Insomnia as an independent predictor of suicide attempts: a nationwide population-based retrospective cohort study

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Abstract

Background: Numerous studies have verified that insomnia is associated with suicidal ideation, suicide attempts, and death by suicide. Limited population-based cohort studies have been conducted to examine the association. The present study aimed to analyze whether insomnia increases the risk of suicide attempts and verify the effects of insomnia on suicide risk.

Methods: This study is a cohort study using 2000–2013 hospitalization data from the National Health Insurance Research Database (NHIRD) to track the rate of suicide attempts among insomnia patients aged 15 years or older. In addition, a 1:2 pairing based on sex, age, and date of hospitalization was conducted to identify the reference cohort (patients without insomnia). Cox proportional hazard model was used to assess the effects of insomnia on suicide risk.

Results: The total number of hospitalized patients aged 15 years or older was 479,967 between 2000 and 2013 (159,989 patients with insomnia and 319,978 patients without insomnia). After adjusting for confounders, suicide risk in insomnia patients was 3.533-fold that of patients without insomnia (adjusted hazard ratio [HR] = 3.533, 95% confidence interval [CI] = 3.059–4.080, P < 0.001). Suicide risk in low-income patients was 1.434-fold (adjusted HR = 1.434, 95% CI = 1.184–1.736, P < 0.001) that of non-low-income patients. Suicide risk in patients with drug dependence and with mental disorders was 1.592-fold (adjusted HR = 1.592, 95% CI = 1.220–2.077, P < 0.001) and 4.483-fold (adjusted HR = 4.483, 95% CI = 3.934–5.109, P < 0.001) that of patients without drug dependence and without mental disorders, respectively. In the female population, suicide risk in insomnia patients was 4.186-fold (adjusted HR = 4.186, 95% CI = 3.429–5.111, P < 0.001) that of patients without insomnia. Among patients aged 25–44 years, suicide risk in insomnia patients was 5.546-fold (adjusted HR = 5.546, 95% CI = 4.236–7.626, P < 0.001) that of patients without insomnia. Furthermore, the suicide risk of insomnia patients with mental disorders was 18.322-fold that of patients without insomnia and mental disorders (P < 0.001).

Conclusion: Insomnia, low income, drug dependence, and mental disorders are independent risk factors for suicide attempts. Female patients and those aged 25–44 years are at high risk of suicide due to insomnia. Insomnia, mental disorders, and low income exhibit a synergistic effect on suicide attempts. Clinicians should pay attention to mental status and income level of insomnia patients.

Keywords: Insomnia, Suicide, National Health Insurance Research Database (NHIRD)

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Background
Suicide is a serious public health problem around the world. The World Health Organization (WHO) reported that approximately 800,000 people worldwide die from suicide every year (an average of one death every 40 s). In addition, suicide is the second leading cause of death among those aged 15–29 years globally, and the fifth leading cause of death among those aged 30–49 years [1]. Compared with death by suicide, there are many more suicide attempts every year [2], and according to a meta-analysis from Japan in 2008, a prior suicide attempt is the most important predictor of suicide [3]. Suicide, moreover, causes immense socioeconomic burdens on families, communities, and nations [4]. In 2016, the number of deaths from suicide in Taiwan was 3765 (a suicide death rate of 16.0 persons per 100,000 population), making suicide the twelfth leading cause of death in Taiwan. The suicide death rate among men was approximately 2.1-times higher than that among women (17.0 men and 7.7 women per 100,000 population). The suicide death rate in all age groups increased with age [5].

Insomnia is one of the most prevalent sleeping disorders in the world [6]. According to epidemiological studies around the world, the prevalence of insomnia in general populations is 10%–25% [7–10]. A 2017 report by the Taiwan Society of Sleep Medicine indicated that a tenth of the population across Taiwan suffers from chronic insomnia (prevalence rate of 11.3%) [11]. Insomnia refers to difficulty falling asleep, remaining awake while trying to sleep, waking up often during the night, or still feeling tired after sleeping for a brief period, factors that subsequently influence daytime activities of daily living for more than four weeks. If insomnia persists for more than six months, it becomes a chronic condition influencing not only the person's psychology and physiology but also his or her health and activities of daily living (e.g., learning and working) [12].

Numerous studies have indicated that insomnia is associated with suicidal ideation [13–16], suicide attempts [17–19], and death by suicide [20–22] among adolescents and adults. These studies laid the foundation for the relationship between insomnia and suicide. However, there are some weakness in methodology and need future research to fill this gap. For example, some studies used a questionnaire asking a single question as criteria for determining insomnia or suicide [22]. Furthermore, mental disorders are a major confounding factor for insomnia and suicide, and over 90% of suicide decedents have at least one mental disorder [23]; however, some studies assessing the link between insomnia and suicide did not adjust for mental disorders [17, 20].

There are still different arguments about whether insomnia is an independent factor of suicide or if insomnia, as a symptom of mental disorders such as depression, increases the risk of suicide. Some studies have found that the relationship between insomnia and suicide is fully mediated by mental disorders [13, 14, 24]. Other studies have shown that insomnia remains an independent factor of suicidal ideation [15, 16], suicide attempt [19], and death by suicide [21, 22] after adjustments for mental disorders, substance abuse and alcohol abuse.

A meta-analysis by Pigeon et al. indicated that insomnia is still a predictor of suicide even after other factors have been adjusted for. However, the authors asserted that the samples collected are non-homogeneous, and samples for observational and clinical studies are confined to adolescents or adults only. Additionally, numerous studies are cross-sectional studies [25], which cannot be used to elucidate whether insomnia increases suicide risk. Hence, we employed the National Health Insurance Research Database (NHIRD) in Taiwan to conduct a retrospective cohort study of whether insomnia increases the risk of suicide attempts.

Methods
Data source
Introduced in Taiwan in 1995, the NHIRD contains the medical records of all insured. The database encompasses the information of more than 99% of the 23 million people who live in Taiwan; thus, the health care information contained therein represents evidence-based data in the medicine and healthcare sector [26]. This study analyzed the 2000–2013 inpatient expenditure files and the registry for contracted medical facilities in the NHIRD.

Design and sample
The retrospective cohort study design method was adopted in this study. Inpatients aged 15 years or older who were newly diagnosed with insomnia (International Classification of Diseases, 9th Revision, Clinical Modifications [ICD-9-CM]: 307.41, 307.42, 780.52) were allocated to the study cohort. To ensure that the patients were newly diagnosed, we also excluded patients who were diagnosed with insomnia between 1997 and 1999. The study period began on January 1, 2000 and ended on December 31, 2013. In total, 159,989 participants were included. The average of follow-up for the diagnosis of insomnia was 6.47 years. To facilitate comparison of the study cohort, propensity score-matching based on sex, age group, and date of hospitalization was performed for a 1:2 pairing to identify the reference cohort (n = 319,978 patients who were diagnosed as not having insomnia). Furthermore, there was no difference in the cause of hospitalization at baseline (Additional file 1: Table S1).

Outcome assessment
All research participants were observed until the incidence of suicide attempts (ICD-9-CM E code: 950–959), loss to follow up, or until the end of December 31, 2013.
Covariates of interest
The participants in this study were allocated into four groups by age: 15–24, 25–44, 45–64, and ≥65 years. Urbanization levels were categorized into high, moderate, and low levels of urbanization [27]. Low income included insured individuals in Category 5 (those to which laws and regulations governing social relief apply). Catastrophic illness was indicated in the copayment remarks of inpatient expenditure files. The Charlson comorbidity index (CCI) was used to weigh the 19 types of diseases by assigning each with a score of 1, 2, or 6, and the scores were summed according to whether each patient had any of these diseases [28, 29]. Because drug dependence, alcohol dependence, and mental disorders are risk factors for suicide attempts, drug dependence (ICD-9-CM 292, 304), alcohol dependence (ICD-9-CM 291, 303), and mental disorders (ICD-9-CM 290–319, excluding 291, 292, 303, 304) were incorporated in the regression model for adjustments.

Statistical analysis
All analyses were performed using SPSS 22 (SPSS, Inc., Chicago, IL, U.S.). The χ² test and Fisher exact test were used to compare the categorical variables of the two groups, and a t-test was conducted to compare the continuous variables of the two groups. With adjustments for sex, age, low income, catastrophic illness, urbanization, CCI, drug dependence, alcohol dependence, and mental disorders, Cox proportional hazards regression analyses were performed to assess the effects of insomnia on the risk of suicide attempts and presented as hazard ratio (HR) with a 95% confidence interval (CI). Furthermore, differences in the risk of suicide attempts between the study and reference cohort were estimated using the Kaplan-Meier method with the log-rank test. A 2-tailed p-value < 0.05 was considered to indicate statistical significance.

Results
Figure 1 shows the case-screening process (inclusion and exclusion) and follow-up results as well as the cumulative risk of suicide incidence between the two groups (patients with/without insomnia). Between 2000 and 2013, 16,716,547 patients were hospitalized, 178,589 of which had insomnia. After exclusion criteria were applied (18,600 excluded), there were a total of 159,989 insomnia
cases (319,978 patients were placed in the reference cohort based on 1:2 pairing) with a subsequent suicide incidence of 0.69% (1098/159,989), whereas the reference cohort exhibited a suicide incidence of 0.09% (297/319,978). The groups demonstrated significant difference (log-rank $P < 0.001$). In other words, the probability (risk) of suicide attempts in insomnia patients was considerably higher than that in patients without insomnia. In addition, both groups exhibited a statistically significant difference after one year follow up (Fig. 2, Table 1). Furthermore, an average follow-up period for the diagnosis of insomnia to suicide attempts was 2.38 years.

Table 2 presents the basic characteristics of the 479,967 patients at the endpoint of the follow-up process (study cohort = 159,989 insomnia patients; reference cohort = 319,978 non-insomnia patients). The study cohort exhibited a substantially higher incidence of suicide attempts compared with the reference cohort (0.7% vs 0.1%; $P < 0.001$). The study cohort comprised significantly higher numbers of low income patients (4.0% vs 1.5%; $P < 0.001$), patients with catastrophic illness (31.2% vs 19.8%; $P < 0.001$), and those who lived in working class urbanized townships (33.4% vs 22.3%; $P < 0.001$) than the reference cohort. The numbers of patients with drug dependence (1.0% vs 0.1%; $P < 0.001$), alcohol dependence (3.5% vs 0.7%; $P < 0.001$), and mental disorders (33.5% vs 7.1%; $P < 0.001$) and other comorbidities in the study cohort were significantly higher than those of the reference cohort.

Table 3 shows the univariate and multivariate analysis results of the factors of suicide attempts. After the research variables (sex, age, low income, catastrophic illness, urbanization, CCI, drug dependence, alcohol dependence, and mental disorders) were adjusted for, the risk of suicide attempts among insomnia patients was 3.533-fold that of non-insomnia patients ($P < 0.001$). The risk of suicide attempts among female patients was 1.522-fold ($P < 0.001$) that of male patients. The risk of suicide attempts among patients aged 15–24, 25–44, 45–64 years was 6.000-fold, 3.581-fold, and 1.595-fold ($P < 0.001$) that of patients aged 65 years or older, respectively. The risk of suicide attempts among low-income patients and patients with catastrophic
Table 2 Characteristics of study in the endpoint

| Variables                  | Total          | With insomnia | Without insomnia | P-value |
|----------------------------|----------------|---------------|------------------|---------|
| Total                      | 479,967 (100.0)| 159,989 (33.3)| 319,978 (66.7)   |         |
| Suicide attempts           |                |               |                  | < 0.001|
| with                       | 1395 (0.3)     | 1098 (0.7)    | 297 (0.1)        |         |
| without                    | 478,572 (99.7)| 158,891 (99.3)| 319,681 (99.9)  |         |
| Gender                     |                |               |                  | 0.880   |
| male                       | 235,771 (49.1)| 78,615 (49.1) | 157,156 (49.1)  |         |
| female                     | 244,196 (50.9)| 81,374 (50.9) | 162,822 (50.9)  |         |
| Age group (years)          |                |               |                  | 0.958   |
| 15–24                      | 15,901 (3.3)   | 5330 (3.3)    | 10,571 (3.3)    |         |
| 25–44                      | 99,058 (20.6)  | 33,030 (20.6) | 66,028 (20.6)   |         |
| 45–64                      | 172,650 (36.0)| 57,504 (35.9)| 115,146 (36.0)  |         |
| ≥65                        | 192,358 (40.1)| 64,125 (40.1)| 128,233 (40.1)  |         |
| Low income                 |                |               |                  | < 0.001|
| yes                        | 11,251 (2.3)   | 6441 (4.0)    | 4810 (1.5)      |         |
| no                         | 468,716 (97.7)| 153,548 (96.0)| 315,168 (98.5) |         |
| Catastrophic illness       |                |               |                  | < 0.001|
| yes                        | 113,150 (23.6)| 49,914 (31.2)| 63,236 (19.8)   |         |
| no                         | 366,817 (76.4)| 110,075 (68.8)| 256,742 (80.2) |         |
| Urbanization               |                |               |                  | < 0.001|
| high                       | 150,124 (31.3)| 41,562 (26.0)| 108,562 (33.9)  |         |
| medium                     | 205,154 (42.7)| 65,049 (40.7)| 140,105 (43.8)  |         |
| low                        | 124,689 (26.0)| 53,378 (33.4)| 71,311 (22.3)   |         |
| CCI                        | 2.36 ± 3.82    | 2.97 ± 4.10   | 2.06 ± 3.63     | < 0.001|
| Drug dependence            |                |               |                  | < 0.001|
| yes                        | 2051 (0.4)     | 1597 (1.0)    | 454 (0.1)       |         |
| no                         | 477,916 (99.6)| 158,392 (99.0)| 319,524 (99.9) |         |
| Alcohol dependence         |                |               |                  | < 0.001|
| yes                        | 7846 (1.6)     | 5674 (3.5)    | 2172 (0.7)      |         |
| no                         | 472,121 (98.4)| 154,315 (96.5)| 317,806 (99.3) |         |
| Mental disorders           |                |               |                  | < 0.001|
| yes                        | 76,322 (15.9)| 53,537 (33.5)| 22,785 (7.1)    |         |
| no                         | 403,645 (84.1)| 106,452 (66.5)| 297,193 (92.9) |         |

P-value (category variable: Chi-square/Fisher exact test; continue variable: t-test)

illness was 1.434-fold and 1.286-fold that of their counterparts, respectively ($P < 0.001$). The risk of suicide attempts among patients with drug dependence and mental disorders was 1.592-fold and 4.483-fold that of their counterparts, respectively ($P < 0.001$).

Table 4 shows the hierarchical analysis of various variables to elucidate the difference in risk of suicide attempts between insomnia and non-insomnia patients. The results indicated that (after other factors were adjusted for), in the female population, suicide risk in insomnia patients was 4.186-fold (adjusted HR = 4.186, 95% CI = 3.429–5.111, $P < 0.001$) that of patients without insomnia. Among patients aged 25–44 years, suicide risk in insomnia patients was 5.546-fold (adjusted HR = 5.546, 95% CI = 4.236–7.262, $P < 0.001$) that of patients without insomnia.

Figures 3 and 4 show the interactive effects of insomnia, mental disorders, and low income after other factors were adjusted for. The risk of suicide attempts in insomnia patients without mental disorders was 4.960-fold that of patients without insomnia and mental disorders.
suicide attempts in non-insomnia patients with mental disorders was 8.237-fold that of patients without insomnia and mental disorders. The risk of suicide attempts in insomnia patients with mental disorders was 18.322-fold that of patients without insomnia and mental disorders (P < 0.001) (Fig. 3). The risk of suicide attempts in non-low-income patients with insomnia was 3.521-fold that of non-insomnia patients who were not in the low-income group. The risk of suicide attempts in low-income patients without insomnia was 1.330-fold that of non-insomnia patients who were not in the low-income group. The risk of suicide attempts in low-income patients with insomnia was 5.084-fold that of non-insomnia patients who were not in the low-income group (P < 0.001) (Fig. 4).

**Table 3** Factors of suicide attempts by Cox proportional hazard model analysis

| Variables           | Crude HR | 95% CI     | P-value | Adjusted HR | 95% CI     | P-value |
|---------------------|----------|------------|---------|-------------|------------|---------|
| Insomnia            |          |            |         |             |            |         |
| no                  | reference| reference  |         |             |            |         |
| yes                 | 7.416    | 6.524      | 8.431   | < 0.001     | 3.533      | 3.059   | 4.080   | < 0.001 |
| Gender              |          |            |         |             |            |         |
| female              | reference| reference  |         |             |            |         |
| male                | 1.289    | 1.159      | 1.433   | < 0.001     | 1.522      | 1.361   | 1.702   | < 0.001 |
| Age group (years)   |          |            |         |             |            |         |
| ≧65                 | reference| reference  |         |             |            |         |
| 15–24               | 6.477    | 5.292      | 7.928   | 0.001       | 6.000      | 4.876   | 7.383   | < 0.001 |
| 25–44               | 4.909    | 4.238      | 5.687   | < 0.001     | 3.581      | 3.069   | 4.177   | < 0.001 |
| 45–64               | 1.747    | 1.487      | 2.052   | < 0.001     | 1.595      | 1.357   | 1.876   | < 0.001 |
| Low income          |          |            |         |             |            |         |
| no                  | reference| reference  |         |             |            |         |
| yes                 | 3.940    | 3.267      | 4.751   | < 0.001     | 1.434      | 1.184   | 1.736   | < 0.001 |
| Catastrophic illness|          |            |         |             |            |         |
| No                  | reference| reference  |         |             |            |         |
| Yes                 | 1.972    | 1.771      | 2.196   | < 0.001     | 1.286      | 1.143   | 1.446   | < 0.001 |
| Urbanization        |          |            |         |             |            |         |
| high                | reference| reference  |         |             |            |         |
| medium              | 1.045    | 0.916      | 1.193   | 0.510       | 1.013      | 0.887   | 1.157   | 0.847   |
| low                 | 1.569    | 1.372      | 1.795   | < 0.001     | 1.073      | 0.937   | 1.230   | 0.309   |
| CCI                 | 0.983    | 0.969      | 0.998   | 0.025       | 0.989      | 0.973   | 1.005   | 0.179   |
| Drug dependence     |          |            |         |             |            |         |
| no                  | reference| reference  |         |             |            |         |
| yes                 | 9.614    | 7.407      | 12.479  | < 0.001     | 1.592      | 1.220   | 2.077   | 0.001   |
| Alcohol dependence  |          |            |         |             |            |         |
| no                  | reference| reference  |         |             |            |         |
| yes                 | 5.872    | 4.899      | 7.038   | < 0.001     | 1.164      | 0.955   | 1.420   | 0.133   |
| Mental disorders    |          |            |         |             |            |         |
| no                  | reference| reference  |         |             |            |         |
| yes                 | 9.742    | 8.723      | 10.879  | < 0.001     | 4.483      | 3.934   | 5.109   | < 0.001 |

**Discussion**

Our study found that insomnia remained an independent risk factor for suicide attempts after adjustments for mental disorders, alcohol dependence, and drug dependence. To the best of our knowledge, our study is the first retrospective cohort study to use population-based data and clinical diagnosis as the criteria for determining insomnia and suicide. The findings can reinforce the deficiencies of other relevant studies.

In recent years, some studies found that mental disorders, alcohol abuse, and drug abuse do not fully mediate the association between insomnia and suicide. A study conducted in 2012 regarding members of the military in the United States reported that insomnia was an independent risk
factor for suicidal ideation after adjustments for depression, hopelessness, post-traumatic stress disorder, anxiety, drug abuse, and alcohol abuse. However, there were no significant associations between insomnia and suicide attempts [15]. Different from the above study, our study used the clinical diagnosis as the criteria for determining insomnia. Another study conducted in the United States in 2016 revealed that insomnia symptoms increase the risk of suicidal ideation and attempts in adolescents with adjustments for mental disorders and substance use disorders [19]. However, the above study [19] had several limitations. First, the study was cross-sectional. No temporal relationships could be established. Second, all measures were based on self-report, thus the data were subject to response and recall bias. Therefore, based on above study [19], we established insomnia as an independent predictor of suicide attempts.

Our study found that insomnia and mental disorders have a synergistic effect on the risk of suicide attempts. The risk of suicide attempts in insomnia patients with mental disorders was 18-fold that of patients without insomnia and mental disorders. However, no comparison with past studies can be made because investigations concerning the interactive effects of insomnia and mental disorders on suicide attempts are lacking. A cross-sectional study conducted in 2018 in the United States reported that poor sleep quality will increase the risk of suicide after adjusting for depression among adolescents. However, sleep problems and depression do not interact with the risk of suicide [30]. Different from above study [30] focus on the association between sleep problems and suicide, our study

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**Table 4** Factors of suicide attempts stratified by variables listed in the table by Cox proportional hazard model analysis

| Variables          | With insomnia | Without insomnia | Ratio | Adjusted HR | 95% CI | P-value |
|--------------------|----------------|-------------------|-------|-------------|--------|---------|
|                    | Event PYs | Rate (per 10^3 PYs) | Event PYs | Rate (per 10^3 PYs) |       |         |
| Total              | 1098     | 953,594           | 115.143 | 297          | 1,913,519 | 15.521 | 7.418 | 3.533 | 3.059 | 4.080 | < 0.001 |
| Gender             |           |                   |       |             |        |         |
| female             | 633      | 468,954           | 134.981 | 153         | 941,910 | 16.244 | 8.310 | 4.186 | 3.429 | 5.111 | < 0.001 |
| male               | 465      | 484,640           | 95.948  | 144         | 971,609 | 14.821 | 6.474 | 2.861 | 2.323 | 3.523 | < 0.001 |
| Age group (years)  |           |                   |       |             |        |         |
| 15–24              | 119      | 42,625            | 279.179 | 33          | 85,435  | 38.626 | 7.228 | 2.778 | 1.747 | 4.417 | < 0.001 |
| 25–44              | 546      | 198,523           | 275.031 | 80          | 400,512  | 19.974 | 13.769 | 5.546 | 4.236 | 7.262 | < 0.001 |
| 45–64              | 264      | 317,825           | 83.065  | 105         | 637,466  | 16.471 | 5.043 | 2.637 | 2.053 | 3.389 | < 0.001 |
| ≥65                | 169      | 394,621           | 42.826  | 79          | 790,106  | 9.999  | 4.283 | 2.900 | 2.188 | 3.844 | < 0.001 |
| Low income         |           |                   |       |             |        |         |
| no                 | 987      | 915,626           | 107.795 | 288         | 1,884,404 | 15.283 | 7.053 | 3.487 | 3.009 | 4.040 | < 0.001 |
| yes                | 111      | 37,968            | 292.351 | 9           | 29,115   | 30.912 | 9.458 | 4.069 | 2.012 | 8.228 | < 0.001 |
| Catastrophic illness|           |                   |       |             |        |         |
| no                 | 635      | 645,939           | 98.306  | 220         | 1,504,259 | 14.625 | 6.722 | 3.769 | 3.171 | 4.479 | < 0.001 |
| yes                | 463      | 307,655           | 150.493 | 77          | 409,260  | 18.814 | 7.999 | 2.916 | 2.255 | 3.770 | < 0.001 |
| Urbanization       |           |                   |       |             |        |         |
| high               | 291      | 243,831           | 119.345 | 84          | 661,398  | 12.700 | 9.397 | 3.911 | 2.964 | 5.161 | < 0.001 |
| medium             | 398      | 369,572           | 107.692 | 132         | 835,185  | 15.805 | 6.814 | 3.320 | 2.655 | 4.153 | < 0.001 |
| low                | 409      | 340,191           | 120.227 | 81          | 416,936  | 19.427 | 6.188 | 3.355 | 2.592 | 4.342 | < 0.001 |
| Drug dependence    |           |                   |       |             |        |         |
| no                 | 1044     | 942,877           | 110.725 | 292         | 1,910,198 | 15.286 | 7.243 | 3.569 | 3.086 | 4.127 | < 0.001 |
| yes                | 54       | 10,717            | 503.872 | 5           | 3321     | 150.557 | 3.347 | 2.079 | 0.817 | 5.286 | 0.207 |
| Alcohol dependence |           |                   |       |             |        |         |
| no                 | 978      | 916,596           | 106.699 | 288         | 1,899,319 | 15.163 | 7.037 | 3.513 | 3.032 | 4.072 | < 0.001 |
| yes                | 120      | 36,998            | 324.342 | 9           | 14,200   | 63.380 | 5.117 | 3.844 | 1.936 | 7.634 | < 0.001 |
| Mental disorders   |           |                   |       |             |        |         |
| no                 | 300      | 618,043           | 48.540  | 181         | 1,763,589 | 10.263 | 4.730 | 4.798 | 3.968 | 5.801 | < 0.001 |
| yes                | 798      | 335,551           | 237.818 | 116         | 149,930  | 77.369 | 3.074 | 2.223 | 1.823 | 2.711 | < 0.001 |
focused on insomnia and suicide attempts. Therefore, it cannot be compared.

The mechanisms underlying the relationship between insomnia and suicide remain unclear. In 2013, a systematic literature review of adult insomnia and suicide conducted in the United States indicated that the mechanisms of insomnia and suicide involve both physiological and psychological dimensions [31]. The physiological mechanism includes a reduction in serotonin [32] and hypothalamic-pituitary-adrenal (HPA) axis dysfunction [33]. The psychological mechanism is associated with dysfunctional beliefs and attitudes about sleep (DBAS) [15] and depression [34]. Furthermore, a previous study found that fatigue resulting from sleep disorders may lead to hopelessness and decrease impulse control, both demonstrated risk factors for suicide [35]. The above possible mechanisms between insomnia and suicide require further studies for confirmation.

Our study determined that the risk of suicide attempts in patients with drug dependence was 1.592-fold that of patients without drug dependence. Several studies have indicated that the abuse of drugs such as marijuana [36], heroin [37, 38], and nicotine [39] are risk factors for suicide. A review article reported that 25% to 50% of people who...
are suicidal are dependent on alcohol or drugs, and the risk of suicide increases if these people have mental disorders [40]. A study conducted in 1995 regarding indigenous peoples in Taiwan reported that the risk of suicide in people with substance dependence and depression was 470.2-fold that of people without substance dependence and depression [41]. In our study, we found that the risk of suicide attempts in patients with both substance dependence and depression was 22.7-fold that of their counterparts who did not fact these issues, which is lower than the finding of the aforementioned study. This difference might be attributable to social group, population, and cultural discrepancies. Another study conducted in the United States in 1993 revealed that the risk of suicide in patients with substance dependence and an emotional disorder was 17.0-fold that of their counterparts [42]. By contrast, we determined that the risk of suicide attempts in patients with both substance dependence and emotional disorder was 23.4-fold that of their counterparts. Similar to the Taiwan study [41], the focus of substance abuse in this study was alcohol and drugs; however, we reported a lower rate of substance abuse (alcohol abuse = 9.2% and drug abuse = 4.3%) compared with that of [41] (alcohol abuse = 24.3% and drug abuse = 13.4%). This difference is possibly attributed to the more stringent standard of substance abuse (individuals exhibiting substance dependence) adopted in the present study.

Our study found that the risk of suicide attempts in patients with mental disorders was 4.483-fold that of patients without mental disorders. In western countries such as the United Kingdom, a study reported that 90% of suicides were associated with a history of mental disorders [43]. In Asian countries such as China, 40% to 70% of suicides were associated with a history of mental disorders [44, 45]. Multiple studies have indicated that different types of mental disorders predict different levels of suicide risk [46–49]. In particular, schizophrenia has the highest risk for suicide [50]. In our study, depression had the highest risk for suicide attempts, followed by schizophrenia. Another study revealed that the risk of suicide in patients with more than one mental disorders was 1.5–2.5-fold that of patients with only one type of mental disorders [43]. We incorporated depression, anxiety, bipolar disorder, and schizophrenia for further analysis and found that for every increase in the number of mental disorders, the risk of suicide attempts increased by 0.6-fold. This finding is somewhat similar to that reported in the aforementioned study.

In 2016, a United Kingdom systematic literature review indicated that low income, low socioeconomic status, and unemployment predicted higher levels of suicidal ideation, suicidal behavior, and death by suicide [51]. Our study found that risk of suicide attempts in low-income patients was 1.434-fold that of their counterparts (adjusted HR = 1.434, 95% CI = 1.182–1.732, P < 0.001). Furthermore, the risk of suicide attempts in insomnia patients who earn low levels of income was 4.960-fold that of their counterparts (P < 0.001). This result implies that under the interactive effects of an economic stressor (low income) and insomnia, suicide became a major option for this particular population (particularly women, people aged 25–44 years, and breadwinners in the family), thereby leading to regrettable circumstances.

Our study has the following limitations. First, due to the limited types of data provided in the NHIRD, we were unable to acquire important suicide-related information (e.g., stigma, family history of suicide, occurrence of material events, and network of support). Second, our study adopted inpatient expenditure files from the NHIRD for analysis; therefore, our study findings may not be generalizable to the entire population without including outpatient cases. Third, all information was collected from medical record (NHIRD). The possibility of information bias cannot be ruled out.

Conclusions
Insomnia is a risk factor for suicide attempts and, indeed, increases the risk for suicide. In addition, low income, drug dependence, and mental disorders are risk factors for suicide attempts. Female patients and those aged 25–44 years are high-risk groups. Risk of suicide attempts is higher in low-income individuals with insomnia and mental disorders. Clinicians should pay attention to the mental status and income level of insomnia patients and implement early suicide prevention intervention. If members of the general public have friends who suffer from insomnia, they should pay attention to the mental and financial status of their friends in order to reduce the probability of suicide.

Additional file

**Additional file 1: Table S1.** Main diagnosis for hospitalization in the baseline. (DOCX 15 kb)

**Abbreviations**
CCI: The Charlson comorbidity index; CI: Confidence interval; HR: Hazard ratio; ICD-9-CM: The International Classification of Diseases, 9th Revision, Clinical Modifications; NHIRD: National Health Insurance Research Database; WHO: World Health Organization

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**Availability of data and materials**
Data are from the National Health Institute Research Database (NHIRD), which are available to researchers in Taiwan and have been extensively used in epidemiologic studies. It is permissible to use these data for academic purposes only after providing proof to National Health Research Institute. Thus, the data cannot be made publicly available. Data requests may be sent to National Health Institute Research Database (http://nhird.nhri.org.tw/) at nhird@nhri.org.tw.
Competing interests

The study protocol was approved by the approval institutional review board, Tri-Service General Hospital. (TSGHIRB No. 1

Ethics approval and consent to participate

The authors declare that they have no competing interests.

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