez 2016). In view of the fact that the Bárány Society criteria are not dissimilar to the AAO-HNS definition, we do not believe that our search results will be materially affected and therefore we opted not to repeat the search or change the scope of this review. However, future research should adhere to the newer Bárány Society diagnostic criteria. Any future trial should also be conducted and reported according to the CONSORT statement.

There are methodological challenges with conducting research into dietary interventions in Ménière's disease. Appropriate placebo groups can be difficult to construct for dietary restrictions. Adherence to diets should be assessed objectively where possible, for example by measurement of urinary sodium levels in the case of low salt diets. Comparison of high and low levels of salt intake has been successfully accomplished, for example when looking at salt restriction for the management of hypertension (Graudal 2017). Unlike hypertension, however, Ménière's disease is relatively uncommon (Murdin 2015), and it has a relapsing

and remitting natural history. Conducting an adequately powered study of a dietary intervention over a suitable time period is therefore challenging. The best evidence may come from a RCT comparing dietary interventions (e.g. low salt versus general healthy diet advice). However, this research question might be more pragmatically addressed by using information from carefully constructed patient registries that include information on dietary intake of substances of interest such as salt, caffeine and alcohol. It would be important to address the question of any possible harms or unwanted effects of dietary modification.

It must, however, also be acknowledged that a Cochrane-style review may not be appropriate for addressing this question (Truswell 2005). Adherence to dietary modification is challenging to implement and assess. However, this does not take away from the need for an appropriate RCT to be conducted.

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## CHARACTERISTICS OF STUDIES

### Characteristics of excluded studies [ordered by study ID]



Study	Reason for exclusion
<a href="#">Acharya 2017</a>	ALLOCATION: Randomised controlled trial PARTICIPANTS: Patients with Ménière's disease according to AAO-HNS 1995 guidelines INTERVENTION: Dietary salt restriction (plus vitamin B complex daily as a placebo) versus amiloride 5 mg and furosemide 40 mg (no advice about intake of salt) versus betahistine 24 mg (no advice about intake of salt); study did not include a no treatment (no dietary restriction) or placebo control group
<a href="#">Bittar 2016</a>	ALLOCATION: Randomised controlled trial PARTICIPANTS: Patients with vestibular disorders: patients with Ménière's disease were excluded INTERVENTION: Fractionated diet with glucose restriction versus no restriction: not salt, caffeine or alcohol restriction

AAO-HNS: American Academy of Otolaryngology – Head and Neck Surgery

## APPENDICES

### Appendix 1. Search strategies

CENTRAL	PubMed	Embase	CINAHL
#1 MeSH descriptor: [Endolymphatic Hydrops] explode all trees	#1 Search "Endolymphatic Hydrops"[Mesh]	1 exp Meniere disease/	S40 S38 OR S39
#2 (meniere or menieres or meniere's):ti,ab,kw	#2 Search meniere*[Title/Abstract]	2 "meniere*".ti,ab.	S39 (MH "Endolymphatic Hydrops+/DH")
#3 ((endolymphatic or cochlea*) and hydrops):ti,ab,kw	#3 Search ((endolymphatic[Title/Abstract] OR cochlea*[Title/Abstract])) AND hydrops[Title/Abstract]	3 ((endolymphatic or cochlea*) and hydrops).ti,ab.	S38 S25 AND S37
#4 ((aural or labyrinth*) and (hydrops or syndrome or vertigo)):ti,ab,kw	#4 Search ((aural[Title/Abstract] OR labyrinth*[Title/Abstract])) AND (hydrops[Title/Abstract] OR syndrome[Title/Abstract] OR vertigo[Title/Abstract])	4 ((aural or labyrinth*) and (hydrops or syndrome or vertigo)).ti,ab.	S37 S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S33 OR S34 OR S35 OR S36
#5 #1 or #2 or #3 or #4	#5 #1 OR #2 OR #3 OR #4	5 exp sodium restriction/	S36 MW "DH"
#6 MeSH descriptor: [Diet, Sodium-Restricted] explode all trees	#6 Search "Diet, Sodium-Restricted"[Mesh]	6 exp feeding behavior/	S35 TX nutrition*
#7 MeSH descriptor: [Feeding Behavior] this term only	#7 Search "Diet, Sodium-Restricted"[Mesh]	7 diet therapy/	S34 TX (diet* or food) and (restrict* or modif* or habit* or free*)
		8 exp sodium intake/	S33 TX Caffein* or decaffein* or "de caffein*" or de-caffein* or coffee or tea or alcohol*
		9 exp caffeine/	
		10 exp drinking behavior/	

(Continued)

#8 MeSH descriptor: [Diet Therapy] this term only	#7 Search "Feeding Behavior"[Mesh:NoExp]	11 (salt* or sodium*).ti,ab.	S32 TX salt* or sodium*
#9 MeSH descriptor: [Sodium, Dietary] explode all trees	#8 Search "Diet Therapy"[Mesh:NoExp]	12 (Caffein* or decaffein* or "de caffein*" or de-caf-fein* or coffee or tea or alcohol*).ti,ab.	S31 (MH "Drinking Behavior+") S30 (MH "Caffeine") S29 (MH "Sodium, Dietary+")
#10 MeSH descriptor: [Caffeine] explode all trees	#9 Search "Sodium, Dietary"[Mesh]	13 ((diet* or food) and (restrict* or modif* or habit* or free*)).ti,ab.	S28 (MH "Diet Therapy") S27 (MH "Food Habits")
#11 MeSH descriptor: [Drinking Behavior] explode all trees	#10 Search "Caffeine"[Mesh]	14 "nutrition*".ti,ab.	S26 (MH "Diet, Sodium-Restricted")
#12 (salt* or sodium*):ti,ab,kw	#11 Search "Drinking Behavior"[Mesh]	15 1 or 2 or 3 or 4	S25 S21 OR S22 OR S23 OR S24
#13 (Caffein* or decaffein* or "decaffein*" or de-caf-fein* or coffee or tea or alcohol*):ti,ab,kw	#12 Search (salt*[Title/Abstract] OR sodium*[Title/Abstract])	16 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14	S24 TX ((aural or labyrinth*) and (hydrops or syndrome or vertigo)) S23 TX ((endolymphatic or cochlea*) and hydrops)
#14 ((diet* or food) and (restrict* or modif* or habit* or free*)):ti,ab,kw	#13 Search (Caffein*[Title/Abstract] OR decaffein*[Title/Abstract] OR de-caf-fein*[Title/Abstract] OR coffee[Title/Abstract] OR tea[Title/Abstract] OR alcohol*[Title/Abstract])	17 15 and 16	S22 TX meniere* S21 (MH "Endolymphatic Hydrops+")
#15 nutrition*:ti,ab,kw	#14 Search ((diet*[Title/Abstract] OR food[Title/Abstract]) AND (restrict*[Title/Abstract] OR modif*[Title/Abstract] OR habit*[Title/Abstract] OR free*[Title/Abstract]))		S20 S18 OR S19
#16 Any MeSH descriptor with qualifier(s): [Diet therapy - DH]	#15 Search nutrition*[Title/Abstract]		S19 (MH "Endolymphatic Hydrops+/DH") S18 S5 AND S17
#17 #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16	#16 Search Diet therapy[sh]		S17 S6 OR S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14 OR S15 OR S16
#18 #5 and #17	#17 Search (#6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16)		S16 MW "DH"
#19 MeSH descriptor: [Endolymphatic Hydrops] explode all trees and with qualifier(s): [Diet therapy - DH]	#18 (#5 and #17)		S15 TX nutrition*
#20 #18 or #19	#19 Search "Endolymphatic Hydrops/diet therapy"[Mesh]		S14 TX (diet* or food) and (restrict* or modif* or habit* or free*)
	#20 (#18 AND #19)		S13 TX Caffein* or decaffein* or "de caffein*" or de-caf-fein* or coffee or tea or alcohol* S12 TX salt* or sodium* S11 (MH "Drinking Behavior+") S10 (MH "Caffeine") S9 (MH "Sodium, Dietary+") S8 (MH "Diet Therapy") S7 (MH "Eating Behavior") S6 (MH "Diet, Sodium-Restricted")

(Continued)

S5 S1 OR S2 OR S3 OR S4

S4 TX ((aural or labyrinth\*) and (hydrops or syndrome or vertigo))

S3 TX ((endolymphatic or cochlea\*) and hydrops)

S2 TX meniere\*

S1 (MH "Endolymphatic Hydrops+")

Web of Science	ICTRP	ClinicalTrials.gov	LILACS
#1 TOPIC: (meniere*)	meniere* AND salt* OR meniere* AND sodium* OR meniere* AND caffein* OR meniere* AND alcohol	<b>Via the Cochrane Register of Studies</b>	(TW:Endolymphatic AND TW:Hydrops) OR (TW:Hidropesía AND TW:Endolinfática) OR TW:meniere*
#2 TOPIC: ((endolymphatic or cochlea*) and hydrops)		1 (meniere or menieres or meniere's) AND INSEGMENT	AND
#3 TOPIC: ((aural or labyrinth*) and (hydrops or syndrome or vertigo))		2 (endolymphatic or cochlea*) and hydrops AND INSEGMENT	Controlled Clinical Trials filter
#4 #3 OR #2 OR #1		3 (aural or labyrinth*) and (hydrops or syndrome or vertigo) AND INSEGMENT	
#5 TOPIC: (salt* or sodium* or Caffein* or decaffein* or "de caffein*" or de-caffein* or coffee or tea or alcohol*)		4 #1 OR #2 OR #3 AND INSEGMENT	
#6 TOPIC: ((diet* or food) and (restrict* or modif* or habit* or free*))		5 salt* or sodium* AND INSEGMENT	
#7 TOPIC: (nutrition*)		6 Caffein* or decaffein* or "de caffein*" or de-caffein* or coffee or tea or alcohol* AND INSEGMENT	
#8 #7 OR #6 OR #5		7 (diet* or food) and (restrict* or modif* or habit* or free*) AND INSEGMENT	
#9 #8 AND #4		8 nutrition* AND INSEGMENT	
		9 #5 OR #6 OR #7 OR #8 AND INSEGMENT	
		10 #4 AND #9 AND INSEGMENT	
		11 (nct*):AU AND INSEGMENT	
		12 #10 AND #11	
		<b>Via Clinicaltrials.org</b>	
		(menieres OR meniere's) AND (salt OR sodium OR caffeine OR alcohol)	

**Search strategies for CENTRAL, PubMed and CINAHL for searches in July 2016 and July 2017**

 See explanation in [Differences between protocol and review](#).

CENTRAL	PubMed	CINAHL
#1 MeSH descriptor: [Endolymphatic Hydrops] explode all trees	#1 Search "Endolymphatic Hydrops"[Mesh]	S40 S38 OR S39
#2 (meniere or menieres or meniere's):ti,ab,kw	#2 Search meniere*[Title/Abstract]	S39 (MH "Endolymphatic Hydrops+/DH")
#3 ((endolymphatic or cochlea*) and hydrops):ti,ab,kw	#3 Search ((endolymphatic[Title/Abstract] OR cochlea*[Title/Abstract])) AND hydrops[Title/Abstract]	S38 S25 AND S37
#4 ((aural or labyrinth*) and (hydrops or syndrome or vertigo)):ti,ab,kw	#4 Search ((aural[Title/Abstract] OR labyrinth*[Title/Abstract])) AND (hydrops[Title/Abstract] OR syndrome[Title/Abstract] OR vertigo[Title/Abstract])	S37 S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S33 OR S34 OR S35 OR S36
#5 #1 or #2 or #3 or #4	#5 (#1 OR #2 OR #3 OR #4)	S36 MW "DH"
#6 MeSH descriptor: [Diet, Sodium-Restricted] explode all trees	#6 Search "Diet, Sodium-Restricted"[Mesh]	S35 TX nutrition*
#7 MeSH descriptor: [Food Habits] explode all trees	#7 Search "Food Habits"[Mesh]	S34 TX (diet* or food) and (restrict* or modif* or habit* or free*)
#8 MeSH descriptor: [Diet Therapy] this term only	#8 Search "Diet Therapy"[Mesh:NoExp]	S33 TX Caffein* or decafein* or "de cafein*" or de-cafein* or coffee or tea or alcohol*
#9 MeSH descriptor: [Sodium, Dietary] explode all trees	#9 Search "Sodium, Dietary"[Mesh]	S32 TX salt* or sodium*
#10 MeSH descriptor: [Caffeine] explode all trees	#10 Search "Caffeine"[Mesh]	S31 (MH "Drinking Behavior+")
#11 MeSH descriptor: [Drinking Behavior] explode all trees	#11 Search "Drinking Behavior"[Mesh]	S30 (MH "Caffeine")
#12 (salt* or sodium*):ti,ab,kw	#12 Search (salt*[Title/Abstract] OR sodium*[Title/Abstract])	S29 (MH "Sodium, Dietary+")
#13 (Caffein* or decafein* or "de cafein*" or de-cafein* or coffee or tea or alcohol*):ti,ab,kw	#13 Search (Caffein*[Title/Abstract] OR decafein*[Title/Abstract] OR "de cafein*" [Title/Abstract] OR de-cafein*[Title/Abstract] OR coffee[Title/Abstract] OR tea[Title/Abstract] OR alcohol*[Title/Abstract])	S28 (MH "Diet Therapy")
#14 ((diet* or food) and (restrict* or modif* or habit* or free*)):ti,ab,kw	#14 Search ((diet*[Title/Abstract] OR food[Title/Abstract])) AND (restrict*[Title/Abstract] OR modif*[Title/Abstract] OR habit*[Title/Abstract] OR free*[Title/Abstract])	S27 (MH "Food Habits")
#15 nutrition*:ti,ab,kw	#15 Search nutrition*[Title/Abstract]	S26 (MH "Diet, Sodium-Restricted")
#16 Any MeSH descriptor with qualifier(s): [Diet therapy - DH]	#16 Search Diet therapy[sh]	S25 S21 OR S22 OR S23 OR S24
#17 #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16	#17 Search (#6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16)	S24 TX ((aural or labyrinth*) and (hydrops or syndrome or vertigo))
#18 #5 and #17	#18 (#5 and #17)	S23 TX ((endolymphatic or cochlea*) and hydrops)
#19 MeSH descriptor: [Endolymphatic Hydrops] explode all trees and with qualifier(s): [Diet therapy - DH]	#19 Search "Endolymphatic Hydrops/diet therapy"[Mesh]	S22 TX meniere*

(Continued)

#20 #18 or #19

#20 (#18 AND #19)

S14 TX (diet\* or food) and (restrict\* or modif\* or habit\* or free\*)

S13 TX Caffein\* or decaffein\* or "de caffein\*" or de-caffein\* or coffee or tea or alcohol\*

S12 TX salt\* or sodium\*

S11 (MH "Drinking Behavior+")

S10 (MH "Caffeine")

S9 (MH "Sodium, Dietary+")

S8 (MH "Diet Therapy")

S7 (MH "Food Habits")

S6 (MH "Diet, Sodium-Restricted")

S5 S1 OR S2 OR S3 OR S4

S4 TX ((aural or labyrinth\*) and (hydrops or syndrome or vertigo))

S3 TX ((endolymphatic or cochlea\*) and hydrops)

S2 TX meniere\*

S1 (MH "Endolymphatic Hydrops+")

## CONTRIBUTIONS OF AUTHORS

Kiran Hussain: screened and selected studies, obtained full texts, extracted data and assessed risk of bias, entered data into RevMan and would have carried out and interpreted the analysis, drafted the final review and will have responsibility for updating and maintaining the review.

Louisa Murdin: screened and selected studies, extracted data and assessed risk of bias, drafted the final review.

Anne GM Schilder: provided advice as needed throughout the review, drafted the final review.

## DECLARATIONS OF INTEREST

Kiran Hussain: Kiran Hussain is a recipient of a small grant from the Ménière's Society (charity).

Louisa Murdin: none known.

Anne GM Schilder: Professor Anne Schilder is joint Co-ordinating Editor of Cochrane ENT, but had no role in the editorial process for this review. Her evidENT team at UCL is supported in part by the National Institute of Health Research University College London Hospitals Biomedical Research Centre. Their research is funded by the NIHR and EU Horizon2020. She is the national chair of the NIHR Clinical Research Network ENT Specialty. She is the Surgical Specialty Lead for ENT for the Royal College of Surgeons of England's Clinical Trials Initiative. She is co-investigator on the NIHR PGfAR grant 'Defining best Management for Adults with Chronic Rhinosinusitis: the MACRO Programme'. In her role as director of the NIHR UCLH BRC Deafness and Hearing Problems Theme, she acts as an advisor on clinical trial design and delivery to a range of biotech companies.

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### Internal sources

- No sources of support supplied

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## DIFFERENCES BETWEEN PROTOCOL AND REVIEW

In March 2018, changes to the searches of CENTRAL, PubMed and CINAHL were required because the National Library of Medicine (NLM) had removed 'Food Habits' from MeSH. The Information Specialist replaced this term with a broader one 'Feeding Behaviour' in CENTRAL and PubMed and 'Eating Behaviour' in CINAHL. The original term was used in previous searches in July 2016 and July 2017. In March 2018, we re-ran the searches over all years to look for duplicates within the previous two searches and removed them. See [Appendix 1](#) for further details.

## INDEX TERMS

### Medical Subject Headings (MeSH)

\*Caffeine; \*Central Nervous System Stimulants; \*Diet, Sodium-Restricted; \*Sodium Chloride, Dietary; Meniere Disease [\*therapy]; Syndrome

### MeSH check words

Humans