



# Hospitalization increases while economic status deteriorates in late stages of chronic obstructive pulmonary disease: the Korean National Health and Nutrition Examination Survey for 2007–2015

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**Background:** Chronic obstructive pulmonary disease (COPD) is associated with frequent hospitalizations, higher mortality, and healthcare costs. Low-income COPD patients have higher rates of emergency department visits and hospitalization due to COPD exacerbation. However, other causes of admissions and their economic burden have not been well-elucidated.

**Methods:** We analyzed the Korean National Health and Nutrition Examination Survey (KNHANES) dataset for 2007–2015. The diagnosis and staging of COPD were based on the Global Initiative for Chronic Obstructive Lung Disease (GOLD) guidelines.

**Results:** Among the 97,622 participants in KNHANES for 2007–2015, we selected 33,963 participants (4,430 with and 29,533 without COPD) aged  $\geq 40$  years, who underwent spirometry, and provided the admission history for the previous year. Participants with COPD had a higher admission rate than those without COPD (12.8% vs. 10.4%,  $P < 0.001$ ). The admission rate increased as the stage of COPD advanced from GOLD 1 to GOLD 4 for total causes (11.5%, 13.6%, 15.1%, and 25.0%, respectively,  $P < 0.001$ ) and respiratory illnesses (0.5%, 1.3%, 4.6%, and 12.5%, respectively,  $P < 0.001$ ). The proportion of the lowest quartile household income increased in the late stages of COPD (GOLD 1–4; 35.2%, 32.1%, 44.9%, and 70.8%, respectively,  $P < 0.01$ ).

**Conclusions:** The hospitalization rate increased in advanced COPD, while GOLD stages 3 and 4 were associated with deterioration in economic status.

**Keywords:** Chronic obstructive pulmonary disease (COPD); hospitalization; severity; economic status

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## Introduction

Chronic obstructive pulmonary disease (COPD) is a common respiratory disease affecting  $\geq 5\%$  of the population (1). The prevalence of COPD among the population aged  $\geq 40$  years ranged between 13.1–14.6% during 2010–2015 in Korea (2). COPD is currently the fourth leading cause of death in the United States (3). It is the 8<sup>th</sup> leading cause of death in Korea with a 5-year mortality rate of 25.4% (29.9% in males and 19.1% in females) (4). Patients with COPD struggle with the disease and the associated financial costs. COPD is a costly disease and the burden increases with increasing prevalence. In 2010, COPD-attributable costs for both direct medical care and absenteeism in the US were estimated USD 36 billion, with an expected rise to USD 49 billion by 2020 (5). The estimated total costs for COPD were approximately 1.245 billion USD in Korea in 2015 and direct non-medical costs, such as nursing costs made up the largest percentage (39%) (6). Meanwhile, the average number of days of out-patient care and hospitalization for COPD were  $40 \pm 36$  and  $11 \pm 33$  days, respectively (7). Lost days for care in COPD lead to an inability to work and unemployment. Predictably, an inverse relationship exists between the prevalence of COPD and income levels. COPD prevalence is 1.5–3 times higher among people with low socioeconomic status than among those with higher status and low-socioeconomic status patients occupy two-thirds of the COPD population despite comprising  $< 20\%$  of the general population (8). COPD patients of a lower socioeconomic status tend to have higher rates of emergency room visits and hospitalizations due to acute exacerbations (9). By the way, COPD is linked to several conditions, such as lung cancer, cardiovascular disease, osteoporosis, psychiatric illness, and cognitive dysfunction that may require hospital admission (10). However, the burden of hospitalization for comorbidities has not been well-elucidated in COPD. In addition, the association between the severity of obstruction and the burden of hospitalization rates and costs has not been well-evaluated in COPD.

The present study evaluated the association between economic status and hospitalization rate with the severity of COPD using data from the Korean National Health and Nutrition Examination Survey (KNHANES) collected for 2007–2015. We present the following article in accordance with the STROBE reporting checklist (available at <http://dx.doi.org/10.21037/jtd-20-2683>).

## Methods

### *Korean National Health and Nutrition Examination Survey (KNHANES)*

We analyzed the KNHANES data for 2007–2015. KNHANES is a national surveillance system that has been assessing the health and nutritional status of Koreans annually since 1998. It is an ongoing, population-based, nationwide, cross-sectional survey conducted by the Korea Centers for Disease Control and Prevention (KCDC). KNHANES encompasses approximately 3,800 households across 576 survey areas selected to represent the civilian non-institutionalized South Korean population. Stratified, multistage, and clustered sampling based on National Census Data is used to ensure that survey results represent the general Korean population (11). Results from each year can be merged into a combined database that allows the review of health trends over long periods. All KNHANES survey protocols were approved by the KCDC institutional review board (2015-01-02-6C). Written informed consent was obtained from all the participants and all procedures were conducted in accordance with the Declaration of Helsinki (as revised in 2013).

### *Diagnosis of COPD*

KNHANES has three component surveys: a health interview, health examination, and nutritional survey. The health examination includes a pulmonary function test (PFT) that is performed in subjects  $\geq 40$  years by well-trained technicians following the American Thoracic Society European Respiratory Society criteria for standardization using a dry rolling-seal spirometer (Vmax-2130, Sensor-Medics, Yorba Linda, CA, USA). The diagnosis of COPD should be based on the post-bronchodilator PFT findings under the criteria of the Global Initiative for Chronic Obstructive Lung Disease (GOLD) guidelines. However, the KNHANES is not specifically designed to the diagnosis of COPD and post-bronchodilator PFT was not available, therefore, pre-bronchodilator PFT findings were used in the present study. When the ratio of forced expiratory volume in one second ( $FEV_1$ ) to functional vital capacity (FVC) was  $< 0.7$ , subjects were considered to have COPD. The severity of airflow limitation was classified according to the GOLD criteria (stage I: mild  $FEV_1 \geq 80\%$ , stage II: moderate  $50\% \leq FEV_1 < 80\%$ , stage III: severe  $30\% \leq FEV_1$

<50%, and stage IV: very severe FEV<sub>1</sub> <30%).

### **Information on hospitalization**

The health interview in KNHANES contains a questionnaire used to collect data regarding hospitalization in the previous year such as the number, duration, cost, satisfaction, and reasons for admission. The reasons for hospital admission described by the participants were transcribed using the Korean Informative Classification of Diseases (KOICD) code by attending statisticians. Among others, the KOICD codes for hospitalization for respiratory diseases include acute upper respiratory tract infection (J00–J01, J05–J06), acute pharyngitis/tonsillitis (J02–J03), acute laryngitis/bronchitis (J04), influenza (J10–J11), pneumonia (J12–J18), acute bronchitis/bronchiolitis (J20–J21), other nasal diseases (J30–J31), chronic sinusitis (J32), other paranasal diseases (J33–J34), chronic tonsillitis/adenoiditis (J35), other upper respiratory diseases (J36–J39), bronchitis/emphysema/other COPDs (J40–J44), asthma (J45–J46), bronchiectasis (J47), pneumoconiosis (J60–J65), and other respiratory conditions (J22 and J66–J99). Moreover, we evaluated hospitalization for gastrointestinal diseases, cardiovascular illnesses, and malignancy according to matched KOICD codes.

### **Statistical analysis**

Continuous variables were presented as means ± standard deviations and categorical data as frequencies and percentiles. Intergroup comparisons of continuous variables and categorical data between the two groups were performed with *t*-tests and chi-square tests, respectively. Multiple group analysis among the four stages of COPD GOLD and non-COPD was done with ANOVA. Binary logistic regression analysis for hospitalization with each stage of GOLD was done and adjusted for age, sex, smoking status, and household income. The results were analyzed using SPSS version 16.0 (SPSS Inc., Chicago, IL, USA). Two-tailed analyses were conducted and *P* values <0.05 were considered significant.

## **Results**

### **Study population**

Of the 97,622 participants in KNHANES for 2007–2015, we selected 33,963 participants who were ≥40 years, underwent spirometry and answered the questionnaire for the information on the history of hospital admission in the

previous year. Among the 33,963 participants, COPD was diagnosed for 4,430 participants (13.0%). GOLD 1 (1,997; 45.1%) and GOLD 2 (2,184; 49.3%) constituted most of the COPD population, while GOLD 3 (219; 4.7%) and GOLD 4 (24; 0.5%) constituted a minor portion (*Figure 1*).

### **Comparison of characteristics between participants with and without COPD**

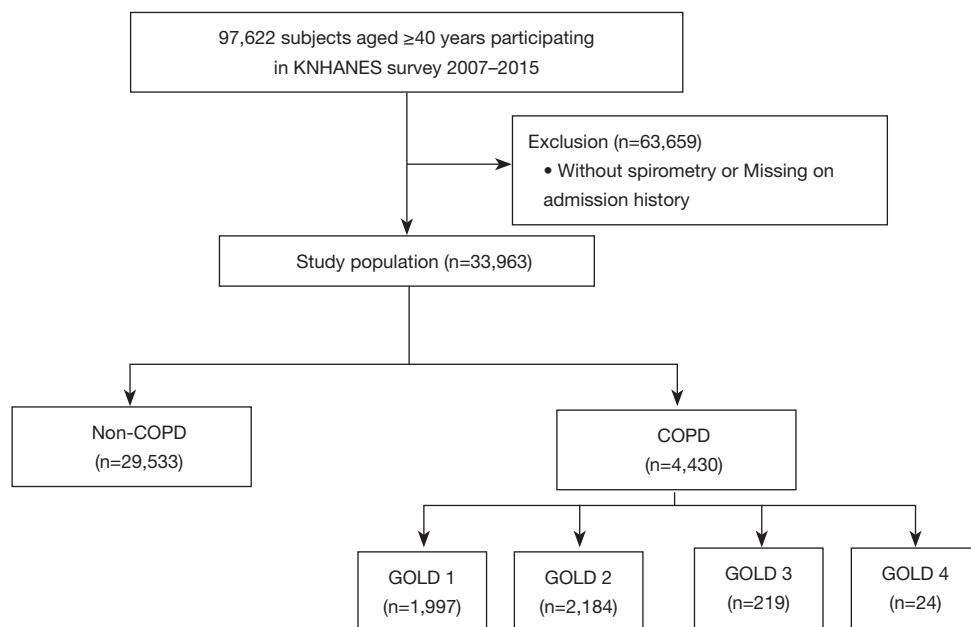
Participants with COPD were older and more likely to be male, and ex- or current-smokers than those without COPD. The lowest quartile of household income occurred with a higher prevalence in the COPD group (34.4% *vs.* 18.1%, *P*<0.001). Participants with COPD had more major comorbidities than those without COPD (*Table 1*).

### **Demographics, pulmonary function, economic status, and comorbidities according to GOLD stage**

As the stage of COPD advanced from GOLD 1 to 4, the current smoking rate increased, while the percent predicted FEV<sub>1</sub> decreased as expected. The proportion of the lowest quartile of household income increased significantly in the late stages of COPD (GOLD 1–4; 35.2%, 32.1%, 44.9%, and 70.8%, respectively, *P*<0.01) (*Table 2*).

### **The rate, number, and causes of hospitalization**

Among the whole study population, 3,647 participants (10.7%) had been hospitalized in the previous year. Most (3,131/3,647, 85.9%) had been admitted once, while others had been hospitalized multiple times (twice in 376/3,647, 10.3%; three times in 87/3,647, 2.4%; more than three times in 52/3,647, 1.4%). Of the 4,430 participants with COPD, 568 had been hospitalized, while 3,070 of the 29,533 participants without COPD had a history of hospitalization in the previous year (12.8% *vs.* 10.4%, *P*<0.01). For causes of admission, patients with COPD had a higher admission rate than those without COPD for respiratory illnesses (1.1% *vs.* 0.4%, *P*<0.01), gastrointestinal diseases (1.3% *vs.* 0.8%, *P*<0.5), and cardiovascular causes (1.3% *vs.* 0.8%, *P*<0.05). Patients with COPD had been hospitalized more frequently than those without COPD for all causes (0.165±0.532 *vs.* 0.125±0.432, *P*<0.01), respiratory causes (0.018±0.246 *vs.* 0.005±0.076, *P*<0.05), gastrointestinal causes (0.015±0.139 *vs.* 0.009±0.104, *P*<0.05), and cardiovascular causes (0.014±0.144 *vs.* 0.009±0.099, *P*<0.05). The five most common diseases for hospitalization



**Figure 1** Flow chart representing the selection of the study population. COPD, chronic obstructive pulmonary disease; GOLD, Global Initiative for Chronic Obstructive Lung Disease.

in each category of respiratory disorders, gastrointestinal illnesses, cardiovascular abnormalities, and neoplasm were summarized at a separate table (Table 3).

#### *Hospitalization rate according to GOLD stage*

As the stage of COPD advanced from GOLD 1 to 4, the admission rate increased for total causes (non-COPD to GOLD 1–4; 10.4%, 11.5%, 13.6%, 15.1%, and 25.0%,  $P < 0.01$ ) and respiratory causes (Non-COPD to GOLD 1–4; 0.4%, 0.5%, 1.3%, 4.6%, and 12.5%,  $P < 0.01$ ) (Figure 2). Binary logistic regression analysis adjusted for age, sex, smoking status, and household income showed that hospitalization for total causes showed an increasing tendency compared to non-COPD as the stage of COPD advanced (Odds ratio of GOLD 1–4; 1.072, 1.325, 1.365, and 2.630) which was statistically significant in GOLD 2 and GOLD 4. Respiratory admission also showed an increasing tendency in accordance with COPD stage compared to non-COPD (odds ratio for GOLD 1–4; 0.862, 2.649, 8.621, and 26.374), which was statistically significant in GOLD 2–4 (Table 4).

## Discussion

The present study evaluated the burden of hospitalization

for the number, rate and economic aspect among KNHANES participants for 2007–2015, depending on the presence and severity of COPD. We showed that participants with COPD had a higher rate and frequency of hospitalization in the previous year than those without COPD. For causes of admission, patients with COPD had a higher rates and frequency of hospital admissions than those without COPD for respiratory illnesses, gastrointestinal diseases, and cardiovascular causes. Of note, the admission rate increased for total and respiratory causes as the stage of COPD advanced from GOLD 1 to 4, while the economic status deteriorated in the late stages of COPD.

As expected, patients with COPD had a higher number and rate of hospitalization in the previous year due to respiratory illnesses. In one large population-based study, COPD always occupied the first place in the frequency of hospital admissions for adult patients among four major respiratory diseases such as COPD, pneumonia, lung cancer, and asthma (12). It is well-known that patients with COPD often suffer from multiple comorbidities (13,14). In particular, COPD is associated with increased an risk of cardiovascular morbidities and approximately one-third of COPD deaths are estimated to be due to cardiovascular causes (15,16). Moreover, physical inactivity accompanying COPD frequently leads to functional constipation,

**Table 1** Demographics, pulmonary function, economic status and comorbidities

| Variable                 | Total          | Non-COPD       | COPD          | P value |
|--------------------------|----------------|----------------|---------------|---------|
| Number                   | 33,963 (100%)  | 29,533 (87.0%) | 4,430 (13.0%) | –       |
| Male sex                 | 14,900 (43.9%) | 11,730 (39.7%) | 3,170 (71.6%) | <0.001  |
| Age (years)              | 54.38±13.24    | 52.89±12.90    | 64.35±10.90   | <0.001  |
| Smoking status           |                |                |               | <0.001  |
| Non-smoker               | 58.3%          | 62.2%          | 32.0%         |         |
| Ex-smoker                | 14.2%          | 12.3%          | 27.0%         |         |
| Current smoker           | 27.5%          | 25.5%          | 40.9%         |         |
| Pulmonary function tests |                |                |               |         |
| FVC (liter)              | 3.48±0.88      | 3.47±0.88      | 3.54±0.92     | <0.001  |
| FVC %                    | 92.31±17.34    | 92.68±17.69    | 89.79±14.56   | <0.001  |
| FEV <sub>1</sub> (liter) | 2.71±0.73      | 2.79±0.71      | 2.24±0.65     | <0.001  |
| FEV <sub>1</sub> %       | 91.76±13.90    | 93.92±12.16    | 77.35±16.01   | <0.001  |
| FEV <sub>1</sub> /FVC    | 0.78±0.08      | 0.80±0.05      | 0.63±0.07     | <0.001  |
| Household income*        |                |                |               | <0.001  |
| Lowest quartile          | 6,778 (20.2%)  | 5,282 (18.1%)  | 1,496 (34.4%) |         |
| Lower middle quartile    | 8,503 (25.4%)  | 7,315 (25.1%)  | 1,188 (27.3%) |         |
| Upper middle quartile    | 8,660 (25.8%)  | 7,791 (26.7%)  | 869 (20.0%)   |         |
| Highest quartile         | 9,596 (28.3%)  | 8,800 (30.1%)  | 796 (18.3%)   |         |
| Comorbidity              |                |                |               |         |
| Hypertension             | 8,933 (51.7%)  | 7,248 (49.7%)  | 1,685 (62.5%) | <0.001  |
| Diabetes                 | 3,310 (24.1%)  | 2,664 (22.9%)  | 646 (31.0%)   | <0.001  |
| Chronic renal disease    | 146 (1.2%)     | 121 (1.2%)     | 25 (1.4%)     | 0.398   |
| Liver cirrhosis          | 117 (1.0%)     | 88 (0.9%)      | 29 (1.7%)     | 0.002   |
| Malignancy               | 556 (1.6%)     | 437 (4.4%)     | 119 (6.9%)    | <0.001  |

\*, household income was measured as combined income from all sources of the respondent and family members and was divided to four quartiles according to criteria of each year.

and the use of inhaled long-acting muscarinic receptor antagonists may cause anti-cholinergic adverse events such as constipation in patients with COPD (17). Corticosteroid use is associated with an increased risk of duodenal ulcer perforation. Because inhaled and systemic corticosteroid are frequently used in the management of COPD, up to 30% of COPD patients have been found to have peptic ulcers, and COPD frequency in peptic ulcer disease is 2–3 times the general population (18). This finding is consistent with our results that participants with COPD had more major comorbidities such as hypertension, diabetes, liver cirrhosis,

and malignancy than those without COPD (*Table 1*). Furthermore, our analysis showed that COPD patients were more frequently hospitalized than those without COPD not only due to respiratory causes but also for cardiovascular and gastrointestinal illnesses.

Previous reports have shown that the severity of COPD is associated with the frequency of hospitalization. Patients in the GOLD D category experienced nearly three times the number of exacerbations and COPD-related admissions as those in the GOLD A category (19). The present analysis showed that both acute exacerbation and respiratory

**Table 2** Demographics, pulmonary function, economic status and comorbidities according to the GOLD stage

| Variable                             | Non-COPD       | GOLD 1         | GOLD 2         | GOLD 3         | GOLD 4         | P value | P value** |
|--------------------------------------|----------------|----------------|----------------|----------------|----------------|---------|-----------|
| Number (%)                           | 29,533 (87.0%) | 1,997 (5.9%)   | 2,184 (6.4%)   | 219 (0.6%)     | 24 (0.1%)      | –       | –         |
| Male sex (%)                         | 11,730 (39.7%) | 1,453 (72.8%)  | 1,540 (70.5%)  | 151 (68.9%)    | 21 (87.5%)     | <0.001  | <0.001    |
| Age (years)                          | 52.89±12.90    | 65.77±10.04    | 63.09±11.39    | 63.82±12.17    | 66.1±8.10      | <0.001  | <0.001    |
| Smoking status<br>(non:ex:current %) | 62.2:12.3:25.5 | 31.2:31.0:37.9 | 33.1:24.1:42.9 | 30.7:21.1:48.2 | 20.8:29.2:50.0 | <0.001  | <0.001    |
| Pulmonary function test              |                |                |                |                |                |         |           |
| FVC (liter)                          | 3.47±0.88      | 3.92±0.86      | 3.29±0.79      | 2.62±0.78      | 2.55±0.79      | <0.001  |           |
| FVC %                                | 92.68±17.69    | 100.01±10.50   | 82.93±10.43    | 68.17±13.83    | 61.72±15.58    | <0.001  |           |
| FEV <sub>1</sub> (liter)             | 2.79±0.71      | 2.60±0.59      | 2.03±0.50      | 1.20±0.29      | 0.82±0.13      | <0.001  |           |
| FEV <sub>1</sub> %                   | 93.92±12.16    | 90.92±8.66     | 68.99±7.70     | 42.38±5.49     | 27.76±2.62     | <0.001  |           |
| FEV <sub>1</sub> /FVC                | 80.29±5.16     | 66.25±3.37     | 62.3±6.54      | 47.63±9.66     | 34.76±9.41     | <0.001  |           |
| Household income*                    |                |                |                |                |                | <0.001  | 0.002     |
| Lowest quartile                      | 5,282 (18.1%)  | 694 (35.2%)    | 686 (32.1%)    | 96 (44.9%)     | 17 (70.8%)     |         |           |
| Lower middle quartile                | 7,315 (25.1%)  | 520 (26.4%)    | 602 (28.2%)    | 62 (29.0%)     | 3 (12.5%)      |         |           |
| Upper middle quartile                | 7,791 (26.7%)  | 372 (18.9%)    | 462 (21.6%)    | 31 (14.5%)     | 3 (12.5%)      |         |           |
| Highest quartile                     | 8,800 (30.1%)  | 386 (19.6%)    | 384 (18.0%)    | 25 (11.7%)     | 1 (4.2%)       |         |           |
| Comorbidity                          |                |                |                |                |                |         |           |
| Hypertension                         | 7,248 (49.7%)  | 747 (59.1%)    | 841 (65.1%)    | 82 (68.3%)     | 11 (68.8%)     | <0.001  | 0.175     |
| Diabetes                             | 2,664 (22.9%)  | 273 (27.1%)    | 339 (34.3%)    | 31 (37.3%)     | 3 (33.3%)      | <0.001  | 0.375     |
| Chronic renal disease                | 121 (1.2%)     | 6 (0.7%)       | 18 (2.2%)      | 1 (1.5%)       | 0 (0%)         | 0.061   | 0.208     |
| Chronic liver disease                | 88 (0.9%)      | 10 (1.2%)      | 17 (2.1%)      | 1 (1.5%)       | 1 (11.1%)      | <0.001  | 0.995     |
| Malignancy                           | 437 (4.4%)     | 55 (6.4%)      | 63 (8.0%)      | 1 (1.5%)       | 0 (0%)         | <0.001  | 0.354     |

\*, household income was measured as combined income from all sources of the respondent and family members and was divided to 4 quartiles according to criteria of each year; \*\*, multivariate linear regression was performed for age, sex, smoking status, household income, and comorbidity. GOLD, Global Initiative for Chronic Obstructive Lung Disease.

illnesses generally increased hospitalization in COPD. Moreover, the total hospitalization rate increased as the stage progressed in COPD (*Figure 2*).

Patients with COPD suffer from the diseases and their associated financial burden. COPD is a disease of the poor. The Center for Disease Control and Prevention reported that 20.4% of those with COPD were unable to work compared to 4.8% without the disease (20). Americans older than 50 years who had COPD were less likely to be employed than those without COPD or those with cancer, heart disease, hypertension, or diabetes (21). Frequent hospitalization places a tremendous financial burden on individuals of low socioeconomic status. Unfortunately, that was what happened in the real world; the hospitalization rate

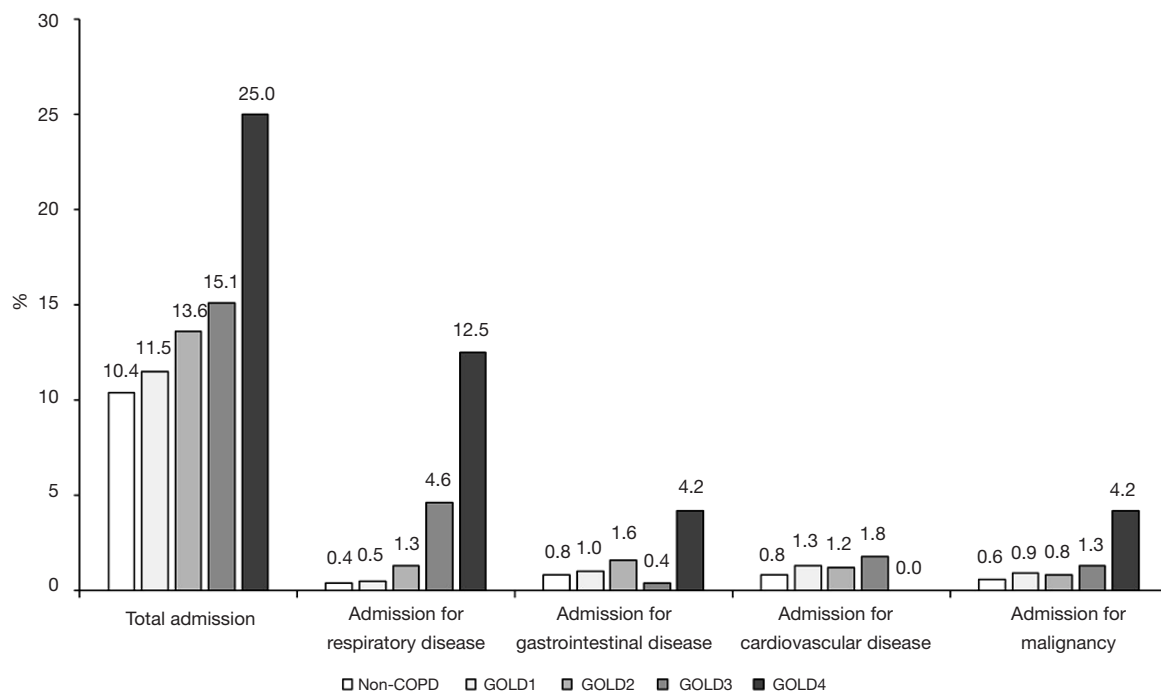
increased as the COPD stage advanced (*Table 3, Figure 2*), while the lowest quartile of household income increased sharply in the late stages of COPD (*Table 2*). A striking observation was that the percentile of the lowest quartile of household income in GOLD 4 was >70%.

The present study has some limitations. First, the KNHANES dataset is a cross-sectional dataset. The positive associations between the number and rates of hospitalization for COPD do not imply causality. Second, although we were able to determine that low economic status was associated with the late stages of COPD, we were unable to determine the underlying causes. Third, the diagnosis of COPD was based on the pre-bronchodilator PFT findings instead of post-bronchodilator ones because it

**Table 3** The five most common diseases of hospitalization in each category of disorder

|   | Respiratory illness                | Gastrointestinal disease                       | Cardiovascular disease                          | Neoplasm                     |
|---|------------------------------------|------------------------------------------------|-------------------------------------------------|------------------------------|
| 1 | Pneumonia [39]                     | Other bowl and peritoneal diseases [113]       | Other ischemic heart disease [46]               | Rectal cancer [50]           |
| 2 | Asthma [25]                        | Cholelithiasis and cholecystitis [67]          | Intracranial hemorrhage [28]                    | Breast cancer [38]           |
| 3 | COPD [20]                          | Appendix disease [46]                          | Acute myocardial infarction [25]                | Leiomyoma of the uterus [36] |
| 4 | Influenza [16]                     | Esophageal, gastric and duodenal diseases [34] | Cerebral infarction [17]                        | Stomach cancer [33]          |
| 5 | Acute pharyngitis, tonsillitis [5] | Other liver diseases [23]                      | Conduction disorder and cardiac arrhythmia [15] | Lung cancer [5]              |

The number in square bracket means the number of hospitalization.



**Figure 2** Hospitalization percentage in the previous year for all causes, respiratory diseases, gastrointestinal diseases, cardiovascular diseases, and malignancy according to the GOLD stage. GOLD, Global Initiative for Chronic Obstructive Lung Disease.

was only available in the KNHANES, which is a deviation from GOLD guideline of COPD diagnosis. Nevertheless, to the best of our knowledge, this is the first nation-wide analysis showing that hospitalization increased in the later stages of COPD not only for respiratory and cardiovascular diseases but also for gastrointestinal and all related causes. Moreover, we were able to determine the burden of

hospitalization for the number, rate, and economic aspects according to the severity of COPD.

In conclusion, the hospitalization rate increased as the stage of COPD advanced, while the economic status of the patients deteriorated in GOLD stages 3 and 4. The burden of admission, including the rate, number, and costs increased in the late stages of COPD.

**Table 4** Association between GOLD stages and the risk of hospitalization

| Admission                | Non-COPD      | GOLD1       | GOLD2       | GOLD3       | GOLD4        |
|--------------------------|---------------|-------------|-------------|-------------|--------------|
| Total causes illness     | 3,079 (10.4%) | 229 (11.5%) | 298 (13.6%) | 33 (15.1%)  | 6 (25.0%)    |
| Odds ratio               | 1             | 1.072       | 1.325       | 1.365       | 2.630        |
| 95% CI                   |               | 0.922–1.246 | 1.157–1.16  | 0.929–2.007 | 1.040–6.648  |
| P value                  |               | 0.367       | <0.001      | 0.113       | 0.041        |
| Respiratory illness      | 128 (0.4%)    | 9 (0.5%)    | 28 (1.3%)   | 10 (4.6%)   | 3 (12.5%)    |
| Odds ratio               | 1             | 0.862       | 2.649       | 8.621       | 26.374       |
| 95% CI                   |               | 0.427–1.740 | 1.701–4.126 | 4.218–1.617 | 7.561–92.004 |
| P value                  |               | 0.678       | <0.001      | <0.001      | <0.001       |
| Gastrointestinal disease | 246 (0.8%)    | 20 (1.0%)   | 35 (1.6%)   | 1 (0.4%)    | 1 (4.2%)     |
| Odds ratio               | 1             | 0.949       | 1.535       | 0.422       | 3.510        |
| 95% CI                   |               | 0.587–1.534 | 1.06–2.253  | 0.059–3.041 | 0.467–26.379 |
| P value                  |               | 0.831       | 0.029       | 0.392       | 0.222        |
| Cardiovascular disease   | 244 (0.8%)    | 26 (1.3%)   | 27 (1.2%)   | 4 (1.8%)    | 0 (0%)       |
| Odds ratio               | 1             | 1.117       | 1.115       | 1.587       | –            |
| 95% CI                   |               | 0.726–1.719 | 0.733–1.697 | 0.579–4.53  | –            |
| P value                  |               | 0.615       | 0.611       | 0.369       | 0.998        |
| Neoplasm                 | 193 (0.6%)    | 19 (0.9%)   | 19 (0.8%)   | 3 (1.3%)    | 1 (4.2%)     |
| Odds ratio               | 1             | 1.333       | 1.266       | 2.278       | 7.579        |
| 95% CI                   |               | 0.788–2.254 | 0.754–2.127 | 0.714–7.269 | 1.004–57.215 |
| P value                  |               | 0.284       | 0.372       | 0.164       | 0.050        |

In this binary logistic regression analysis, results were adjusted by age, sex, smoking status, and household income. GOLD, Global Initiative for Chronic Obstructive Lung Disease.

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## Footnote

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uniform disclosure form (available at <http://dx.doi.org/10.21037/jtd-20-2683>). The authors have no conflicts of interest to declare.

**Ethical Statement:** The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). All KNHANES survey protocols were approved by the KCDC institutional review board (2015-01-02-6C). Written informed consent was obtained from all the participants.

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