

More Attacks and Analgesic Use in Old Age: Self-Reported Headache Across the Lifespan in a German Sample

Müller, Britta; Dresler, Thomas; Gaul, Charly; Glass, Änne; Jürgens, Tim P.; Kropp, Peter; Ruscheweyh, Ruth; Straube, Andreas; Förderreuther, Stefanie

Veröffentlichungsversion / Published Version

Zeitschriftenartikel / journal article

Empfohlene Zitierung / Suggested Citation:

Müller, B., Dresler, T., Gaul, C., Glass, Ä., Jürgens, T. P., Kropp, P., ... Förderreuther, S. (2019). More Attacks and Analgesic Use in Old Age: Self-Reported Headache Across the Lifespan in a German Sample. *Frontiers in Neurology*, 10, 1-10. <https://doi.org/10.3389/fneur.2019.01000>

Nutzungsbedingungen:

Dieser Text wird unter einer CC BY Lizenz (Namensnennung) zur Verfügung gestellt. Nähere Auskünfte zu den CC-Lizenzen finden Sie hier: <https://creativecommons.org/licenses/by/4.0/deed.de>

Terms of use:

This document is made available under a CC BY Licence (Attribution). For more information see: <https://creativecommons.org/licenses/by/4.0>



More Attacks and Analgesic Use in Old Age: Self-Reported Headache Across the Lifespan in a German Sample

Britta Müller^{1*}, Thomas Dresler^{2,3}, Charly Gaul⁴, Änne Glass⁵, Tim P. Jürgens⁶, Peter Kropp¹, Ruth Ruscheweyh⁷, Andreas Straube⁷ and Stefanie Förderreuther⁷

¹ Institute of Medical Psychology and Medical Sociology, University Medicine Rostock, Rostock, Germany, ² Department of Psychiatry and Psychotherapy, University Hospital Tübingen, Tübingen, Germany, ³ LEAD Graduate School & Research Network, University of Tübingen, Tübingen, Germany, ⁴ Migraine and Headache Clinic Königstein, Königstein, Germany, ⁵ Institute for Biostatistics and Informatics in Medicine and Ageing Research, University Medicine Rostock, Rostock, Germany, ⁶ Department of Neurology, University Medicine Center Rostock, Rostock, Germany, ⁷ Department of Neurology, Ludwig Maximilian University of Munich, Munich, Germany

OPEN ACCESS

Edited by:

Massimiliano Valeriani,
Bambino Gesù Children Hospital
(IRCCS), Italy

Reviewed by:

Simona Guerzoni,
University of Modena and Reggio
Emilia, Italy
Piero Barbanti,
San Raffaele Pisana (IRCCS), Italy

*Correspondence:

Britta Müller
britta.mueller@med.uni-rostock.de

Specialty section:

This article was submitted to
Headache Medicine and Facial Pain,
a section of the journal
Frontiers in Neurology

Received: 05 July 2019

Accepted: 02 September 2019

Published: 17 October 2019

Citation:

Müller B, Dresler T, Gaul C, Glass Ä, Jürgens TP, Kropp P, Ruscheweyh R, Straube A and Förderreuther S (2019) More Attacks and Analgesic Use in Old Age: Self-Reported Headache Across the Lifespan in a German Sample. *Front. Neurol.* 10:1000. doi: 10.3389/fneur.2019.01000

Background: Reliable population-based data on the prevalence and characteristics of primary headache across the lifespan are essential. However, robust data are lacking.

Methods: We utilized questionnaire data from a random general population sample in Germany, that comprised 2,478 participants aged ≥ 14 years. A standardized questionnaire addressing headache and headache treatment was filled in during the face-to-face survey.

Results: The 6-month prevalence of self-reported headache in the total sample amounted to 39.0% (known diagnosis of migraine 7.2%; tension-type headache 12.4%; another diagnosis or unknown diagnosis 23.4%). Age-specific prevalence rates were 37.9% (14–34 years), 44.6% (35–54 years), 38.5% (55–74 years), and 26.9% (≥ 75 years). Compared to age group 14–34, participants aged 35–54 were more ($OR = 1.29$, 95%- CI 1.05–1.60, $p = 0.018$) and those aged ≥ 75 were less ($OR = 0.55$, 95%- CI 0.40–0.76, $p < 0.001$) likely to have any headache. Of the participants with headache, 79.5% reported headache on < 4 days per month, 15.6% on 4–14 days per month and 4.9% on > 14 days per month. The frequency of headache did not differ significantly between age groups in men [$\chi^2_{(3, N = 384)} = 1.45$, $p > 0.05$], but in women [$\chi^2_{(3, N = 651)} = 21.57$, $p < 0.001$]: women aged ≥ 75 years were over-represented in the group reporting 4–14 headache days per month. The analgesic use (days per month) differed significantly between age groups among participants with headache on < 4 days per month and on > 14 days per month: 1.8 (14–34 years), 2.5 (35–54 years), 3.2 (55–74 years), and 3.4 (≥ 75 years), respectively 7.9 (14–34 years), 11.4 (35–54 years), 18.4 (55–74 years), and 22.8 (≥ 75 years).

Conclusions: In general, the prevalence of headache decreases with age. However, older women suffer from more frequent attacks and older participants take analgesics

on more days per month than younger participants. This might put them at risk of medication overuse which may lead to medication overuse headache. More research is needed to understand these specifics in headache frequency and treatment behavior in older people.

Keywords: migraine, tension-type headache, lifespan, population-based study, 6-month prevalence, headache attacks, headache impact, use of analgesics

INTRODUCTION

Headache is the most frequent neurological disorder worldwide (1). According to estimates, the 1-year prevalence is around 38% for tension-type headache (TTH), and 10% for migraine (2, 3). These two types of headache account for 13% of all disability-adjusted life-years (DALYs) caused by neurological disorders (1). For purposes of healthcare planning, reliable population-based data on the characteristics of headache across the lifespan, such as prevalence, frequency of attacks and headache impact, are essential. Additional data on behavioral patterns across the lifespan in people with headache, such as physician consultation and use of medication, are also important. There are hardly any robust research results available.

Previous findings in prevalence refer almost exclusively to children and adolescents as well as young and middle-aged adults (4). Prevalence studies including older persons suffer from different limitations. First, a number of studies date back at least two decades (5–8). Second, some studies do not differentiate between different age groups among elderly people (6, 8–11). Third, many study samples make it difficult to generalize the findings to the whole population because they exclude either rural (12, 13) or urban residents (14–16). Age-specific patterns of attack frequency, medical use and headache impact have rarely been studied.

The aim of the present analysis is to (a) describe the 6-month prevalence of self-reported headache across the lifespan in general and (b) to identify age-specific characteristics of frequency of headache, headache impact and frequency of analgesic use within the German-speaking population aged ≥ 14 years.

METHODS

Participants

The analysis is based on data from a random general population sample in Germany with participants aged 14–94 years. A cross-sectional questionnaire survey was performed through face-to-face contacts conducted by an independent demographic consulting company (USUMA, Berlin, Germany). The sociodemographic data was collected using a standardized face-to-face interview. For the collection of headache-related data we used standardized questionnaires filled in by the participants themselves. For sample selection, random multi-stage sampling procedures were employed. First, 258 regional sample points

in Germany were determined (stage 1). Subsequently, 19 households per sample point were selected using the random-route procedure (stage 2) (17). Members of households, who met the inclusion criteria (age above 14, able to read, and understand German) were randomly selected using the Kish-selection-grid technique (stage 3) (18). The sample was designed to be representative in terms of age, sex, household size, and population by federal state. In total, 4,838 subjects were selected for the study, of whom 52% participated ($N = 2,510$). Reasons for non-participation were refusal to take part ($n = 1,453$), three unsuccessful attempts to contact the household member ($n = 863$), and illness of the selected subjects or his/her incapacity to follow the interview ($n = 8$). The final sample consisted of 2,510 interviewed participants. From the current analysis 32 subjects were excluded [reasons for exclusion were missing answers to following questions: (a) occurrence of headache in the last 6 months $n = 5$; (b) headache diagnosis $n = 13$; (c) headache frequency $n = 15$]. Thus, $n = 2,478$ interviewees were entered into the present analysis. A comparison of included subjects vs. drop-outs revealed no differences regarding sex composition, $\chi^2_{(1, N = 2,510)} = 1.09$, $p = 0.373$ and mean age, $U = 35947.50$; $p = 0.364$, $r = 0.02$. An adjustment factor was calculated on the basis of the German population structure regarding age, sex, household size, and population by federal state. German population parameters were provided by data from the 2016 Microcensus conducted by the German Federal Statistical Office. Based on the adjustment factor a weighted random sample was created, the structure of which corresponds to that of the German population (19). Data collection took place from September to November 2016. All participants gave their written informed consent. The Ethics Committee of the Faculty of Medicine, University of Leipzig, reviewed and approved the study (297/16-ek).

Questionnaire

The standardized questionnaire about headache and headache treatment consisted of 36 items. The introductory screening question was “Did you have a headache in the last 6 months?” The advantage over lifetime prevalence is that data depicting headache in a maximum of the last 12 months show a higher reliability due to a lower recall bias, particularly among older subjects (4). The headache frequencies were assessed using a five-point ordinal scale: (1) <1 day per month; (2) 1–3 days per month; (3) 4–14 days per month; (4) >14 days per month, but not daily; (5) daily. For statistical analysis we summarized the five categories into three classes: <4 days per month, 4–14 days per month, >14 days per month. In

Abbreviations: ICHD, International Classification of Headache Disorders; MOH, medication overuse headache; TTH, tension-type headache.

those participants having indicated the presence of headache, data regarding headache types was assessed by self-reporting using an item allowing multiple answers. The question used here was: “Do you know the diagnosis of your headache?” Response options were “migraine,” “TTH,” “cluster headache,” “other headache,” and “unclear headache (unknown diagnosis).” We summarized the headache types “cluster headache,” “other headache,” and “unclear headache (unknown diagnosis)” to the category “other headache.” As a reliable distinction between migraine with and without aura cannot be made solely by such a questionnaire, it was not assessed in our study (4). The impact of headache on daily life was assessed using the German version of the Headache Impact Test (HIT-6) (20). The HIT-6 is a six-item, self-administered questionnaire with three items assessing the impact of headache during the past 4 weeks and three items without a specific timeframe. Five response categories were given: “never,” “rarely,” “sometimes,” “very often,” and “always.” The total score can range from 36 to 78. Higher scores indicate a greater impact of headache on the ability to function on the job, at school, at home and in social situations. The HIT-6 provides a grading indicating four levels of headache impact: little or no impact (<50), some impact (50–55), substantial impact (56–59), and severe impact (≥ 60). To determine lifetime physician consultation for headache, participants were asked if they ever consulted a physician (or more than one) because of their headaches (yes/no). Analgesic use was assessed with the question: “How many days per month do you use analgesics on average?” Additionally, we used a standardized sociodemographic questionnaire to assess following parameters: age, sex, school education, and living alone (yes/no). Data regarding the community size was taken from the sampling plan.

Statistical Analysis

The data of the weighted random sample were analyzed using descriptive statistics. The 6-month prevalence of self-reported headache was estimated with 95% confidence interval (CI). To test for differences between sociodemographic characteristics and sex, we used Fisher’s exact test, Welch’s *t*-test and Pearson’s chi-squared test. To identify the factors associated with headache, we conducted a multivariate binary logistic regression that included sex and age group. We conducted ANOVAs to assess the effects of sex and age group on frequency of headache, analgesic use and headache impact (21). Differences were considered significant if the two-tailed *p*-value was $p < 0.05$. To analyze categorical variables we calculated adjusted Pearson residuals (22). We tested these residuals in the cells using a Bonferroni correction. For 6-month prevalence of headache, scatterplot smoothing LOESS curves were fitted to both men and women, with non-linear regression. Statistical analyses were performed using IBM SPSS Statistics 23.

RESULTS

Sociodemographic Characteristics

The sociodemographic characteristics of the weighted sample stratified for sex are presented in **Table 1**. The age of the participants ranged from 14 to 94 years.

6-Month Prevalence of Self-Reported Headache in Men and Women Across the Lifespan

In our sample, the weighted 6-month prevalence of self-reported headache amounted to 39.0%. 7.2% indicated to have received a diagnosis of migraine, 12.4% a diagnosis of TTH, and 23.4% had another diagnosis or had not received a diagnosis. 50.7% of the participants reporting headache had consulted a physician because of their headache, women (54.1%) more often than men (45.3%), $\chi^2_{(1, N = 950)} = 6.9, p = 0.009$.

Logistic regression analysis was employed to predict the probability of having headache in the last 6 months. Predictor variables were sex and age group (14–34 years, 35–54 years, 55–74 years, ≥ 75 years). The overall significance of the model was $\chi^2_{(4, N = 2478)} = 3194.99, p < 0.001$ (Likelihood-Ratio-Test). The analysis revealed both predictors remaining significant in the equation: sex (reference men) ($OR = 2.22, 95\%-CI 1.88-2.62, p < 0.001$) and age groups (reference 14–34 years): 35–54 years ($OR = 1.29, 95\%-CI 1.05-1.60, p = 0.018$), 55–74 years ($OR = 0.97, 95\%-CI 0.78-1.21, p = 0.800$), ≥ 75 years ($OR = 0.55, 95\%-CI 0.40-0.76, p < 0.001$). The odds ratio for sex indicated that women were 2.2 times more likely to have a headache than men. The age group variable was dummy coded using age group 14–34 years as the reference group. Participants aged 35–54 years were 1.29 times more likely to have a headache than participants aged 14–34 years. The odds of headache in age group 14–34 years were 1.82 times higher than for participants aged ≥ 75 years (inverted odds ratio for age group ≥ 75 years).

Univariate analyses indicated that the weighted prevalence of headache was higher in women compared to men in all headache categories: all headache types (male-to-female ratio 1:1.6), $\chi^2_{(1, N = 2,477)} = 85.6, p < 0.001$; migraine (male-to-female ratio 1:2.7), $\chi^2_{(1, N = 2,477)} = 42.1, p < 0.001$; TTH (male-to-female ratio 1:2), $\chi^2_{(1, N = 2,477)} = 42.1, p < 0.001$; other headaches (male-to-female ratio 1:1.3), $\chi^2_{(1, N = 2,477)} = 15.2, p < 0.001$. Prevalence rates of reported headache were 37.9% (14–34 years), 44.6% (35–54 years), 38.5% (55–74 years), and 26.9% (≥ 75 years). Participants aged 35–54 years were over-represented among headache participants (adjusted residual: 4.0), while participants aged ≥ 75 years less likely to have headache (adjusted residual: -4.4). In **Figure 1**, weighted age-specific 6-month prevalences are depicted, represented by scatterplot smoothing LOESS curves for men and women. Up to the age of around 45 years, the prevalence rose with age. After that age, prevalence declined with a steeper slope starting around 65 years (for more information see **Supplement Table 1**).

Frequency of Headache in Men and Women Across the Lifespan

Of the participants with headache, 79.5% reported headache on <4 days per month, 15.6% on 4–14 days per month and 4.9% on >14 days per month. We conducted a non-parametric two-way ANOVA to assess the effects of sex and age groups on the frequency of headache. The frequency of headache differed significantly for sex, $\chi^2_{(1, N = 1,035)} = 20.73, p < 0.001$, and age groups, $\chi^2_{(3, N = 1,035)} = 7.20, p < 0.05$. We found a significantly

TABLE 1 | Sociodemographic characteristics of the study sample according to sex.

Variable	Total sample	Men	Women	<i>p</i> -Value
	<i>n</i> = 2,477	<i>n</i> = 1,215	<i>n</i> = 1,262	
Age, mean (<i>SD</i>)	48.71 (19.15)	47.47 (18.91)	49.90 (19.31)	0.002 ^a
Age group, <i>n</i> (%)				0.059 ^b
14–24 years	319 (12.9)	166 (13.7)	153 (12.1)	
25–34 years	359 (14.5)	196 (16.1)	163 (12.9)	
35–44 years	360 (14.4)	172 (14.2)	188 (14.9)	
45–54 years	449 (18.1)	225 (18.5)	224 (17.7)	
55–64 years	388 (15.7)	191 (15.7)	197 (15.6)	
65–74 years	327 (13.2)	143 (11.8)	184 (14.6)	
75 years or older	275 (11.1)	122 (10.0)	153 (12.1)	
High school graduation (yes), <i>n</i> (%)	520 (21.1)	283 (23.3)	237 (18.9)	0.008 ^c
Marital status, <i>n</i> (%)				<0.001 ^b
Unmarried	721 (29.2)	407 (33.6)	314 (25.0)	
Married	1,266 (51.3)	641 (52.4)	625 (51.0)	
Divorced	258 (10.5)	119 (9.8)	139 (11.1)	
Widowed	223 (9.0)	44 (3.6)	179 (14.2)	
Living with partner (yes), <i>n</i> (%)	1,501 (61.1)	759 (63.1)	742 (59.1)	0.047 ^c
Living in a city (yes), <i>n</i> (%)	1,378 (55.7)	690 (56.8)	688 (54.6)	0.275 ^c
Community size, <i>n</i> (%)				0.468 ^b
<5,000 residents	101 (4.1)	48 (4.0)	53 (4.2)	
5,000–19,999 residents	227 (9.2)	119 (9.8)	108 (8.6)	
20,000–99,999 residents	430 (17.4)	199 (16.4)	231 (18.3)	
≥1,00,000 residents	1,718 (69.4)	849 (69.9)	869 (68.9)	

Weighted random sample.

^aWelch's *t*-test.

^bPearson's chi-squared test.

^cFisher's exact test; *SD*, standard deviation.

interaction between sex and age groups, $\chi^2_{(3, N = 1,035)} = 13.92$, $p < 0.05$. This interaction was due to the fact that women were under-represented among those with headache on <4 days per month (adjusted residual: -2.7). Given the hybrid type of interaction we conducted two separate non-parametric one-way ANOVAs for men and women. The frequency of headache did not differ significantly for age groups in men, $\chi^2_{(3, N = 384)} = 1.45$, $p > 0.05$, but in women, $\chi^2_{(3, N = 651)} = 21.57$, $p < 0.001$ (see **Figure 2A**). Women aged 35–54 years (adjusted residual: 3.0) were more likely and women aged ≥75 years (adjusted residual: -4.0) were less likely to have headache on <4 days per month. Additionally, women aged ≥75 years (adjusted residual: 3.1) were observed significantly more than expected among those with headache on 4–14 days per month. Detailed information on the frequency of headache regarding age group and sex is provided in the **Supplement Table 1**.

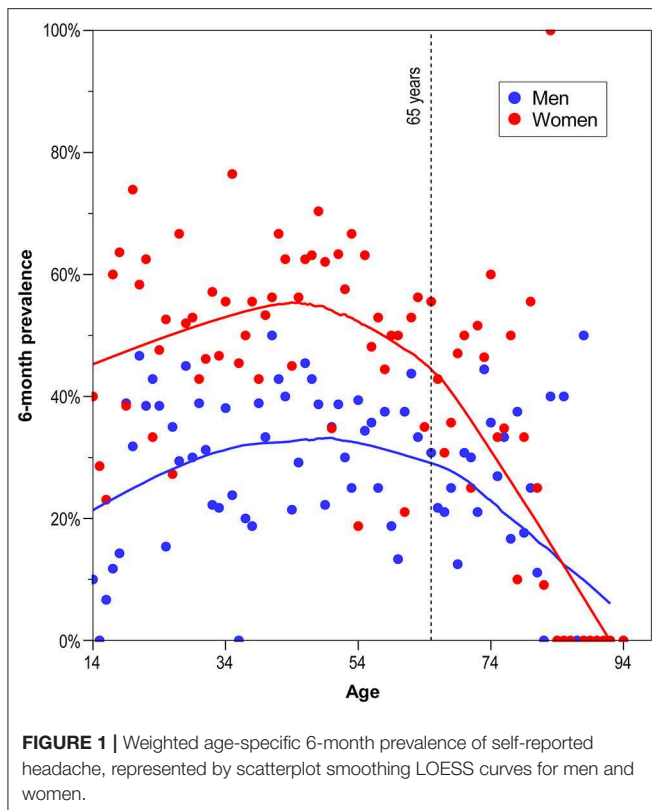
Headache Impact in Men and Women Across the Lifespan

41.6% of the participants with headache reported no or little headache impact, 25.1% some impact, 12.2% substantial impact and 21.1% severe impact (HIT-6). We conducted a parametric two-way ANOVA to assess the effects of sex and age groups on the total score of HIT-6. The headache impact differed significantly for sex, $F_{(1; 1,005)} = 36.34$, $p < 0.001$. Women (mean: 52.78, *SD*: 8.66) reported a stronger impact than men (mean: 49.24, *SD*:

8.08). Age groups did not differ significantly, $F_{(3; 1,005)} = 1.30$, $p = 0.274$. Mean values for headache impact for the four age groups 14–34 years, 35–54 years, 55–74 years, ≥75 years were 51.29 (*SD*: 8.72), 51.52 (*SD*: 8.08), 51.14 (*SD*: 8.90), and 53.32 (*SD*: 10.51). There was no significant interaction effect between sex and age groups, $F_{(3; 1,005)} = 0.98$, $p = 0.402$.

Frequency of Analgesic Use in Men and Women Across the Lifespan

Of the participants with headache, 43.3% reported taking analgesics on <2 days per month, 50.8% on 2–10 days per month and 5.9% on more than 10 days per month. A non-parametric two-way ANOVA was employed to assess the effects of sex and age groups on the frequency of analgesic use. The frequency of analgesic use differed statistically significant for sex, $\chi^2_{(1, N = 999)} = 30.80$, $p < 0.001$ and age groups $\chi^2_{(1, N = 999)} = 9.52$, $p = 0.025$. There was no significant interaction between sex and age groups to the frequency of analgesic use, $\chi^2_{(3, N = 999)} = 2.50$, $p > 0.05$ (see **Figure 2B**). The significant association between sex and analgesic use was due to the smaller numbers of women (adjusted residual: -5.6) within those with <2 days per month and the increased numbers of women (adjusted residual: 4.6) among those with analgesic use on 2–10 days per month and >10 days per month (adjusted residual: 2.1). The significant association between age groups and analgesic use was attributable to the higher than expected number of participants aged ≥75 years (adjusted residual: 4.0) among those



with analgesic use on >10 days per month (for more information see the **Supplement Table 1**).

Frequency of Analgesic Use in Men and Women Across the Lifespan Depending on the Frequency of Headache

In the following, we tested whether there is an association between age group and analgesic use, respectively sex and analgesic use even if the participants belong to the same headache frequency category. We conducted parametric two-way ANOVAs for this purpose to assess the effects of sex and age groups on analgesic use (number of days per month) in three different headache frequency categories: <4 days per month, 4–14 days per month, and >14 days per month.

In the headache frequency category “<4 days per month,” the analgesic use differed statistically significant for age groups $F_{(3; 788)} = 4.40, p = 0.004$. The age group means showed a linear trend ($p = 0.006$). According to the Tukey-HSD, age group 14–34 years, mean: 1.75 ($SD: 2.28$), differed significantly from age group 55–74 years, mean: 3.22 ($SD: 5.10$) ($-1.47, p = 0.002$). Sex did not differ significantly, $F_{(1; 788)} = 2.44, p = 0.119$. There was no significant interaction effect between sex and age groups, $F_{(3; 788)} = 0.35, p = 0.790$.

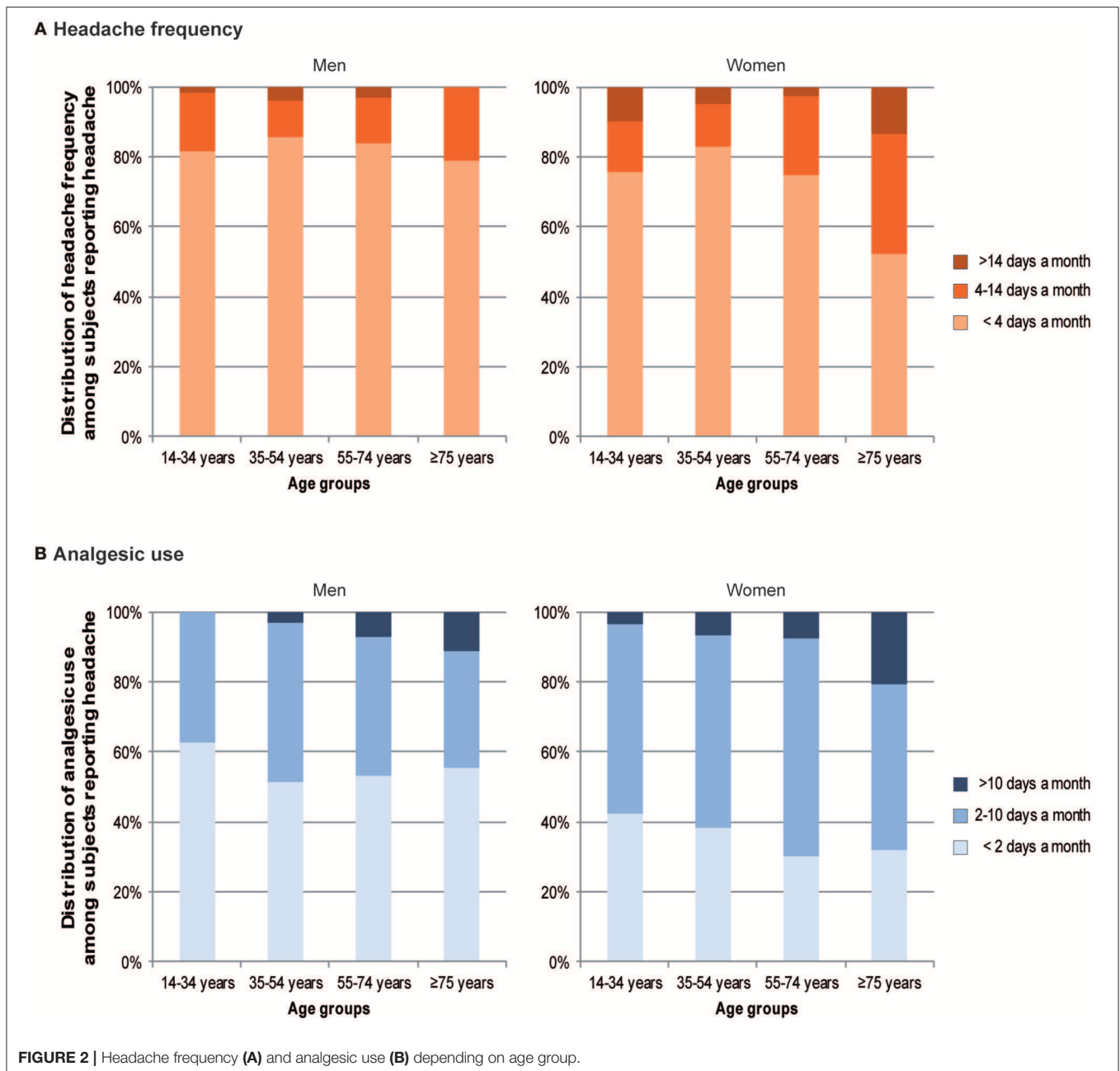
In the headache frequency category “4–14 days per month,” the analgesic use differed neither in age groups, $F_{(3; 153)} = 1.08, p = 0.361$, nor in sex, $F_{(1; 153)} = 1.30, p = 0.256$. There was no significant interaction effect between sex and age groups, $F_{(3; 153)} = 0.36, p = 0.790$.

In the headache frequency category “>14 days per month,” the analgesic use differed statistically significant for age groups, $F_{(3; 43)} = 4.09, p = 0.012$. The age group means showed a linear trend ($p = 0.003$). According to the Tukey-HSD, age group 14–34 years, mean: 7.89 ($SD: 6.96$), differed significantly from age group ≥ 75 years, mean: 22.83 ($SD: 11.84$) ($-14.94, p = 0.014$). Sex did not differ significantly, $F_{(1; 43)} = 1.77, p = 0.191$. There was no significant interaction effect between sex and age groups, $F_{(2; 43)} = 0.78, p = 0.465$ (see **Figure 3**).

DISCUSSION

We analyzed the 6-month prevalence and characteristics of self-reported headache across the lifespan in a random general population sample in Germany aged 14–94 years.

In our study, the overall headache prevalence in the past 6 months in Germany, as obtained with a questionnaire, was 39%. This frequency is slightly lower than the mean prevalence in Europe (53%) reported by Stovner and Andree (4). They evaluated 49 studies with different timeframes of headache prevalence (a third of them contained lifetime prevalence data) and different methods of data collection (nearly a third of studies with a timeframe for headache ≤ 1 year were based on personal interviews). Both, lifetime prevalence of headache (compared to a timeframe for headache ≤ 1 year) and prevalence rates based on personal interviews (compared to questionnaires) result in higher figures. The prevalence rate in our German study is comparable to Norway (38%) (23), Estonia (41%) (24), and Italy (43%) (12). Furthermore, we found a 6-month prevalence of self-reported diagnosed migraine of 7.2% with a male-to-female ratio of 1:2.7. The mean prevalence of migraine in Europe is two times higher (14.7%) with a quite similar male-to-female ratio of 1:2.2 (14.7%) (4). The difference is most likely due to half of the men and women with migraine in Germany not knowing their headache diagnosis. This result supports findings of other studies (25, 26). The 6-month prevalence of self-reported diagnosed TTH in our study (12.4%) is approximately five times lower than the mean prevalence in Europe (62.6%) (4). The male-to-female ratio in TTH of our sample (1:2) differs only slightly from that in other European studies (1:1.3). The proportions of women and men with TTH who do not know their diagnosis are similarly high. It is estimated that people with headache underreport TTH more often than migraine due to the lesser severity of symptoms and impact of TTH (27–30). 23.4% of all participants had another diagnosis or did not know their diagnosis, which corresponds to 60% of all those suffering from headache. This result is at glance surprising, as maximally 8–10% of all headache patients suffer from secondary headaches only, and just about 1–2% of the headache patients suffer from primary headaches other than migraine or TTH (31–36). Our conclusion is that about 35% of the population in Germany has primary headaches and that about 44% of the patients with primary headache are not aware of their specific diagnosis. This fits well with the result that 49% of the participants with headache had not seen a physician because of their headache (and consequently not received a diagnosis). However, diagnosing or classifying the individual

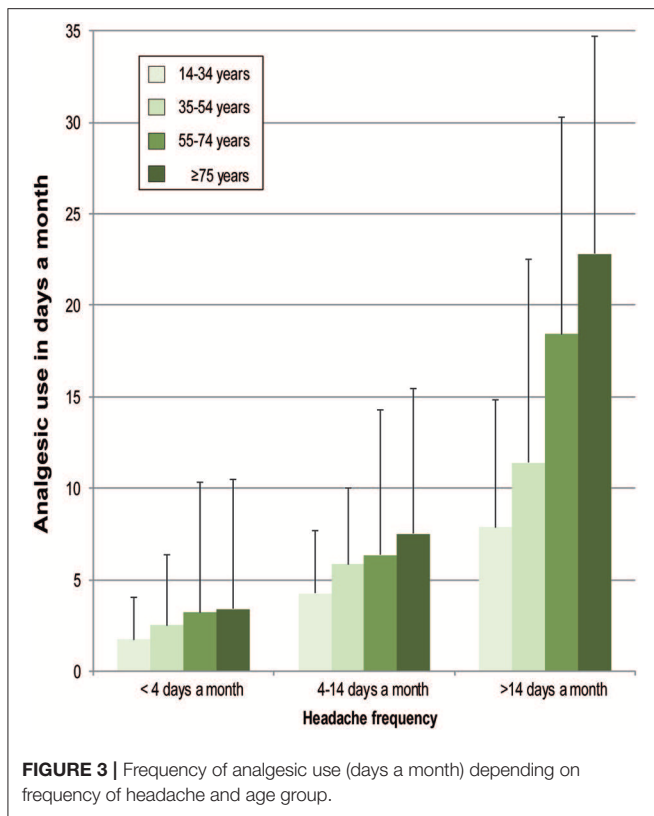


headache allows for an appropriate treatment and management (37–41). The low consultation rate probably reflects the general opinion that headaches cannot be treated specifically, or that self-medication is sufficient.

We found that the headache prevalence was associated with sex and age group. Higher headache prevalence in women is well-documented in the literature (4, 16, 42, 43). Compared to the age group 14–34 years, participants aged 35–54 years are more and those aged ≥ 75 years are less likely to have headache (inverted u-shape, see Figure 1). This likely reflects the known age-specific prevalence rates of migraine and TTH. Migraine reaches a peak during the fourth decade of life and remains constant during the

fifth decade, TTH peaks between the ages of 30 and 39 years (44, 45). The lowest prevalence of TTH and migraine is at age 70 years and older (13, 42, 44, 46, 47).

Of the participants with headache, 80% reported headache on < 4 days per month. Lipton et al. (29) reported a very similar proportion (79%) for participants with migraine in an American sample. Although in clinical practice, in addition to attack frequency, other criteria (e.g., duration of attack, impairment of quality of life) are considered when deciding on preventive treatment, we assume from an epidemiological perspective that about 80% of all participants with headache do not need preventive therapy (48). In our sample, of the



participants with headache, 5% reported chronic headache (>14 days per month), which is about 2% of our total sample. Stovner et al. (49) estimated, based on data from 8 European countries, that about 4% of the adult population in Europe suffers from chronic headache. Data from Germany were not included. Our data suggest that the proportion among the population aged ≥ 14 in Germany is lower.

In the following we discuss the three most important results of our study:

First, we could show that in women, but not in men, the frequency of headache attacks differs between age groups. Women in old age (≥ 75 years) tend to have more headache days (per month) than younger women. This tendency is known for migraine patients (15, 50). However, it has not yet been analyzed in the literature whether this trend applies to both men and women. It is possible that the association between age and frequency of migraine attacks exists only in women. The cause of higher attack frequency in older patients is not known. It could be hypothesized that only patients with primarily severe migraine still have it in older age and that more patients with less severe migraine become headache-free in older age. In this case, the higher headache frequency in old age would be the result of a selection effect. Furthermore, the polypharmacy in older patients may exacerbate a pre-existing headache or trigger medication overuse headache (MOH), e.g., regular intake of analgesics for chronic arthritis pain (46, 51, 52).

Second, the impact of headache did not differ between age groups. We found no other study in which the impact of

headache was analyzed across the lifespan. Given that among women the frequency of attacks increases with age, the finding of a stable headache impact appears paradoxical. Plausible would have been an increase in headache impact with age. We suppose that psychological adaptation processes, according to the theory of Rothermund and Brandstädter (53) about coping with deficits and losses in later life, play an important role in the subjective perception of the headache impact. It is possible that, based on the longer duration of the disease, headache sufferers in old age are better able to cope with their condition than younger ones. They may be more efficient able to generate affectively relieving cognitions.

Third, we could show that among participants with a headache frequency of >14 days per month those aged ≥ 75 years use analgesics on more days per month than younger participants. The reason is unclear. In our study, we used a categorical variable to measure headache frequency. We do not have data about the actual number of headache days per month. It is possible that older people with >14 headache days per month have a higher frequency of headache than younger people. In this case, the higher headache frequency could result in a higher burden of disease, which is compensated by a more frequent intake of analgesics. It is also conceivable, however, that older people with the same headache frequency as younger people may take more analgesics than younger ones. However, it could also be due to the intake of analgesics for treatment of concomitant (non-headache) painful conditions, which are more frequent with increasing age. Furthermore, the more frequent intake of analgesics can also be the reason for higher blood pressure in elderly persons (54). Studies indicate that roughly half of the individuals with headache on >14 days per month use analgesics daily or almost daily (55, 56). Our data suggest that the risk may be higher in old age.

Physicians should pay attention to the frequency of headache attacks and medication use in older people. The more frequent intake of analgesics might put older people at risk of developing medication overuse and subsequently MOH (57). Especially for older women who consult a physician for non-headache pain, taking the history should also include questions about self-medication to treat headache. General practitioners play a key role, as they are often consulted by older patients. They should be aware of the possibility of MOH and refer the patients to a specialist if necessary. Based on the assumption that the higher analgesic use of older subjects was mainly caused by headache, our data implies an unmet need of pharmacological management of headache in the elderly. Pharmacological headache trials should therefore not be limited to the age of 65, but should include also individuals aged 70–75 years or even older.

LIMITATIONS

An important limitation is the fact that our headache type classification was based on participants' self-reports of prior headache diagnoses, necessarily underestimating migraine and TTH prevalence. It is possible that the category "other" includes many undiagnosed migraine and TTH patients. In this regard,

our results on prevalence rates are less accurate than studies that reported prevalence based on ICHD III beta criteria. Furthermore, we cannot say whether the attacks and more frequent use of analgesics in old age are due to migraine or TTH, nor whether they are due to primary or secondary headache. We assume that the response to the question about the use of analgesics was influenced by an anchoring effect (58): the headache-related questions were preceded by an introductory sentence (“The following questions deal with headaches. Please answer as accurately as you can.”); the question about analgesic use was asked in the latter part of the headache-related questionnaire and followed immediately after three questions related directly to the treatment of headache. Nevertheless, we cannot rule out the possibility that the analgesics may have been taken because of other painful, not headache-related symptoms. Especially for chronic headache, strong associations to musculoskeletal symptoms and to fibromyalgia have been reported (59, 60). It is still unclear whether these relationships are moderated by age.

CONCLUSION

In our representative German sample of participants aged ≥ 14 years and living in private households, about 39% suffer from headache, and many participants with headache do not know their diagnosis. We estimate this proportion at about 44%, based on the proportions of participants reporting migraine and TTH diagnoses compared to the known prevalences of these disorders. Headache prevalence peaks in the age group 35–54 years, the lowest prevalence was found in the age group 75 years and more. Women in old age have more headache days per month than younger women. Among participants with a headache frequency of >14 days per month, those aged ≥ 75 years use analgesics on more days per month than younger participants, putting them at risk for MOH. More research is needed to

understand these specifics of headache in old age. As a first step toward an effective treatment of primary headaches in the general population, it also seems necessary to encourage people suffering from headache to contact a physician and receive a headache diagnosis.

DATA AVAILABILITY STATEMENT

The datasets generated for this study are available on request to the corresponding author.

AUTHOR CONTRIBUTIONS

TD, CG, TJ, PK, AS, and SF contributed conception of the study. BM and ÄG performed the statistical analysis. BM wrote the first draft of the manuscript. BM, TD, CG, TJ, PK, RR, SF, and AS reviewed and edited the manuscript. All authors approved the final version of the manuscript.

FUNDING

This study was funded by the German Migraine and Headache Society (DMKG e.V., <http://dmkg.de/>).

ACKNOWLEDGMENTS

We would like to thank all participants of this study.

SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: <https://www.frontiersin.org/articles/10.3389/fneur.2019.01000/full#supplementary-material>

Supplement Table 1 | Percentage of participants with headache reporting selected characteristics (in %, with 95% CI). Weighted random sample.

REFERENCES

- Feigin VL, Abajobir AA, Abate KH, Abd-Allah F, Abdulle AM, Abera SF. Global, regional, and national burden of neurological disorders during 1990–2015: a systematic analysis for the Global Burden of Disease Study 2015. *Lancet Neurol.* (2017) 16:877–97. doi: 10.1016/S1474-4422(17)30299-5
- Woldeamanuel YW, Cowan RP. Migraine affects 1 in 10 people worldwide featuring recent rise: a systematic review and meta-analysis of community-based studies involving 6 million participants. *J Neurol Sci.* (2017) 372:307–15. doi: 10.1016/j.jns.2016.11.071
- Stovner Lj, Hagen K, Jensen R, Katsarava Z, Lipton R, Scher A, et al. The global burden of headache: a documentation of headache prevalence and disability worldwide. *Cephalalgia.* (2007) 27:193–210. doi: 10.1111/j.1468-2982.2007.01288.x
- Stovner LJ, Andree C. Prevalence of headache in Europe: a review for the Eurolight project. *J Headache Pain.* (2010) 11:289. doi: 10.1007/s10194-010-0217-0
- Waters W, O’connor P. Prevalence of migraine. *J Neurol Neurosurg Psychiatry.* (1975) 38:613–6. doi: 10.1136/jnnp.38.6.613
- Nikiforow R. Headache in a random sample of 200 persons: a clinical study of a population in northern Finland. *Cephalalgia.* (1981) 1:99–107. doi: 10.1111/j.1468-2982.1981.tb00016.x
- D’Alessandro R, Benassi G, Lenzi PL, Gamberini G, Sacquegna T, De Carolis P, et al. Epidemiology of headache in the Republic of San Marino. *J Neurol Neurosurg Psychiatry.* (1988) 51:21–7. doi: 10.1136/jnnp.51.1.21
- Castillo J, Muñoz P, Guitera V, Pascual J. Epidemiology of chronic daily headache in the general population. *Headache.* (1999) 39:190–6. doi: 10.1046/j.1526-4610.1999.3903190.x
- Katsarava Z, Dzagnidze A, Kukava M, Mirvelashvili E, Djibuti M, Janelidze M, et al. Primary headache disorders in the Republic of Georgia Prevalence and risk factors. *Neurology.* (2009) 73:1796–803. doi: 10.1212/WNL.0b013e3181c34abb
- Dahlöf C, Linde M. One-year prevalence of migraine in Sweden: a population-based study in adults. *Cephalalgia.* (2001) 21:664–71. doi: 10.1046/j.1468-2982.2001.00218.x
- Boardman HF, Thomas E, Millson DS, Croft PR. One-year follow-up of headache in an adult general population. *Headache.* (2005) 45:337–45. doi: 10.1111/j.1526-4610.2005.05071.x
- Taga A, Russo M, Manzoni GC, Torelli P. The PACE study: lifetime and past-year prevalence of headache in Parma’s adult general population. *Neurol Sci.* (2017) 38:789–95. doi: 10.1007/s10072-017-2845-5

13. Schwaiger J, Kiechl S, Seppi K, Sawires M, Stockner H, Erlacher T, et al. Prevalence of primary headaches and cranial neuralgias in men and women aged 55–94 years (Bruneck Study). *Cephalalgia*. (2009) 29:179–87. doi: 10.1111/j.1468-2982.2008.01705.x
14. Camarda R, Monastero R. Prevalence of primary headaches in Italian elderly: preliminary data from the Zabut Aging Project. *Neurol Sci*. (2003) 24:s122–4. doi: 10.1007/s100720300058
15. Prencipe M, Casini AR, Ferretti C, Santini M, Pezzella F, Scaldaferrì N, et al. Prevalence of headache in an elderly population: attack frequency, disability, and use of medication. *J Neurol Neurosurg Psychiatry*. (2001) 70:377–81. doi: 10.1136/jnnp.70.3.377
16. Zhang Y, Shi Z, Hock D, Yue W, Liu S, Zhang Y, et al. Prevalence of primary headache disorders in a population aged 60 years and older in a rural area of Northern China. *J Headache Pain*. (2016) 17:83. doi: 10.1186/s10194-016-0672-3
17. Hoffmeyer-Zlotnik JH. Random-route-stichproben nach ADM. In: *Stichproben in der Umfragepraxis*. Oxford: Springer (1997). p. 33–42. doi: 10.1007/978-3-322-86533-5_4
18. Gaziano C. Comparative analysis of within-household respondent selection techniques. *Public Opin Q*. (2005) 69:124–57. doi: 10.1093/poq/nfi006
19. Schmich P, Johannes L, Marie-Luise Z, Anja M, Jennifer A, Matthias W. *Ad-hoc-Studien im Robert Koch-Institut* (2018).
20. Kosinski M, Bayliss MS, Bjorner JB, Ware JE Jr, Garber WH, Batenhorst A, et al. A six-item short-form survey for measuring headache impact: the HIT-6TM. *Qual Life Res*. (2003) 12:963–74. doi: 10.1023/A:1026119331193
21. Winer BJ. *Statistical Principles in Experimental Design* (1991).
22. Agresti A, Kateri, M. *Categorical Data Analysis*. Oxford: Springer (2011). doi: 10.1007/978-3-642-04898-2_161
23. Hagen K, Zwart JA, Vatten L, Stovner LJ, Bovim G. Prevalence of migraine and non-migrainous headache—head-HUNT, a large population-based study. *Cephalalgia*. (2000) 20:900–6. doi: 10.1046/j.1468-2982.2000.00145.x
24. Toom K, Raidvee A, Allas KH, Floria E, Juhkami K, Klimušev G, et al. The prevalence of primary headache disorders in the adult population of Estonia. *Cephalalgia*. (2019) 39:883–91. doi: 10.1177/0333102419829909
25. Rasmussen BK, Jensen R, Olesen J. Impact of headache on sickness absence and utilisation of medical services: a Danish population study. *J Epidemiol Community Health*. (1992) 46:443–6. doi: 10.1136/jech.46.4.443
26. Bigal ME, Krymchantowski AV, Lipton RB. Barriers to satisfactory migraine outcomes. What have we learned, where do we stand? *Headache*. (2009) 49:1028–41. doi: 10.1111/j.1526-4610.2009.01410.x
27. Lavados P, Tenhamm E. Consulting behaviour in migraine and tension-type headache sufferers: a population survey in Santiago, Chile. *Cephalalgia*. (2001) 21:733–7. doi: 10.1046/j.1468-2982.2001.00217.x
28. Edmeads J, Findlay H, Tugwell P, Pryse-Phillips W, Nelson RF, Murray TJ. Impact of migraine and tension-type headache on life-style, consulting behaviour, and medication use: a Canadian population survey. *Can J Neurol Sci*. (1993) 20:131–7. doi: 10.1017/S0317167100047697
29. Lipton RB, Bigal ME, Diamond M, Freitag F, Reed ML, Stewart WF, et al. Migraine prevalence, disease burden, and the need for preventive therapy. *Neurology*. (2007) 68:343–9. doi: 10.1212/01.wnl.0000252808.97649.21
30. Bussone G, Usai S, Grazi L, Rigamonti A, Solari A, D'Amico D. Disability and quality of life in different primary headaches: results from Italian studies. *Neurol Sci*. (2004) 25:s105–7. doi: 10.1007/s10072-004-0263-y
31. Dodick DW. Pearls: headache. *Semin Neurol*. (2010) 30:74–81. doi: 10.1055/s-0029-1245000
32. Katsarava Z, Obermann M, Yoon MS, Dommes P, Kuznetsova J, Weimar C, et al. Prevalence of cluster headache in a population-based sample in Germany. *Cephalalgia*. (2007) 27:1014–9. doi: 10.1111/j.1468-2982.2007.01380.x
33. Ekblom K, Svensson DA, Pedersen NL, Waldenlind E. Lifetime prevalence and concordance risk of cluster headache in the Swedish twin population. *Neurology*. (2006) 67:798–803. doi: 10.1212/01.wnl.0000233786.72356.3e
34. Sjaastad O, Bakkeiteig L. Cluster headache prevalence. Vågå study of headache epidemiology. *Cephalalgia*. (2003) 23:528–33. doi: 10.1046/j.1468-2982.2003.00585.x
35. Torelli P, Beghi E, Manzoni GC. Cluster headache prevalence in the Italian general population. *Neurology*. (2005) 64:469–74. doi: 10.1212/01.WNL.0000150901.47293.BC
36. Antonaci F, Sjaastad O. Chronic paroxysmal hemicrania (CPH): a review of the clinical manifestations. *Headache*. (1989) 29:648–56. doi: 10.1111/j.1526-4610.1989.hed2910648.x
37. Gaul C, Liesering-Latta E, Schäfer B, Fritsche G, Holle D. Integrated multidisciplinary care of headache disorders: a narrative review. *Cephalalgia*. (2016) 36:1181–91. doi: 10.1177/0333102415617413
38. Sorgenfrei V, Kropp P, Straube A, Ruscheweyh R. High achievement motivation is not related to increased use of acute headache medication in migraine: a cross-sectional observational cohort study. *Headache*. (2018) 58:1629–38. doi: 10.1111/head.13431
39. Kropp P, Meyer B, Dresler T, Fritsche G, Gaul C, Niederberger U, et al. Relaxation techniques and behavioural therapy for the treatment of migraine: guidelines from the German Migraine and Headache Society. *Schmerz*. (2017) 31:433–47. doi: 10.1007/s00482-017-0214-1
40. Meyer B, Keller A, Wöhlbier HG, Overath CH, Müller B, Kropp P. Progressive muscle relaxation reduces migraine frequency and normalizes amplitudes of contingent negative variation (CNV). *J Headache Pain*. (2016) 17:37. doi: 10.1186/s10194-016-0630-0
41. Martelletti P, Jensen RH, Antal A, Arcioni R, Brighina F, de Tommaso M, et al. Neuromodulation of chronic headaches: position statement from the European Headache Federation. *J Headache Pain*. (2013) 14:86. doi: 10.1186/1129-2377-14-86
42. Straube A, Aicher B, Förderreuther S, Eggert T, Köppel J, Möller S, et al. Period prevalence of self-reported headache in the general population in Germany from 1995–2005 and 2009: results from annual nationwide population-based cross-sectional surveys. *J Headache Pain*. (2013) 14:11. doi: 10.1186/1129-2377-14-11
43. Rastenyte D, Mickevičiene D, Stovner LJ, Thomas H, Andrée C, Steiner TJ. Prevalence and burden of headache disorders in Lithuania and their public-health and policy implications: a population-based study within the Eurolight project. *J Headache Pain*. (2017) 18:53. doi: 10.1186/s10194-017-0759-5
44. Jensen R, Stovner LJ. Epidemiology and comorbidity of headache. *Lancet Neurol*. (2008) 7:354–61. doi: 10.1016/S1474-4422(08)70062-0
45. Radtke A, Neuhauser H. Prevalence and burden of headache and migraine in Germany. *Headache*. (2009) 49:79–89. doi: 10.1111/j.1526-4610.2008.01263.x
46. Tonini MC, Bussone G. Headache in the elderly: primary forms. *Neurol Sci*. (2010) 31:67–71. doi: 10.1007/s10072-010-0278-5
47. Straube A, Andreou A. Primary headaches during lifespan. *J Headache Pain*. (2019) 20:35. doi: 10.1186/s10194-019-0985-0
48. Diener H, Gaul C, Kropp P. Therapie der Migräneattacke und Prophylaxe der Migräne, S1-Leitlinie. In: Deutsche Gesellschaft für Neurologie, editor. *Leitlinien für Diagnostik und Therapie in der Neurologie*. (2018). Available online at: <https://www.dgn.org/leitlinien> (accessed July 22, 2018).
49. Stovner LJ, Zwart JA, Hagen K, Terwindt GM, Pascual J. Epidemiology of headache in Europe. *Eur J Neurol*. (2006) 13:333–45. doi: 10.1111/j.1468-1331.2006.01184.x
50. Bigal ME, Liberman JN, Lipton RB. Age-dependent prevalence and clinical features of migraine. *Neurology*. (2006) 67:246–51. doi: 10.1212/01.wnl.0000225186.76323.69
51. Lipton RB, Pfeiffer D, Newman LC, Solomon S. Headaches in the elderly. *J Pain Symptom Manag*. (1993) 8:87–97. doi: 10.1016/0885-3924(93)90106-6
52. Song TJ, Kim YJ, Kim BK, Kim BS, Kim JM, Kim SK, et al. Characteristics of elderly-onset (≥ 65 years) headache diagnosed using the International Classification of Headache Disorders, beta version. *J Clin Neurol*. (2016) 12:419–25. doi: 10.3988/jcn.2016.12.4.419
53. Rothermund K, Brandstädter J. Coping with deficits and losses in later life: from compensatory action to accommodation. *Psychol Aging*. (2003) 18:896. doi: 10.1037/0882-7974.18.4.896

54. Wilson SL, Poulter NR. The effect of non-steroidal anti-inflammatory drugs and other commonly used non-narcotic analgesics on blood pressure level in adults. *J Hypertens.* (2006) 24:1457–69. doi: 10.1097/01.hjh.0000239278.82196.a5
55. Zwart JA, Dyb G, Hagen K, Svebak S, Stovner LJ, Holmen J. Analgesic overuse among subjects with headache, neck, and low-back pain. *Neurology.* (2004) 62:1540–4. doi: 10.1212/01.WNL.0000123262.96132.FC
56. Straube A, Pfaffenrath V, Ladwig KH, Meisinger C, Hoffmann W, Fendrich K, et al. Prevalence of chronic migraine and medication overuse headache in Germany—the German DMKG headache study. *Cephalalgia.* (2010) 30:207–13. doi: 10.1111/j.1468-2982.2009.01906.x
57. Natoli JL, Manack A, Dean B, Butler Q, Turkel CC, Stovner L, et al. Global prevalence of chronic migraine: a systematic review. *Cephalalgia.* (2010) 30:599–609. doi: 10.1111/j.1468-2982.2009.01941.x
58. Furnham A, Boo HC. A literature review of the anchoring effect. *J Socio-Economics.* (2011) 40:35–42. doi: 10.1016/j.socsec.2010.10.008
59. Hagen K, Einarsen C, Zwart JA, Svebak S, Bovim G. The co-occurrence of headache and musculoskeletal symptoms amongst 51 050 adults in Norway. *Eur J Neurol.* (2002) 9:527–33. doi: 10.1046/j.1468-1331.2002.00451.x
60. Onder H, Hamamci M, Alpua M, Ulusoy EK. Comorbid fibromyalgia in migraine patients: clinical significance and impact on daily life. *Neurol Res.* (2019) 2019:1–7. doi: 10.1080/01616412.2019.1630164

Conflict of Interest: TD has received honoraria for consulting within the past 3 years from Novartis Pharma. CG has received honoraria for consulting and lectures within the past 3 years from Allergan Pharma, Bayer vital, Boehringer Ingelheim Pharma, Cerbotec, Desitin Arzneimittel, electroCore, Grünenthal, Hormosan Pharma, Lilly Germany, Novartis Pharma, Ratiopharm, Reckitt Benckiser, Sanofi Aventis, and TEVA. TJ has received honoraria for consulting and lectures from Allergan Pharma, Autonomic Technologies, Desitin Arzneimittel, Lilly Germany, Novartis Pharma, and TEVA. PK has received honoraria for consulting and lectures from Novartis Pharma and Shire. RR has received honoraria for consulting and lectures within the past 3 years from Allergan Pharma, Hormosan Pharma, Novartis Pharma, Lilly Germany, and TEVA. AS has received honoraria for consulting and educational lectures from Allergan Pharma, Bayer, Desitin Arzneimittel, electroCore, Lilly Germany, Novartis Pharma, Sanofi Aventis, and TEVA. SF has received honoraria for consulting and lectures within the past 3 years from Allergan Pharma, Astra Zeneca, Hormosan Pharma, Lilly Germany, Novartis Pharma, Sanofi Aventis, and TEVA.

The remaining authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Copyright © 2019 Müller, Dresler, Gaul, Glass, Jürgens, Kropp, Ruscheweyh, Straube and Förderreuther. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.