Healthcare Expenditures Associated with Comorbid Anxiety and Depression among Adults with Migraine

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Abstract

Objective. To examine direct healthcare expenditures among adults with migraine alone or with comorbid anxiety and/or depression.

Methods. This was a retrospective cross-sectional study using the 2012, 2014, and 2016 Medical Expenditure Panel Survey data. Adult patients aged ≥22 years with migraine headache were included in the study. The direct healthcare expenditures of four migraine groups (migraine alone, migraine and anxiety, migraine and depression, migraine and both conditions) were compared.

Results. There were 1,556 patients met the inclusion criteria and were eventually enrolled in the study. Approximately, 50% of the study sample had migraine with comorbid depression and/or anxiety (16.1% have depression, 12.3% have anxiety disorder, and 13.9% have both). The mean total healthcare expenditures of adults with migraine alone ($6,461) were significantly lower than those with comorbid depression and anxiety ($11,102), comorbid anxiety ($10,817), and comorbid depression ($14,577). Migraine with comorbid anxiety and depression was significantly associated with incremental costs of $1,027 in outpatient’s and $662 emergency room healthcare expenditures, compared to the migraine alone group after controlling for a multitude of covariates. Besides, migraine with comorbid anxiety and/or depression was significantly associated with incremental costs of prescription drug, compared to the migraine alone group.

Conclusions. The healthcare expenditures associated with migraine with comorbid depression and/or anxiety are significantly higher than those without mental health comorbidities. Therefore, regular depression and anxiety screening for patients with migraine may help in reducing the healthcare expenditures associated with
depression and/or anxiety comorbidities as well as in improving the quality of care.

Introduction

Migraine headache is a highly prevalent chronic condition among adults in the United States [1]. The World Health Organization (WHO) ranked migraine headache as the third cause of disability measured in years lived with disability (YLDs) [1]. Migraine is associated with poor Health-Related Quality of Life [2, 3], disability [3, 4], and missed working days. Furthermore, it is associated with a great economic burden, in terms of direct medical costs [5-7] and indirect costs due to absenteeism in the workplace [8]. Individuals with migraine are more likely to have higher healthcare expenditures compared to those without migraine [5, 6]. Also, individuals with migraine have a higher rate of healthcare services utilization, such as emergency room and physician office visits, compared to those without migraine [5, 9].

Depression and/or anxiety are common among patients with migraine [10]. The likelihood of having major depressive disorder among migraine is significantly higher than their counterparts without migraine [11]. Furthermore, it has been estimated that adults with migraine are two to four times more likely to suffer from depression and anxiety in comparison to the general population [12-15]. The comorbidity of depressive and anxiety disorders among migraine patients is associated with a high prevalence rate of disability [16], poor Health-Related Quality of Life (HRQoL) [3, 12, 16], and high rate of healthcare utilization and costs [17].

The economic burden of migraine is high with an estimated annual cost of $17 billion in the U.S. alone [18]. However, this economic burden is significantly higher
among patients with comorbid depression and/or anxiety [17, 19]. Several studies have indicated that the healthcare expenditures associated with migraine with comorbid depression and/or anxiety can be double that of patients with migraine alone [17, 20]. In a cohort study among adults with migraine using a one year data of a large employer-based insurance database, the presence of depression and/or anxiety along with migraine have resulted in higher direct medical costs than those with migraine alone ($5589.5 migraine alone vs $10223.4 with anxiety and $10582.2 with depression) [17]. Moreover, in a cross-sectional study that used the Medical Expenditure Panel Survey (MEPS) data and included 2,400 subjects with migraine between 2006-2012, the mean annual total healthcare expenditures as well as migraine-related healthcare expenditures were significantly higher among migraine patients with depression compared to those without a comorbid depression ($10,012 vs. $4,740, \( P < .001 \)) and ($723 vs $499, \( P=0.014 \)), respectively [20]. However, studies that compared the differences in healthcare expenditures among adults with migraine with or without comorbid anxiety and/or depression using nationally representative data are scarce. Besides, the impacts of anxiety and anxiety with depression comorbidity on healthcare expenditures among migraine patients have not been identified before. Thus, the aim of this study was to examine the differences in the total healthcare expenditures as well as the types of healthcare expenditures among adults with migraine alone as well as among those with comorbid anxiety and/or depression taking into consideration a myriad of factors that are likely to influence healthcare expenditures among this segment of the patient population.

Methods
Study Design

This was a retrospective cross-sectional study using the Medical Expenditure Panel Survey (MEPS) data of 2012, 2014, and 2016. MEPS is a nationally representative survey conducted by the Agency for Healthcare Research and Quality (AHRQ) of the US noninstitutionalized civilian population (10). MEPS has information on demographics, socioeconomic characteristics, medical conditions, health status, and other health-related data. The International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM) codes was used to report the medical conditions. Beginning in 2016, the International Classification of Diseases, Tenth Revision, Clinical Modification (ICD-10-CM) codes used to report medical conditions.

Study Population

Adults aged 22-64 years with migraine headache (based upon ICD-9-code of ‘346’, ICD-10-code of ‘G43’) who were alive during the calendar years of 2012, 2014, and 2016, were included in the study.

Measures

Dependent Variables

Type and total healthcare expenditures

The types of healthcare expenditures, such as inpatient, outpatient, prescription, emergency room, and other healthcare expenditures (e.g., dental, vision, and durable medical equipment use, and others), were included in the analysis. Furthermore, the total healthcare expenditures, which consisted of the sum of all types of healthcare expenditures, were estimated. All healthcare expenditures were adjusted using the consumer price index and were expressed in 2016 constant dollars as provided by the US Bureau of Labor Statistics [21].
Key Independent Variable

The primary independent variable was the presentation of migraine, which included four mutually exclusive groups (migraine alone, migraine and anxiety, migraine and depression, migraine with both conditions).

Other Independent Variables

The other independent variables that were considered were the sociodemographic characteristics, which included gender, age in years (22-39, 40-49, and 50-64), race/ethnicity (White, African American, Latino and other), marital status (married, widowed, separated/divorced, and never married), education (less than high school, high school, and above high school), region of residence (Northeast, Midwest, South, and West), employment (unemployed, employed), health insurance (public, private, and uninsured), prescription medication insurance (insured, uninsured), and poverty status. Other independent variables included personal health practices such as physical activity (3 times/week, no exercise), smoking (smoker, non-smoker), perceived physical health (excellent/very good, good, and fair/poor) and comorbid chronic health conditions such as diabetes, hypertension, and hyperlipidemia.

Statistical analysis

Descriptive statistics were used to describe the study sample across the four migraine groups. One-way analysis of variance (ANOVA) was used to compare the unadjusted means of the total healthcare expenditures across the four migraine groups. Generalized linear model (GLM) regressions with log link were used to estimate the healthcare expenditures associated with comorbid anxiety and depression among individuals with migraine after adjusting for the confounder (sociodemographic characteristics, health insurance status, and personal health practices, perceived physical health, and comorbid chronic health conditions). The
GLM is an attractive alternative to OLS regressions on log-transformed expenditures because it corrects for heteroscedasticity and avoids the retransformation bias [22]. We used GLM with log link and gamma family distribution to estimate the adjusted medical expenditures associated with comorbid anxiety and depression. We also used a two-part model to estimate inpatient expenditures as the majority of the adults in the sample had zero inpatient expenditures. The first-part of the model estimates the probability of having zero expenditure versus positive expenditures. The second part of the model uses GLM to estimate the expenditures conditional on having positive inpatient expenditures. A P value of < 0.05 was considered statistically significant. Primary sampling unit, strata, and weights provided in the MEPS were all used in the analysis. All analyses were performed using survey procedures in the Statistical Analysis System SAS 9.4 (SAS Institute Inc., Cary, NC, USA) and the Software for Statistics and Data Science §TATA 15.1).

Results

Description of the Study Sample

The characteristics of the study sample who met the inclusion criteria (n = 1,556) are shown in Table 1. The vast majority of patients were women (83.0%), white (74.3%), and belong to the 22–39 years age group (40.3%). Approximately, 16% of the study sample had depression, 12.3% had an anxiety disorder, and 13.9% had both conditions. Sociodemographic and economic factors, personal health practices, and coexisting chronic conditions were significantly different across the four migraine groups (P < 0.05). For example, the percentage of patients who reported employment among patients with migraine alone (64.2%) is significantly higher than their counterparts from other migraine groups, which were 14.3%, 11.6%, and 9.9%
among patients with comorbid depression, comorbid anxiety, and comorbid depression and anxiety, respectively.

Healthcare expenditures by migraine groups

The differences in healthcare expenditures associated with migraine between adults with migraine alone and those with comorbid anxiety and/or depression are shown in Table 2. The mean total healthcare expenditures (inpatient, outpatient, prescription drugs, emergency visits, and other expenditures) among patients with migraine alone were significantly lower than their counterparts with comorbid anxiety and/or depression. Similarly, the mean inpatient, outpatient, prescription, and emergency healthcare expenditures were significantly lower among patients with migraine alone compared to other migraine groups. For example, total healthcare expenditures for individuals with migraine alone were $6,461, which were significantly lower than the total healthcare expenditures for individuals with migraine and either comorbid anxiety ($11,102) or depression ($10,817), or with both comorbid anxiety and depression ($14,577).

Adjusted healthcare expenditures by migraine groups

The adjusted differences in healthcare expenditures between patients with migraine alone and those with comorbid anxiety and/or depression are shown in Table 3. Migraine with comorbid anxiety and depression was significantly associated with incremental costs of $1,027 in outpatient’s and $662 emergency room healthcare expenditures, compared to the migraine alone group after controlling for a multitude of covariates. Besides, migraine with comorbid anxiety and/or depression was significantly associated with incremental costs of prescription drug, compared to the migraine alone group after controlling for a multitude of covariates. However,
migraine with comorbid depression and/or anxiety was not associated with significant incremental inpatient, or other healthcare costs. Moreover, the incremental total healthcare costs were not significant for migraine patients with comorbid depression and/or anxiety.

Discussion

The present study compared healthcare expenditures among subjects with migraine when diagnosed alone or with comorbid anxiety and/or depression. The study findings indicate that around half of adults with migraine had anxiety and/or depression, this is consistent with the findings of previously published studies [16]. The high prevalence of depression and anxiety among adults with migraine and the significantly higher healthcare-related costs associated with it compared to the general population emphasizes the need for incorporating depression and anxiety disorders screening into the routine clinical practice of managing migraine among adults. Adults with migraine who were diagnosed with comorbid anxiety and/or depression incurred two to three times higher total healthcare expenditures compared to those with migraine alone. This is consistent with Pesa et al. study which found that the presence of depression and/or anxiety along with migraine resulted in significantly higher total direct medical costs ($5,589 migraine alone vs $10,223 with anxiety and $10,582 with depression) [17]. Further, Wu et al. reported a higher mean annual total health expenses for those with depression ($4,740 migraine alone vs. $10,012 with depression, \( P<.001 \)) [20].

What distinguishes the current investigation from the previously published studies is the fact that the healthcare expenditures for anxiety and anxiety with depression comorbidity were identified after adjusting for a myriad of variables [17, 20].
adjusted analysis of healthcare expenditures found significant differences between migraine groups with respect to outpatient healthcare expenditures and prescription drugs only. Migraine with comorbid anxiety and depression was significantly associated with incremental costs of $1,027 in outpatient and $662 emergency room healthcare expenditures, and $1,128 prescription drug, compared to the migraine alone group. The positive association between comorbid anxiety and depression and healthcare expenditures among adults with migraine has many potential explanations. For example, depression and/or anxiety may exacerbate migraine symptoms and increase the rate of healthcare utilization resulting in higher overall healthcare expenditures. Multiple observational studies with nationally representative samples among adult patients with migraine have reported that the comorbidity of migraine and depression was associated with higher rates of emergency department visits and all-cause office visits compared to those without depression [15, 20]. Besides, it is well-established that the comorbidity of migraine with depression and/or anxiety is associated with a higher rate of prescription drugs utilization as reported by SMILE that was conducted among migraine patients visiting primary care clinics in France [23]. Effective management of depressive and anxiety disorder among patients with migraine may reduce the rate of healthcare utilization and expenditures and improve patient quality of life. Therefore, future studies should examine the economic impact of proper depression and/or anxiety management among patients with migraine and whether this will result in lower overall healthcare expenditures.

It has to be noted, however, that migraine with comorbid depression and/or anxiety was associated with an increase in the total healthcare expenditures, inpatient and other expenditures resulting in higher total healthcare expenditures compared to
the migraine alone group. This association did not reach a statistically significant level in the adjusted analysis, where no significant difference in total healthcare expenditures, inpatient, and other expenditures was observed between the migraine groups. One unexpected finding, was the lower inpatient costs among patients with depression comorbidity, although depression comorbidity is associated with higher rate of hospitalization and emergency room visits compared to migraine patients without depression [15, 20]. These unexpected findings warrant further examination in future studies.

Several practical implications can emanate from the current study. The higher outpatient expenditures among migraine patients with comorbid depression and/or anxiety highlight the need to improve the quality of care provided to this segment of the patient population to improve their adherence to prescribed drug regimens which will eventually lower the rate of emergency room visits and hospitalization resulting in lower overall healthcare costs. Besides, regular depression and anxiety screening for patients with migraine may also reduce the healthcare expenditures associated with migraine and improve the quality of care through early detection and management of these conditions.

Although this study has multiple merits, such as the use of a nationally representative data, the control for a multiple medical and socioeconomic factors, and the robustness of the findings using different statistical models, it should be interpreted in the context of some limitations. These limitations such as the lack of control for the severity of illness and duration of illness should be acknowledged.

Conclusion

The current study has provided an updated estimate of the healthcare expenditures
associated with migraine management with or without comorbid depression and/or anxiety among adult U.S. population. Its findings suggest that healthcare providers should be aware of the high prevalence of anxiety and depression among migraine patients that necessitates early detection and management of these comorbid conditions alongside migraine given the higher outpatient healthcare expenditures associated with these comorbid conditions.

Abbreviations

WHO World Health Organization, YLDs Years Lived with Disability, HRQoL Health-Related Quality of Life, MEPS Medical Expenditure Panel Survey, AHRQ Agency for Healthcare Research and Quality, ICD-9-CM The International Classification of Diseases, Ninth Revision, Clinical Modification, ANOVA Analysis of variance, GLM Generalized linear model, SAS Statistical Analysis System, STATA Software for Statistics and Data Science.

Declarations

Ethics approval and consent to participate

A secondary database “the Medical Expenditure Panel Survey (MEPS) database” a publically available database was used, therefore no ethical approval or consent to participate is required.

Consent for publication

Not applicable.

Availability of data and materials

The dataset supporting the conclusions of this article is available from the Medical Expenditure Panel Survey (MEPS) database, and openly made available for
researchers at the following website: https://meps.ahrq.gov/data_stats/download_data_files.jsp.

Competing interests

The authors declare that there is no conflict of interests regarding the publication of this paper.

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Authors’ Contributions

All authors contributed to the study conception and design. Data analysis was performed by Monira Alwhaibi and Abdulkarim M. Meraya. The first draft of the manuscript was written by Monira Alwhaibi and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript.

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Tables

Table 1
Characteristics of the Study Sample and Number and Row Percentage of Characteristics by Migraine Group among Medical Expenditure Panel Survey data of 2012, 2014, 2016

|                  | Total Sample | Migraine Alone | Migraine & Depression | Migraine & Anxiety |
|------------------|--------------|----------------|-----------------------|-------------------|
|                  | N  | Wt.% | N  | Wt.% | N  | Wt.% | N  | Wt.% |
| All              | 1,556 | 100 | 912 | 57.7 | 243 | 16.1 | 185 | 12.3 |
| Age in years     |     |     |     |      |     |      |     |      |
| 22-39            | 661 | 40.3 | 419 | 62.0 | 79  | 13.1 | 90  | 13.6 |
| 40-49            | 421 | 26.9 | 240 | 57.7 | 70  | 16.9 | 43  | 9.9  |
| 50-64            | 474 | 32.8 | 253 | 52.5 | 94  | 19.1 | 52  | 12.6 |
| Gender           |     |     |     |      |     |      |     |      |
| Women            | 1,307 | 83.0 | 743 | 56.2 | 205 | 15.9 | 167 | 13   |
| Men              | 249  | 17.0 | 169 | 65.2 | 38  | 17.0 | 18  | 8.9  |
| Race/ethnicity   |     |     |     |      |     |      |     |      |
| White            | 850  | 74.3 | 458 | 55.3 | 141 | 16.8 | 116 | 12.7 |
| African American | 251  | 8.6  | 168 | 65.6 | 43  | 18.6 | 22  | 8.5  |
| Latino           | 357  | 12.1 | 221 | 63   | 47  | 12.4 | 36  | 11.7 |
| others           | 98   | 5.0  | 65  | 68.2 | 12  | 9.1  | 11  | 13.2 |
| Marital status   |     |     |     |      |     |      |     |      |
| Married          | 782  | 56.8 | 505 | 62.4 | 105 | 14.7 | 80  | 10.0 |
| Widow            | 229  | 13.1 | 106 | 40.3 | 49  | 21.2 | 30  | 15.7 |
| Sep/Div          | 387  | 21.2 | 204 | 52.3 | 61  | 17.1 | 58  | 17.2 |
| Education level | Never married | 158 | 8.9 | 97 | 66.7 | 28 | 15.2 | 17 | 9.8 | 1  
|----------------|--------------|-----|-----|----|------|----|------|----|------|---|
| LT HS          |              | 142 | 5.8 | 77 | 50.2 | 37 | 28.1 | 11 | 7.7  | 1  
| HS             |              | 198 | 11.8| 110| 58   | 34 | 16.4 | 23 | 11.5 | 3  
| > HS           |              | 922 | 64.3| 531| 56.7 | 127| 14.8 | 117| 13   | 14|
| Missing        |              | 294 | 18.1| 194| 63.7 | 45 | 16.7 | 34 | 11.7 | 2  
| Region         |              |     |     |    |      |    |      |    |      |   
| Northeast      |              | 268 | 17.6| 156| 58.9 | 34 | 11.6 | 38 | 14.3 | 4  
| Midwest        |              | 367 | 25.0| 199| 53.9 | 70 | 18.8 | 41 | 12   | 5  
| South          |              | 504 | 32.9| 307| 59.8 | 76 | 17.1 | 55 | 10.5 | 6  
| West           |              | 417 | 24.5| 250| 58.0 | 63 | 15.1 | 51 | 13.3 | 5  
| Employment     |              |     |     |    |      |    |      |    |      |   
| Employed       |              | 1,061| 73.9| 708| 64.2 | 134| 14.3 | 125| 11.6 | 9  
| Not employed   |              | 495 | 26.1| 204| 39.4 | 109| 21   | 60 | 14.3 | 12 |
| Poverty status†|              |     |     |    |      |    |      |    |      |   
| Poor           |              | 327 | 14.2| 141| 38.6 | 69 | 20.9 | 43 | 15.9 | 7  
| Near Poor      |              | 317 | 15.1| 184| 53.1 | 48 | 15.8 | 38 | 14.6 | 4  
| Middle         |              | 440 | 27.8| 273| 57.2 | 59 | 16.8 | 53 | 12.1 | 5  
| High Income    |              | 472 | 42.9| 314| 66.1 | 67 | 14.1 | 51 | 10.3 | 4  
| Health Insurance|             |     |     |    |      |    |      |    |      |   
| Private        |              | 1,002| 75.0| 643| 62.6 | 138| 15.0 | 121| 11.8 | 10 |
| Public         |              | 416 | 18.1| 182| 38.9 | 85 | 20.2 | 55 | 16.2 | 9  
| Uninsured      |              | 138 | 6.8 | 87 | 54.5 | 20 | 17.2 | 9  | 6.8  | 2  
| Rx Insurance   |              |     |     |    |      |    |      |    |      |   
| Rx insurance   |              | 884 | 68.1| 569| 62.8 | 115| 14.5 | 114| 12.2 | 8  
| No Rx insurance|              | 672 | 31.9| 343| 47.0 | 128| 19.5 | 71 | 12.3 | 13|
| General health |              |     |     |    |      |    |      |    |      |   
| Ex/very good   |              | 604 | 43.0| 454| 75.2 | 60 | 10.7 | 54 | 8.2  | 3  
| Good           |              | 519 | 32.2| 298| 53.4 | 89 | 17.9 | 66 | 14.6 | 6  
| Fair/poor      |              | 433 | 24.8| 160| 33.2 | 94 | 23   | 65 | 16.3 | 11 |
|                                | Yes       | No       | Total | Mean | SD    |
|--------------------------------|-----------|----------|-------|------|-------|
| **Physical activity**          |           |          |       |      |       |
| 3/week                         | 657       | 895      | 1,552 | 42.9 | 6.8   |
| No                             | 895       | 1,189    | 2,084 | 56.8 | 5.7   |
| **Smoking**                    |           |          |       |      |       |
| Current Smoker                 | 271       | 1,189    | 1,460 | 16.3 | 3.3   |
| Others                         | 895       | 1,189    | 2,084 | 56.8 | 5.7   |
| **Heart**                      |           |          |       |      |       |
| Yes                            | 154       | 1,402    | 1,556 | 8.9  | 2.8   |
| No                             | 1,402     | 1,189    | 2,591 | 91.1 | 6.9   |
| **Hypertension**               |           |          |       |      |       |
| Yes                            | 393       | 1,163    | 1,556 | 24.0 | 3.2   |
| No                             | 1,163     | 775      | 1,938 | 76.0 | 4.5   |
| **Diabetes**                   |           |          |       |      |       |
| Yes                            | 128       | 1,428    | 1,556 | 7.2  | 1.9   |
| No                             | 1,428     | 775      | 2,203 | 92.8 | 4.7   |
| **Hyperlipidemia**             |           |          |       |      |       |
| Yes                            | 781       | 775      | 1,556 | 48.3 | 3.1   |
| No                             | 775       | 1,016    | 1,791 | 51.7 | 3.4   |
| **Asthma**                     |           |          |       |      |       |
| Yes                            | 240       | 1,316    | 1,556 | 14.8 | 2.6   |
| No                             | 1,316     | 1,016    | 2,332 | 85.2 | 2.5   |
| **COPD**                       |           |          |       |      |       |
| Yes                            | 255       | 1,301    | 1,556 | 16.3 | 2.8   |
| No                             | 1,301     | 1,016    | 2,317 | 83.7 | 2.5   |
| **Arthritis**                  |           |          |       |      |       |
| Yes                            | 540       | 1,016    | 1,556 | 34.8 | 4.2   |
| No                             | 1,016     | 694      | 1,710 | 65.2 | 4.2   |
| **GERD**                       |           |          |       |      |       |
| Yes                            | 204       | 1,352    | 1,556 | 13.5 | 2.1   |
| No                             | 1,352     | 828      | 2,180 | 86.5 | 2.9   |
| **Cancer**                     |           |          |       |      |       |
| Yes                            | 96        | 1,460    | 1,556 | 7.2  | 1.8   |
| No                             | 1,460     | 877      | 2,337 | 92.8 | 2.9   |

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Note: Based on 1,556 adults with migraine headache, aged ≥22 years who were alive during the calendar years 2012–2016.

Missing indicators for education, smoking were used but not presented in the table.
† Poor (< 100% federal poverty line), near poor (100% to < 200%), middle income (200% to < 400%), and high income (≥ 400%).
COPD: Chronic obstructive pulmonary disease; GERD: Gastroesophageal reflux disease; HS: High School; Widow/Sep/Div: Widow/separated/divorced.

Table 2
Mean Healthcare Expenditures by Migraine Groups among Adults with Migraine
Medical Expenditure Panel Survey data of 2012, 2014, and 2016

| Expenditures          | Migraine Alone | Migraine & Depression | Migraine & Anxiety | Migraine & Anxiety |
|-----------------------|----------------|-----------------------|--------------------|--------------------|
|                       | Mean ($)       | SE                    | Mean ($)           | SE                 | Mean ($)           | SE                 |
| Total                 | 6,461          | 634                   | 11,102             | 1446               | 10,817             | 1382               |
| Inpatient             | 1,391          | 353                   | 1,250              | 421                | 1,845              | 592                |
| Outpatient            | 2,360          | 207                   | 4,836              | 1216               | 4,374              | 781                |
| Prescription          | 1,619          | 224                   | 3,922              | 542                | 3,234              | 557                |
| Emergency Room        | 439            | 66                    | 535                | 192                | 578                | 122                |
| Other                 | 652            | 86                    | 785                | 83                 | 710                |

Note: Based on 1,556 adults with migraine headache, aged 22 years and older and were alive during the calendar years 2012–2016.
p-value represent significant mean differences by migraine groups using Anova.
Other expenditures included dental, vision, durable medical equipment use, and others.
SE: Standard Error
Table 3

Intercept and parameter estimates for the migraine groups from generalized linear models with log link and gamma distribution among Adults with Migraine, Medical Expenditure Panel Survey

|                          | Migraine & Depression | Migraine & Anxiety |                  |                  |                  |                  |                  |
|--------------------------|-----------------------|--------------------|------------------|------------------|------------------|------------------|------------------|
|                          | Intercept  (SE)        | β                  | SE               | Incremental cost | p-value          | β               | SE               | Incremental cost | p-value          |
| Unadjusted Generalized Linear Model with Log Link and Gamma Distribution |                       |                    |                  |                  |                  |                  |                  |
| Total                    | 8.77 (0.09)           | 0.54               | 0.15             | $4,641           | 0.002            | 0.51            | 0.16             | $4,356           |
| Inpatient                | 9.78 (0.78)           | -0.28              | 0.42             | -$141            | 0.799            | 0.04            | 0.33             | $454             |
| Outpatient               | 7.76 (0.08)           | 0.71               | 0.21             | $2,476           | 0.026            | 0.62            | 0.20             | $2,014           |
| Prescription             | 7.38 (0.14)           | 0.88               | 0.19             | $2,302           | <0.000           | 0.69            | 0.23             | $1,616           |
| Emergency                | 6.08 (0.15)           | 0.27               | 0.27             | $95              | 0.639            | 0.27            | 0.27             | $140             |
| Other                    | 6.47 (0.13)           | -0.15              | 0.19             | -$91             | 0.432            | 0.18            | 0.40             | $133             |
| Adjusted Generalized Linear Model with Log Link and Gamma Distribution |                       |                    |                  |                  |                  |                  |                  |
| Total                    | 10.61 (0.41)          | 0.11               | 0.11             | 1025             | 0.330            | 0.09            | 0.12             | $871             |
| Inpatient                | 10.8(0.68)            | -0.48              | 0.27             | -               | 0.029            | -0.1            | 0.34             | -$707            |
| Outpatient               | 8.46 (0.43)           | 0.15               | 0.12             | $1,332           | 0.224            | 0.23            | 0.14             | $816             |
| Prescription             | 8.75 (0.46)           | 0.43               | 0.14             | $1,296           | 0.005            | 0.52            | 0.16             | $1,598           |
| Emergency                | 7.89 (0.92)           | 0.37               | 0.29             | $213             | 0.281            | 0.68            | 0.36             | $460             |
| Other                    | 6.11 (0.58)           | -0.16              | 0.16             | -$109            | 0.307            | -0.32           | 0.18             | -$197            |

Note: Based on 1,556 adults with migraine headache, aged 22 years and older and were alive during the calendar year. p-values denote statistical significance in parameter estimates from generalized linear models on expenditures with log link. Other expenditures included dental, vision, durable medical equipment use, and others. Reference group for migraine groups= Migraine alone

B: Coefficient; Substandard Error