BMJ Open Natural history of tinnitus in adults: a cross-sectional and longitudinal analysis

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ABSTRACT

Objective To assess incidence and changes in tinnitus and bothersome tinnitus as well as associated risk factors in a large sample of UK adults.

Design Prospective cohort study.

Setting UK.

Participants For cross-sectional analysis, a group of 168 348 participants aged between 40 and 69 years with hearing and tinnitus data from the UK Biobank resource. Longitudinal analysis included a subset of 4746 people who attended a 4-year retest assessment.

Main outcome measures Presence and bothersomeness of tinnitus.

Results 17.7% and 5.8% of participants reported tinnitus or bothersome tinnitus, respectively. The 4-year incidence of tinnitus was 8.7%. Multivariate logistic regression models suggested that age, hearing difficulties, work noise exposure, ototoxic medication and neuroticism were all positively associated with both tinnitus and bothersome tinnitus. Reduced odds of tinnitus, but not bothersome tinnitus, was seen in alcohol drinkers versus non-drinkers. Male gender was associated with increased odds of tinnitus, while female gender was associated with increased odds of bothersome tinnitus. At followup, of those originally reporting tinnitus, 18.3% reported no tinnitus. Of those still reporting tinnitus, 9% reported improvement and 9% reported tinnitus becoming more bothersome, with the rest unchanged. Male gender and alcohol consumption were associated with tinnitus being reported less bothersome, and hearing difficulties were associated with the odds of tinnitus being reported as more bothersome.

Conclusions This study is one of the few to provide data on the natural history of tinnitus in a non-clinical population, suggesting that resolution is relatively uncommon, with improvement and worsening of symptoms equally likely. There was limited evidence for any modifiable lifestyle factors being associated with changes in tinnitus symptoms. In view of the largely persistent nature of tinnitus, public health strategies should focus on: (1) primary prevention and (2) managing symptoms in people that have tinnitus and monitoring changes in bothersomeness.

INTRODUCTION

Tinnitus (the subjective experience of sound perception when there is no external source) can be a troublesome experience, and when severe, it can be associated with insomnia, poor concentration, anxiety and/ or depression.¹ Around 10%–15% of adults

Strengths and limitations of this study

- This study investigated both the prevalence and incidence of tinnitus and its correlates in a sample drawn from the UK Biobank resource.
- The study includes both a cross-sectional analysis of a large sample (168348 participants), with the longitudinal component based on a smaller sample (4746 participants).
- The UK Biobank resource, from which the data were drawn, is not completely representative of the UK population.
- A wide range of relevant risk factors were available for the analysis.
- Lack of a consensus on the definition of tinnitus hampers comparison across the literature.

have tinnitus, and although cost-effective, the cost of care of patients with tinnitus is high.² The question of the natural history of tinnitus in adults is of major importance for both patients and clinicians,³ but data regarding the natural history of tinnitus in adults are scant. A few studies have investigated tinnitus in various populations longitudinally (table 1). Estimates of the incidence of tinnitus vary depending on the age of the population and the definition of tinnitus in each study. Such studies are useful in demand forecasting for diagnostic and therapy services but do not support the counselling of existing tinnitus cases regarding their prognosis. The main limitation of many studies examining changes in tinnitus over time is that they were conducted with specific populations, clinical samples or with people taking part in tinnitus research and so may not be representative of the general population.

Clifford *et al*[‡] reported on the progression of tinnitus in a US Marine cohort, indicating that worsening tinnitus was associated with the presence of post-traumatic stress disorder and moderate/severe traumatic brain injury. One other study reported a modest improvement in the bothersomeness of tinnitus at follow-up 4.9 years after treatment by a clinical psychologist, the majority (59%) having received cognitive–behavioural therapy for

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Table 1 Prevalence and/or incidence rates of tinnitus

Study	Definition of tinnitus	Population	Prevalence of tinnitus at baseline	Follow-up interval; incident tinnitus
Gopinath et al ¹³	Tinnitus over the past year lasting 5 min or longer.	Australia Age: ≥55 years. Baseline (n=2006). Follow-up (n=1292, female=58.9%).	37% (481).*	5-year follow-up; Persisting: 82% (346). Resolved: 18% (78). Incidence: 18% (156).
Nondahl <i>et al</i> ¹²	Tinnitus over the past year of (1) at least moderate severity or causing difficulty with sleep and (2) mild tinnitus	USA. Age: 48–92 years. Baseline (n=3753,	8.2% (moderate tinnitus) (308).	5-year follow-up; incidence of significant tinnitus: 5.7% (143).
	not affecting sleep.	female=57.7%). Follow-up (n=2800, female=58.6%).	20.2% (mild tinnitus) (754).	5-year follow-up; persisting: 68.6% (394). Resolved: 31.4% (180).
Nondahl <i>et al⁶⁴</i>	Tinnitus over the past year of at least moderate severity or causing difficulty with sleep.	USA. Age: 48–92 years. Follow-up (n=2922, female: 59.3%).	N.A.	10-year follow-up; 12.7% (371).
Lee et al ¹⁴	Based on record of health service utilisation for 'tinnitus' through the Korean National Health Insurance Service.	South Korea. Age: all ages. Follow-up (n=51 million, female=not reported).	12.58–14.62 per 1000 from 2006 to 2015.	Yearly incidence; 8.26–9.49 per 1000 from 2006 to 2015 9.1% 10-year incidence.
Martinez <i>et al</i> ¹⁵	Any tinnitus: based on health service utilisation for 'tinnitus' gathered through the UK Clinical Practice Research Datalink. Significant tinnitus: as above but with related follow-up within 28 days. See Martinez <i>et al</i> ¹⁵ for full definition.	UK. Age: ≤85 years. Follow-up (n=4.7 million, female=50.5%).	N.A.	10-year follow-up; incidence significant tinnitus 5.4 per 10 000 person years. Incidence any tinnitus 47.3 per 10 000 person years.

tinnitus.³ Another study reported that severity of symptoms tended to be more severe, with tinnitus of longer duration among patients presenting for tinnitus therapy.⁵ A systematic review and meta-analysis reporting the experiences of patients with tinnitus who were research participants enrolled in control (waiting list) arms of clinical trials reported a small, statistically significant improvement in tinnitus symptoms over time, though clinical significance of these improvements was unclear.⁶ Placebo groups in controlled clinical trials of tinnitus treatments have also reported reduced bothersomeness of tinnitus immediately and up to 14-week postplacebo treatment.⁷⁻⁹ In a conference report, Smith and Coles¹⁰ reported data regarding tinnitus from the UK National Study of Hearing.¹¹ Participants reporting tinnitus experiences were asked to retrospectively rate their tinnitus loudness and annovance at two stages of their tinnitus experience: 'onset to middle' and 'middle/recent or end'. These illdefined sample points render the data in this report are hard to interpret, but it appears that in a small number (8.5%), the tinnitus had resolved completely and that there was a general trend over time towards decreased annoyance.

Large-scale data regarding longitudinal experiences of a general population regarding tinnitus has not been reported to date, with the closest examples examining samples an order of magnitude smaller than the present study.^{12 13} The only large-scale population data available used indirect measures of tinnitus based on clinical record or health claim data.^{14 15} In the present study, the aim was to establish the proportions of people who experience tinnitus and bothersome tinnitus and changes in tinnitus and bothersome tinnitus over time and to determine demographic, health and lifestyle correlates in order to inform patient counselling and identify potential avenues for prevention and treatment of tinnitus.

METHODS

Participants

Participants were drawn from the UK Biobank, an international resource for studying the genetic, environmental and lifestyle causes of diseases of middle and older age.¹⁶ Participant recruitment was conducted via the UK National Health Service and aimed to be as inclusive as possible of the UK population. In total, 9.2 million invitations were sent to recruit 503 325 participants who were aged between 40 and 69 years between 2006 and 2010, a response rate of 5.47%. The UK Biobank sample contains a higher proportion of women, people reporting white British ethnic background and people living in less deprived areas than the general population.¹⁷ The UK Biobank is not representative of the UK general population, but the disease–exposure relationships are thought to generalisable due to the size and inclusiveness of the sample. Hearing and tinnitus measures were included part way through data collection, so information on tinnitus at baseline was available for 168348 participants.

Participants attended an assessment centre where data on demographic, health, environmental and lifestyle factors were collected via computerised questionnaire along with physical measures including hearing testing during assessments of around 90min in duration. Further information on procedures and the data collected is contained on the UK Biobank website (http://www.ukbiobank.ac.uk/). During 2012 and 2013, 17 819 participants attended a retest assessment, with a 21% response rate. All baseline measures were repeated, including hearing and tinnitus. The mean retest interval was 4.3 years (range 2–7 years); retest tinnitus data were available for 4746 participants (for further details of the repeat assessment, see http://biobank.ctsu.ox.ac.uk/~ bbdatan/Repeat_assessment_doc_v1.0.pdf). All participants provided written informed consent.

Information on sex and ethnicity (based on 2001 UK Census categories) and area of residence was collected. Area of residence was used to determine a Townsend deprivation score. The Townsend deprivation score is a proxy for socioeconomic status and is applicable across the countries of the UK.¹⁸ Townsend scores are based on four variables: unemployment, non-car ownership, nonhome ownership and household overcrowding. Each variable is normalised relative to national levels and summed to provide an overall deprivation index. Higher scores represent more deprived (less affluent) socioeconomic status. A score of 0 represents the national mean with an SD of 1. Townsend scores for areas of residence ranged between 14.01 and -5.59 in the 2011 census.¹⁹ In the regression analyses below, Townsend scores were grouped from least to most deprived quartiles in the study sample.

Tinnitus

Participants were asked 'Do you get or have you had noises (such as ringing or buzzing) in your head, or in one or both ears, that lasts for more than five min at a time?'. In this analysis, tinnitus was identified based on responses of 'yes most of the time', 'yes a lot of the time' or 'yes some of the time', similar to criteria used in other studies of the epidemiology of tinnitus.^{20–22} If a participant reported that they did experience tinnitus that lasted for more than 5 min at a time, they were asked 'How much do these noises worry, annoy or upset you when they are at their worst?'; severely, moderately, slightly or not at all. In this analysis, 'bothersome' tinnitus was identified on the basis of responses of either 'moderately' or 'severely'.

Incident tinnitus was identified if a person who did not report tinnitus at baseline reported tinnitus at least some of the time at retest. Among those who reported tinnitus at baseline, 'Worse tinnitus' was identified if someone reported their tinnitus as not being bothersome at baseline (ie, 'slightly' or 'not at all') but reported their tinnitus being bothersome at follow-up (ie, 'moderately' or 'severely').

Hearing

Participants completed an English version of the Digit Triplet Test (DTT), a test of speech recognition in noise developed for large-scale hearing screening.^{23 24} The DTT correlates strongly with audiometric thresholds. The DDT is described elsewhere (http://biobank.ctsu. ox.ac.uk/crystal/label.cgi?id=100049). In short, 15 sets of three monosyllabic digits (eg, 6-1-3) were presented over circumaural headphones with the volume of presentation set to a comfortable level. Digits were presented in background noise shaped to match the spectrum of the spoken digits. Noise levels varied adaptively to track an Signal-to-noise ratio (SNR) for the 50% speech recognition threshold (SRT), which was based on the mean SNR for the last eight triplets. Lower (more negative scores) indicates better performance. Hearing was additionally indexed by self-reported hearing status with the question 'Do you have any difficulty with your hearing?'.

Occupation-related and music-related noise exposure, ototoxic medication, metabolic syndrome, physical activity, smoking, alcohol consumption and hearing aid use

The potential associations between tinnitus and risk factors were explored using a previously identified list and discussion between the authors.¹ Occupational and music-related noise exposure was identified on the basis of any reported exposure in response to the questions 'Have you ever worked in a noisy place where you had to shout to be heard?' and 'Have you ever listened to music for more than 3 hours per week at a volume which you would need to shout to be heard or, if wearing headphones, someone else would need to shout for you to hear them?'. The criterion for work-related and music-related noise corresponds to exposure estimated to exceed 85 dB(A).²⁵ Use of ototoxic medication was identified on the basis of reported regular (daily, weekly or monthly) use of medications known to have ototoxic properties (including loop diuretics, aminoglycoside antibiotics, quinine derivatives, non-steroidal antiinflammatories and salicylates). Metabolic syndrome was identified based on the Adult Treatment Panel III report of the National Cholesterol Education Program criteria²⁶; positive risk was identified on the basis of three or more of waist circumference of $\geq 102 \text{ cm}$ in men and $\geq 88 \text{ cm}$ in women; participant report of high cholesterol or if the participant reported they were currently taking medication for high cholesterol; measured systolic blood pressure greater than 130 mm Hg or diastolic pressure greater than 85 mm Hg; and participant report of diabetes or the use of medication for diabetes. Participants were identified as being physically 'active' if they reported doing more than 10 min of physical activity in relation to the question 'Yesterday, about how long did you spend doing activities that needed moderate effort, making you

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somewhat short of breath? For example walking upstairs, going to the gym, jogging, energetic dancing, aerobics, most sports, using heavy power tools and other physically demanding DIY & gardening' 'Inactive' participants were identified on the basis of physical activity of 10 min or less. Current or previous tobacco smoking was identified on the basis of positive responses to two questions: 'Do you smoke tobacco now?' and 'In the past, how often have you smoked tobacco?'. Alcohol drinkers were identified on the basis of any report of current alcohol consumption ('About how often do you drink alcohol?', 'Special occasions only', 'One to three times a month', 'One or twice a week', 'Three or four times a week' or 'Daily or almost daily'). Non-drinkers were categorised based on a response of 'Never'. Hearing aid use was identified on the basis of a 'yes' response to 'Do you use a hearing aid most of the time?'.

Neuroticism

Neuroticism scores were based on summed positive responses to 12 items from the Eysenck Personality Questionnaire Revised,²⁷ including 'Does your mood often go up and down?'; 'Do you ever feel "just miserable" for no reason?'; 'Are you an irritable person?'; 'Are your feelings easily hurt?'; 'Do you often feel "fed-up"?'; 'Would you call yourself a nervous person?'; 'Are you a worrier?'; 'Would you call yourself tense or "highly strung"?'; 'Do you worry too long after an embarrassing experience?'; 'Do you suffer from "nerves'?'; 'Do you often feel lonely?'; and 'Are you often troubled by feelings of guilt?'. Scores are summed to provide an integer score between 1 and 12 representing the number of neurotic traits present, with higher scores indicating greater neuroticism.

Data analysis

Cross-tabulations performed to describe characteristics of those who reported tinnitus versus no tinnitus and the subset of people with tinnitus who reported 'bothersome' tinnitus. Demographic, health, lifestyle and psychological characteristics were selected on the basis of previously being linked to tinnitus.^{21 28 29} There were missing data for some measures primarily due to measures being added to the study protocol at different time points during data collection (see table 2). Because the reason for the missing data was not systematically related to the outcomes of interest in this study, it was assumed that the data were missing completely at random. Missing variable analysis did not identify any pattern to the missing data. Multinomial logistic regression was used to model crosssectional baseline associations between demographic, hearing, noise exposure, health and lifestyle factors and tinnitus (vs no tinnitus) and bothersome tinnitus (vs nonbothersome tinnitus). A Cox proportional hazard model was used to model the incidence of tinnitus and more bothersome tinnitus at 4-year follow-up. All the statistical assumptions for performing the Cox proportional hazard model were met. Analyses were performed with IBM SPSS V.23.³⁰

Table 2 Sample characteristics				
	No tinnitus	Tinnitus		
		Any tinnitus (not bothersome† bothersome)	Not bothersome	Bothersome
	n=1 38 487	n=29861	n=20110	n=9751
Age (168 348)*	56.4 (SD 8.2)	58.7 (SD 7.58)	57.5 (SD 8.22)	58.0 (SD 7.78)
Sex (male; 168 348) (%)	44.0	52.80	51.0	45.70
Social economic status score† (168 079)	-1.12 (SD 2.92)	-0.99 (SD 3.01)	-1.02 (SD 3.00)	-0.66 (SD 3.16)
SRT better ear (157 574)	-7.43 (SD 1.62)	-7.07 (SD 1.96)	-7.26 (SD 1.80)	-6.91 (SD 2.13)
Hearing difficulties (168 348) (%)	21.3	56.4	43.4	63.3
Work noise exposure (166 805) (%)	20.5	34.4	32.2	37.0
Music noise exposure (165 977) (%)	11.5	16.6	17.4	18.5
Physical activity (106 989) (%)	71.1	71.7	71.5	69.3
Ototoxic medication (168 348) (%)	39.2	46.5	44.6	53.1
Alcohol drinker (168 201) (%)	91.5	90.1	91.2	88.1
Current or previous smoking (167 725) (%)	43.9	48.5	47.4	50.7
Metabolic risk (168 348) (%)	9.1	12.0	10.7	13.3
Neuroticism score (136 600)	3.98 (SD 3.22)	4.63 (SD 3.41)	4.44 (SD 3.33)	5.64 (SD 3.47)

*The number in brackets indicates the number of participants that completed each measure.

+Social economic status indexed by Townsend deprivation index score; lower (more negative) score indicates less deprived (more affluent) status.

SRT, speech recognition threshold.

Patient and public involvement

Patient and public involvement is reported according to the Guidance for Reporting Involvement of Patients and the Public – short form.³¹ (1) Aim: UK Biobank consulted with stakeholders including the public at several times during the planning stages and postimplementation to obtain guidance and feedback in relation to: consent, confidentiality, access, commercialisation and oversight/ monitoring. The conception of the project and its aim sprung directly from public enquiries fielded by the British Tinnitus Association, a partner in this project. (2) Methods: for the UK Biobank, a key element in the public consultation process was an initial workshop that included 20 members of the public in the study target age range and 10 outside the target age range. Sessions key points were noted, and sessions tape were recorded; a postworkshop questionnaire was sent to all attendees and to any stakeholders who were not able to attend the workshops in person (to increase representativeness). (3) Results: public opinion across many areas addressed in the aims was diverse; a full report can be found at http://www. ukbiobank.ac.uk/public-consultation/; (4) Discussion: public input influenced ethical considerations, access to data, the consent process, the commercialisation of the resource and oversight/monitoring. (5) Reflections/ critical perspective: public and other stakeholder input into the study was essential to ensure public confidence in the study conduct and to respond to public concerns

with the resource. While efforts were taken to consider public input, the diversity of opinion meant that not all perspectives were equally influential on the UK Biobank's design and conduct.

RESULTS

Cross-sectional analysis

In this sample of adults aged 40–69 years, 17.7% (n=29861) reported tinnitus and 5.8% (n=9751) reported bothersome tinnitus. Table 2 summarises the characteristics of participants who reported that they experienced tinnitus versus those who did not report tinnitus. Characteristics of those who reported tinnitus are broken down further with respect to whether participants reported their tinnitus as being bothersome or not bothersome.

All variables were entered simultaneously into multivariable logistic regression model for tinnitus versus no tinnitus (of the original sample of 168 348, after excluding all participant with missing data 80 380 participants were included in the analysis). A multivariable logistic regression model was also conducted to compare bothersome tinnitus versus not bothersome tinnitus (of the original sample of 29 861 tinnitus sufferers after excluding all participants with missing data, 21 690 were included in the analysis) (table 3). Similar patterns of association were observed for tinnitus and bothersome tinnitus. The Nagelkerke r^2 was 0.143 for tinnitus and

Table 3 Cross-sectional correlates of tinnitus and bothersome tinnitus								
	Tinnitus n=80380 χ²(15)=7110.23, p<0.001			Bothersome tinnitus n=21 690 χ²(15)=912.89, p<0.001				
		95% CI for Exp(B)			95% CI for Exp(B)			
	Exp(B)	Lower	Upper	P value	Exp(B)	Lower	Upper	P value
Age	1.03	1.03	1.03	0.000	1.00	1.00	1.01	0.176
Sex (male)	1.20	1.15	1.25	0.000	0.74	0.68	0.80	0.000
Social economic status				0.294				0.023
First quartile 1 (reference)				-				-
Second quartile	1.02	0.97	1.08	0.484	0.95	0.86	1.05	0.297
Third quartile	0.98	0.93	1.04	0.480	1.03	0.93	1.14	0.587
Fourth quartile (most deprived)	1.03	0.98	1.09	0.262	1.11	1.00	1.23	0.043
SRT better ear	1.03	1.02	1.04	0.000	1.02	1.00	1.05	0.020
Hearing difficulties (yes)	3.75	3.60	3.90	0.000	2.07	1.92	2.23	0.000
Work noise exposure (yes)	1.40	1.34	1.47	0.000	1.15	1.06	1.25	0.001
Music noise exposure (yes)	1.39	1.31	1.47	0.000	1.03	0.93	1.13	0.576
Physical activity (yes)	1.05	1.00	1.09	0.038	0.97	0.89	1.04	0.372
Ototoxic medication (yes)	1.18	1.13	1.23	0.000	1.19	1.11	1.28	0.000
Alcohol drinker (yes)	0.87	0.80	0.93	0.000	0.89	0.78	1.01	0.070
Current or previous smoking (yes)	0.99	0.95	1.03	0.500	1.04	0.97	1.12	0.300
Metabolic risk (yes)	1.05	0.98	1.12	0.210	0.97	0.86	1.09	0.601
Neuroticism score	1.05	1.04	1.06	0.000	1.10	1.08	1.11	0.000

SRT, speech recognition threshold.

0.067 for bothersome tinnitus. Older age, male sex, poorer SRT, hearing difficulties, work noise exposure, music noise exposure, physical activity, regular use of ototoxic medication and neuroticism were associated with tinnitus. Alcohol consumption was associated with lower odds of tinnitus. Female sex, most deprived social economic status, poorer SRT, hearing difficulties, work noise exposure, ototoxic medication and neuroticism were associated with bothersome tinnitus. The sample included 1013 hearing aid users. The analyses were rerun to check for interactions with hearing difficulties and hearing aid use on tinnitus and bothersome tinnitus. There was no significant hearing aid by hearing difficulties interaction for tinnitus (OR 0.50, 95% CI 0.21 to 1.21, p=0.125) or bothersome tinnitus (OR 0.87, 95% CI 0.18 to 4.18, p=0.888).

Longitudinal analysis

A subset of participants (3997 people) who had completed the questions about tinnitus and the full set of correlates of interest were included in the longitudinal analysis, with a mean retest interval of 4.3 years (2–7 years range). There were 276 cases of incident tinnitus among the 3177 people who did not report tinnitus at baseline; a 4-year incidence of 8.7%. The Cox proportional hazard model for incident tinnitus between baseline and 4-year follow-up was not statistically significant ($\chi^2(15)=21.6$, p=0.119). Among the 820 people who reported tinnitus at baseline and completed responses at follow-up, 150 reported no tinnitus at follow-up (including 63 who claimed never to have had tinnitus); 18.3% of people reporting tinnitus at baseline did not report tinnitus at follow-up. Of the 820 tinnitus sufferers completing follow-up, after excluding cases of missing data, 565 were included in the Cox proportional hazard model. The analysis suggests that only hearing difficulties were associated with reduced likelihood of no tinnitus at follow-up (table 4).

Among the 1039 people who completed questions about tinnitus annoyance at baseline and follow-up, 850 (81.8%) reported no change, 93 (9%) reported that their tinnitus was more bothersome and 93 (9%) reported their tinnitus being less bothersome.

In a Cox proportional hazard model, of those who completed follow-up and who reported 'not at all' or 'slightly' bothersome tinnitus at baseline cases of tinnitus being reported as being more bothersome (vs those reporting no change) were associated with higher (poorer) better ear SRT, non-drinking and female gender (table 5). The sample included 27 hearing aid users, and the model was rerun to check for an interaction with hearing aid use and speech reception threshold. The interaction was not statistically significant (OR 0.88, 95% CI 0.67 to 1.14, p=0.337). The model for reduced bothersomeness was not statistically significant χ^2 (15)=24.1, p=0.063.

Table 4 Cox proportional hazard model for no tinnitus at follow-up (ie, resolved tinnitus)						
	Resolved tinnitus n=565 (χ²(15)=26.7, p=0.031)					
		95% CI				
	β	Lower	Upper	P value		
Age	0.99	0.96	1.03	0.683		
Sex (male)	0.96	0.62	1.47	0.841		
Social economic status				0.558		
First quartile 1 (reference)				-		
Second quartile	1.11	0.69	1.80	0.671		
Third quartile	0.72	0.39	1.30	0.273		
Fourth quartile (most deprived)	1.01	0.55	1.85	0.972		
SRT better ear	0.98	0.88	1.10	0.735		
Hearing difficulties (yes)	0.48	0.32	0.74	0.001		
Work noise exposure (yes)	1.47	0.95	2.26	0.080		
Music noise exposure (yes)	1.26	0.76	2.09	0.379		
Physical activity (yes)	1.48	0.84	2.58	0.173		
Ototoxic medication (yes)	0.83	0.55	1.25	0.383		
Alcohol drinker (yes)	0.66	0.30	1.47	0.309		
Current or previous smoking (yes)	1.19	0.79	1.78	0.399		
Metabolic risk (yes)	0.66	0.35	1.25	0.203		
Neuroticism score	1.05	0.99	1.12	0.101		

SRT, speech recognition threshold.

Table 5 Cox proportional hazard model for tinnitus bothersomeness worse							
	Bothersomeness worse n=404 χ^2 (15)=34.4, p=0.003						
		95% CI					
	β	Lower	Upper	P value			
Age	0.98	0.93	1.04	0.481			
Sex (male)	0.44	0.22	0.86	0.017			
Social economic status				0.258			
First quartile 1 (reference)				-			
Second quartile	1.66	0.80	3.48	0.176			
Third quartile	0.77	0.27	2.23	0.633			
Fourth quartile (most deprived)	0.77	0.27	2.19	0.621			
SRT better ear	1.13	1.02	1.27	0.026			
Hearing difficulties (yes)	2.01	0.96	4.20	0.063			
Work noise exposure (yes)	1.41	0.71	2.83	0.329			
Music noise exposure (yes)	1.15	0.50	2.63	0.738			
Physical activity (yes)	0.88	0.39	1.96	0.752			
Ototoxic medication (yes)	1.24	0.65	2.34	0.513			
Alcohol drinker (yes)	0.30	0.11	0.87	0.026			
Current or previous smoking (yes)	1.61	0.83	3.11	0.156			
Metabolic risk (yes)	0.67	0.22	2.04	0.485			
Neuroticism score	0.95	0.86	1.06	0.381			

SRT, speech recognition threshold.

DISCUSSION

In cross-sectional analysis, 17.7% of adults aged 40-69 years old reported tinnitus, with 5.8% reporting that tinnitus was bothersome. The 4-year incidence of tinnitus in this sample was 8.7%. The study offered some cause for optimism with respect to the natural history of tinnitus; around 18% of people who reported tinnitus at baseline did not report tinnitus at follow-up, an average of 4 years later. For those who continued to experience tinnitus, 81.8% reported that tinnitus bothersomeness was unchanged after 4 years, 9% reported tinnitus became worse (previously not bothersome, now bothersome) and in 9% better (previously bothersome, now not). The strengths of the study include the large inclusive sample, which was not derived from a specific tinnitus nor hearing study. The availability of longitudinal data was a significant strength. Longitudinal tinnitus data are available in a very small number of other studies. The use of standard tinnitus phenotype questions allowed comparison of these results with those of other studies. Although accounted for in the model, the variability in time elapsed at retest (2-7 years) may be a limitation. However, the minimum of 2 years is longer than the period of most intervention studies and provides time to observe natural variation in tinnitus. In terms of patient counselling about long-term prognoses for tinnitus, the 4-year mean follow-up period limits the certainty of any opinion in relation to longer-term outcomes. One significant limitation of the study is the possibility that a person

may have received clinical help for tinnitus during the intervening years. Most people seek help within the first year of onset,³² so this is unlikely to have been the case for a large proportion of participants here. Unfortunately, information about receiving clinical help and the duration of tinnitus was not available in this study, and this limits our confidence in stating that all cases of spontaneous recovery were indeed spontaneous.

Tinnitus correlates

Poorer hearing (better ear SRT and self-reported hearing difficulties) was associated with the presence of tinnitus and bothersome tinnitus. Hearing difficulties were associated with lower likelihood of resolved tinnitus and SRT with lower likelihood of tinnitus being less bothersome over time. However, there was no relationship between SRT or hearing difficulties and incident tinnitus. A relationship between tinnitus and hearing loss is consistently reported^{21 33-36} with hearing loss being proposed as a trigger for tinnitus, which then persists due to maladaptive plasticity in the central auditory and associated systems.³⁷ The lack of a relationship between hearing and incident tinnitus may be due to the much smaller sample for the longitudinal analysis versus the cross-sectional analysis. Work noise exposure was associated with prevalent tinnitus and bothersome tinnitus. Music exposure was associated with prevalent tinnitus. Noise exposure is the primary modifiable risk factor for tinnitus^{34 38 39} and the pathophysiological impact can be either cochlear hair cell dysfunction, leading to a subjective hearing loss, and/or cochlear synaptopathy, the effects of which are more subtle. 40

Alcohol consumption was associated with reduced odds of tinnitus, but not of bothersome tinnitus. Moderate alcohol consumption has been suggested to have a protective effect on hearing, perhaps via cardiovascular pathways.^{41 42} Other studies reported no association between alcohol consumption and tinnitus,^{43 44} with one study reporting increased risk of tinnitus with alcohol consumption.⁴² There are several difficulties with disentangling the effect of alcohol consumption on tinnitus. First, alcohol consumption is highly confounded with sociocultural factors that may also impact on health, including hearing.⁴⁵ Second, impacts of alcohol consumption may be dose dependent; heavy alcohol consumption is certainly bad for general health, including hearing.⁴⁶ Impacts may be different for moderate or light levels of consumption.⁴⁶⁻⁴⁸ Third, comparing health outcomes in drinkers versus non-drinkers may give the false impression that alcohol consumptions is linked to better health outcomes due to the inclusion of people who have given up drinking due to poor health in the non-drinker group ('sick-quitters').^{41 49} The detailed level of analysis in relation to these questions is beyond the scope of the present paper and should be the subject of future investigation.

Interestingly, men were more likely to report tinnitus, but women more likely to report tinnitus being bothersome. One explanation may be that because men are more likely to have hearing loss,⁵⁰ they are more likely to experience tinnitus. However, men may be less likely to report tinnitus as being bothersome due to differences in socialisation that leads to men being less likely to acknowledge and report discomfort in relation to physical symptoms in general.⁵¹ Higher neuroticism scores were also associated with increased odds of both tinnitus and bothersome tinnitus, in line with previous research.⁵² The lack of association between neuroticism and increased bothersomeness of tinnitus over time suggests that neuroticism and psychological discomfort may be a consequence of rather than a risk for bothersome tinnitus.

Use of ototoxic medication was associated with prevalent tinnitus and bothersome tinnitus, but not with incidence of tinnitus or changes in bothersomeness. The association between tinnitus and ototoxic medication is supported by other literature⁵³ as is the adverse effect on quality of life that results.⁵⁴ Though an association between smoking and tinnitus has been reported previously,^{28 55} there were no associations with tinnitus in the present analysis. Both metabolic syndrome and lack of physical activity have previously been associated with tinnitus⁵⁶ and tinnitus bothersomeness,^{57 58} while physical activity was weakly associated with tinnitus in the current study. The relatively low baseline for being physically active (10 min daily moderate activity), and self report measure in the current study may help to explain this apparent anomaly. Previous studies using accelerometers have indicated that applying higher thresholds for

physical activity produced more pronounced associations in older adult populations. 56

Taken together, there is an indication that generally healthy lifestyle may be linked to reduced likelihood of tinnitus. Variations in findings relating to both factors across studies may relate to differences in measurement and the fact that both are also strongly associated with age, socioeconomic status and sex. A limitation of the study is that the sample sizes were substantially lower for the longitudinal analyses; lack of longitudinal associations may be due to lack of statistical power. The analysis did not include potentially important explanatory factors (eg, personality factors besides neuroticism, leisure noise and genetic factors) and some factors may not have been well captured by the measures available in this data set. For example, work-related and music-related noise exposure was based on a self-report measure, which is estimated to correspond to noise levels above 85 dB(A).²⁵ However, the measure does not account for levels that may substantially exceed 85 dB(A) nor for the use or nonuse of ear protection.

A key limitation of the present study—and all other tinnitus research—is the lack of a reliable objective measure of tinnitus and no agreement about the validity or characterisation of tinnitus phenotypes.⁵⁹ This point was highlighted in the present study: of those originally reporting tinnitus with subsequent cessation at follow-up, over one-third now claimed never to have had tinnitus. This finding calls into question the reliability of the current self-report measures of tinnitus utilised in epidemiological research and suggests that a collaborative effort to arrive at a refined definition and appropriate measure of tinnitus should be made.

There were no interactions between prevalent tinnitus, tinnitus bothersomeness or change in tinnitus bothersomeness with hearing aid use. These data suggest that poor hearing is the main driver of the risk of tinnitus, but that this is not offset by hearing aid use. Clinical experience, case series⁶⁰ and retrospective studies⁶¹ indicate that hearing aids can reduce or inhibit tinnitus, although to date no controlled trials have shown the benefits of hearing aids on tinnitus.⁶² The UK Biobank data did not include information on other tinnitus therapies, used either individually, or in combination with hearing aids, some of which have shown promising results.⁶³ Given the modest and uncertain impact of hearing aids, public health approaches should focus on primary prevention of hearing loss in order to reduce the impacts of tinnitus. Additionally, given the largely persistent nature of tinnitus shown in the study, further attention should be paid to effectively managing symptoms in people with tinnitus and ultimately to finding a cure.

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REFERENCES

- Baguley D, McFerran D, Hall D. Tinnitus. *The Lancet* 2013;382:1600–7.
- 2 Stockdale D, McFerran D, Brazier P, et al. An economic evaluation of the healthcare cost of tinnitus management in the UK. BMC Health Serv Res 2017;17:577.
- 3 Andersson G, Vretblad P, Larsen HC, et al. Longitudinal followup of tinnitus complaints. Arch Otolaryngol Head Neck Surg 2001;127:175–9.
- 4 Clifford RE, Baker D, Risbrough VB, *et al.* Impact of TBI, PTSD, and hearing loss on tinnitus progression in a US marine cohort. *Mil Med* 2019;184:839–46.
- 5 Stouffer JL, Tyler RS, Kileny PR, et al. Tinnitus as a function of duration and etiology: counselling implications. Am J Otol 1991;12:188–94.
- 6 Phillips JS, McFerran DJ, Hall DA, et al. The natural history of subjective tinnitus in adults: a systematic review and meta-analysis of no-intervention periods in controlled trials. Laryngoscope 2018;128:217–27.
- 7 Duckert LG, Rees TS. Placebo effect in tinnitus management. *Otolaryngol Head Neck Surg* 1984;92:697–9.
- 8 Drew S, Davies E. Effectiveness of Ginkgo biloba in treating tinnitus: double blind, placebo controlled trial. *BMJ* 2001;322:73.
- 9 Robinson SK, Viirre ES, Bailey KA, *et al.* Randomized placebocontrolled trial of a selective serotonin reuptake inhibitor in the treatment of nondepressed tinnitus subjects. *Psychosom Med* 2005;67:981–8.
- 10 Smith P, Coles R. Epidemiology of tinnitus: an update. In: Feldmann H, ed. Proceedings of the third International tinnitus seminar. Karlsruhe: Harsch, 1987: 147–53.
- 11 Davis A. Hearing in adults: the prevalence and distribution of hearing impairment and reported hearing disability in the MRC Institute of hearing research's national study of hearing. London: Whurr Publishers, 1995.
- 12 Nondahl DM, Cruickshanks KJ, Wiley TL, et al. Prevalence and 5-year incidence of tinnitus among older adults: the epidemiology of hearing loss study. J Am Acad Audiol 2002;13:323–31.
- 13 Gopinath B, McMahon CM, Rochtchina E, *et al.* Incidence, persistence, and progression of tinnitus symptoms in older adults: the blue Mountains hearing study. *Ear Hear* 2010;31:407–12.

- 14 Lee HM, Han Kdo, Kong SK, et al. Epidemiology of clinically significant tinnitus: a 10-year trend from nationwide health claims data in South Korea. Otol Neurotol 2018;39:680–7.
- 15 Martinez C, Wallenhorst C, McFerran D, et al. Incidence rates of clinically significant tinnitus: 10-year trend from a cohort study in England. *Ear Hear* 2015;36:e69.
- 16 Collins R. What makes UK Biobank special? *Lancet* 2012;379:1173–4.
- 17 Fry A, Littlejohns TJ, Sudlow C, et al. Comparison of sociodemographic and health-related characteristics of UK Biobank participants with those of the general population. Am J Epidemiol 2017;186:1026–34.
- 18 Norman P. Identifying change over time in small area socio-economic deprivation. *Applied Spatial Analysis and Policy* 2010;3:107–38.
- 19 Yousaf S, Bonsall A. Uk Townsend deprivation scores-townsend deprivation scores report. 2017. Colchester: UK Data Service, 2011.
- 20 Davis A, Rafaie EA. Epidemiology of tinnitus. In: Tyler RS, ed. *Tinnitus Handbook*. Sandiego, CA: Singular, 2000: 1–23.
- 21 Gopinath B, McMahon CM, Rochtchina E, et al. Risk factors and impacts of incident tinnitus in older adults. Ann Epidemiol 2010;20:129–35.
- 22 Davis AC. The prevalence of hearing impairment and reported hearing disability among adults in Great Britain. *Int J Epidemiol* 1989;18:911–7.
- 23 Smits C, Kapteyn TS, Houtgast T. Development and validation of an automatic speech-in-noise screening test by telephone. *Int J Audiol* 2004;43:15–28.
- 24 Hall J. The development of a new English sentence in noise test and an English number recognition test. University of Southampton: Southampton, 2006.
- 25 Health and Safety Executive. *Noise at work. guidance on regulation.* London: HMSO, 1989.
- 26 National Cholesterol Education Program. Expert panel on detection, evaluation, and treatment of high blood cholesterol in adults (adult treatment panel III). third report of the National cholesterol education program (NCEP) expert panel on detection, evaluation, and treatment of high blood cholesterol in adults (adult treatment panel III) final report. *Circulation* 2002;106:3143–421.
- 27 Eysenck SBG, Eysenck HJ, Barrett P. A revised version of the psychoticism scale. *Pers Individ Dif* 1985;6:21–9.
- 28 Shargorodsky J, Curhan GC, Farwell WR. Prevalence and characteristics of tinnitus among US adults. *Am J Med* 2010;123:711–8.
- 29 McCormack A, Edmondson-Jones M, Fortnum H, et al. The prevalence of tinnitus and the relationship with neuroticism in a middle-aged UK population. J Psychosom Res 2014;76:56–60.
- 30 IBM Corp. IBM SPSS statistics for windows, version 23.0. Armonk, NY: IBM Corp, 2015.
- 31 Staniszewska S, Brett J, Simera I, *et al.* GRIPP2 reporting checklists: tools to improve reporting of patient and public involvement in research. *Res Involv Engagem* 2017;3:13.
- 32 McFerran D, Hoare DJ, Carr S, et al. Tinnitus services in the United Kingdom: a survey of patient experiences. BMC Health Serv Res 2018;18:110.
- 33 Kim H-J, Lee H-J, An S-Y, *et al*. Analysis of the prevalence and associated risk factors of tinnitus in adults. *PLoS One* 2015;10:e0127578.
- 34 Nondahl DM, Cruickshanks KJ, Huang G-H, et al. Tinnitus and its risk factors in the beaver dam offspring study. Int J Audiol 2011;50:313–20.
- 35 Gallus S, Lugo A, Garavello W, et al. Prevalence and determinants of tinnitus in the Italian adult population. *Neuroepidemiology* 2015;45:12–19.
- 36 Michikawa T, Nishiwaki Y, Kikuchi Y, et al. Prevalence and factors associated with tinnitus: a community-based study of Japanese elders. J Epidemiol 2010;20:271–6.
- 37 Shore SE, Roberts LE, Langguth B. Maladaptive plasticity in tinnitus-triggers, mechanisms and treatment. *Nat Rev Neurol* 2016;12:150–60.
- 38 Bhatt JM, Lin HW, Bhattacharyya N, Prevalence BN. Prevalence, severity, exposures, and treatment patterns of tinnitus in the United States. JAMA Otolaryngol Head Neck Surg 2016;142:959–65.
- 39 Axelsson A, Prasher D. Tinnitus induced by occupational and leisure noise. *Noise Health* 2000;2:47.
- 40 Liberman MC. Noise-induced and age-related hearing loss: new perspectives and potential therapies. *F1000Res* 2017;6:927.
- 41 Dawes P, Cruickshanks KJ, Moore DR, et al. Cigarette smoking, passive smoking, alcohol consumption, and hearing loss. J Assoc Res Otolaryngol 2014;15:663–74.
- 42 Quaranta A, Assennato G, Sallustio V. Epidemiology of hearing problems among adults in Italy. *Scand Audiol Suppl* 1996;42:9–13.

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- 43 Sindhusake D, Golding M, Wigney D, et al. Factors predicting severity of tinnitus: a population-based assessment. J Am Acad Audiol 2004;15:269–80.
- 44 Sindhusake D, Golding M, Newall P, et al. Risk factors for tinnitus in a population of older adults: the blue Mountains hearing study. Ear Hear 2003;24:501–7.
- 45 Rimm EB, Giovannucci EL, Willett WC, *et al*. Prospective study of alcohol consumption and risk of coronary disease in men. *The Lancet* 1991;338:464–8.
- 46 Popelka MM, Cruickshanks KJ, Wiley TL, et al. Moderate alcohol consumption and hearing loss: a protective effect. J Am Geriatr Soc 2000;48:1273–8.
- 47 Gopinath B, Flood VM, McMahon CM, *et al.* The effects of smoking and alcohol consumption on age-related hearing loss: the blue Mountains hearing study. *Ear Hear* 2010;31:277–82.
- 48 Itoh A, Nakashima T, Arao H, et al. Smoking and drinking habits as risk factors for hearing loss in the elderly: epidemiological study of subjects undergoing routine health checks in Aichi, Japan. Public Health 2001;115:192–6.
- 49 Hines LM, Rimm EB. Moderate alcohol consumption and coronary heart disease: a review. *Postgrad Med J* 2001;77:747–52.
- 50 Cruickshanks KJ, Wiley TL, Tweed TS, et al. Prevalence of hearing loss in older adults in Beaver dam, Wisconsin. The epidemiology of hearing loss study. Am J Epidemiol 1998;148:879–86.
- 51 Barsky AJ, Peekna HM, Borus JF. Somatic symptom reporting in women and men. *J Gen Intern Med* 2001;16:266–75.
- 52 McKee GJ, Stephens SD. An investigation of normally hearing subjects with tinnitus. *Audiology* 1992;31:313–7.
- 53 Dille MF, Konrad-Martin D, Gallun F, et al. Tinnitus onset rates from chemotherapeutic agents and ototoxic antibiotics: results of a large prospective study. J Am Acad Audiol 2010;21:409–17.

- 54 Pearson SE, Taylor J, Patel P, et al. Cancer survivors treated with platinum-based chemotherapy affected by ototoxicity and the impact on quality of life: a narrative synthesis systematic review. Int J Audiol 2019;58:685–95.
- 55 Veile A, Zimmermann H, Lorenz E, et al. Is smoking a risk factor for tinnitus? A systematic review, meta-analysis and estimation of the population attributable risk in Germany. BMJ Open 2018;8:e016589.
- 56 Loprinzi PD, Lee H, Gilham B, et al. Association between accelerometer-assessed physical activity and tinnitus, NHANES 2005-2006. Res Q Exerc Sport 2013;84:177–85.
- 57 Carpenter-Thompson JR, McAuley E, Husain FT. Physical activity, tinnitus severity, and improved quality of life. *Ear Hear* 2015;36:574–81.
- 58 Stuerz K, Lafenthaler M, Pfaffenberger N, et al. Body image and body concept in patients with chronic tinnitus. *Eur Arch Otorhinolaryngol* 2009;266:961–5.
- 59 McFerran DJ, Stockdale D, Holme R, et al. Why is there no cure for tinnitus? Front Neurosci 2019;13:802.
- 60 Trotter MI, Donaldson I. Hearing AIDS and tinnitus therapy: a 25-year experience. *J Laryngol Otol* 2008;122:1052–6.
- 61 Searchfield GD, Kaur M, Martin WH. Hearing aids as an adjunct to counseling: tinnitus patients who choose amplification do better than those that don't. *Int J Audiol* 2010;49:574–9.
- 62 Hoare DJ, Edmondson-Jones M, Sereda M, et al. Amplification with hearing AIDS for patients with tinnitus and co-existing hearing loss. *Cochrane Database Syst Rev* 2014:CD010151.
- 63 Grewal R, Spielmann PM, Jones SEM, et al. Clinical efficacy of tinnitus retraining therapy and cognitive behavioural therapy in the treatment of subjective tinnitus: a systematic review. J Laryngol Otol 2014;128:1028–33.
- 64 Nondahl DM, Cruickshanks KJ, Wiley TL, et al. The ten-year incidence of tinnitus among older adults. Int J Audiol 2010;49:580–5.