Factors Associated with Depressive Symptom in Chronic Obstructive Pulmonary Disease Based on the Third (2005) and Fourth (2008) Korea National Health and Nutritional Examination Survey (KNHANES III, IV)

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Background: Depressive symptoms accompanied by chronic obstructive pulmonary disease (COPD) can be influenced by socioeconomic status, associated chronic diseases and the current smoking status. This study was conducted to assess factors that are associated with depressive symptoms accompanied by COPD, using the data obtained from the Korea National Health and Nutritional Survey (KNHANES) conducted in 2005 and 2008.

Methods: From the third (2005) and the fourth (2008) KNHANES, 407 (0.9%) with physician-diagnosed COPD were selected. Of the 407 subjects, only 180 (0.4%) who reported having depressive symptoms were included in this study. The associations of depressive symptoms with socioeconomic status, associated chronic diseases and smoking status were investigated.

Results: Of the total 180 subjects, 45 (25%) had depressive symptoms. There were 102 males (55%) and 78 females (45%) with a slight predilection for males. In multivariate analysis, significant predictors of depressive symptoms were dependent activities of daily living (odds ratio [OR], 2.42; 95% confidence interval [CI], 2.06 to 2.84) and association with number of chronic diseases (OR of one, two, and three, 1.40, 1.72, 2.60; 95% CI of one, two, and three, 1.20 to 1.63, 1.41 to 2.10, 1.99 to 3.39).

Conclusion: This study provides the basis for managing COPD patients in a clinical setting by understanding the number and characteristics of COPD patients with depressive symptoms. The results of this study suggest that primary physicians should manage COPD patients with consideration of risk factors for depressive symptom.

Keywords: Pulmonary Disease, Chronic Obstructive; Depression; Smoking; KNAHNES

Received: August 27, 2010, Accepted: May 16, 2011

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INTRODUCTION

Chronic obstructive lung disease (COPD) is extremely difficult to treat successfully. It is expected to rank third among the causes of death in 2020.1 In Korea, the prevalence of COPD has increased to 17.2% in subjects aged ≥ 45 years.2 Since smoking is known to be the most important etiology of COPD and to affect prognosis, smoking cessation is most important in the prevention...
of COPD.

Mood disorders are frequently associated with COPD. In particular, 6% to 46% of COPD patients had associated depression. The success of smoking cessation depends considerably on the absence or presence of associated depression, and the success rate of quitting smoking is lower in smokers with depression than in those without it. Associated mood disorders may influence emotional status, physical activity and quality of life as well as clinical course and mortality. For these reasons, knowledge of the association of depression symptom with COPD is important in the management of COPD patients. Depressive symptoms accompanied by COPD can be influenced by various factors, such as socioeconomic status, associated chronic disease and the current smoking status. It is essential to understand factors that are associated with depressive symptoms and to maintain the effects of smoking cessation in patients with COPD.

Although there have been numerous studies on the early diagnosis, prevention and treatment of COPD, few reports have been found on the prevalence of associated depressive symptoms and therapeutic approaches to the socio-psychological aspect. Furthermore, there have been only a few studies in Korean subjects and those who have largely been localized. For this reason, these studies are difficult to apply in the general population.

This study was conducted to assess factors that are associated with depressive symptom accompanied by COPD, using the data obtained from the 2005 and 2008 Korea National Health and Nutritional Survey (KNHANES) and to assist for the management of depressive symptom in COPD patients.

METHODS

1. Study Subjects

The KNHANES has been conducted every 3 years using a probability cluster sample that is representative of Korean people in accordance with the National Health Promotion Law legislated in 1995. The first KNHANES was conducted in 1998, the second in 2001, the third in 2005 and the fourth in 2008.

The number of subjects who responded to the third (2005) and the fourth (2008) KNHANES was 43,889 in total. Of the 43,889 subjects, 407 (0.9%) who reported having depressive symptom were included in this study.

2. Methods

As for depressive symptom, subjects were asked to answer the question, “Have you felt sad or desperate for ≥ 2 weeks during the past 1 year so as to disturb everyday life?” Subjects who reported “yes” were regarded as having depressive symptom. Subjects were asked to report their smoking status as one of the following: current smoking, past smoking and others. Subjects who reported “current smoking” were regarded as current smokers. Current and past smokers were asked to report their smoking amounts as follows: < 10 cigarettes per day and ≥ 10 cigarettes per day. Their intention of smoking cessation was assessed in terms of time point: “within 1 month,” “within 6 months,” “not decided” and “no intention.” Their experience with smoking cessation campaigns was assessed. Socio-demographic factors, including age, sex, marital status, educational level and household income, were assessed. The subjects were divided into 3 groups: the ≤ 64, 65–74, and ≥ 75 age groups. Marital status was reported as “married,” “unmarried,” “separated,” “widowed” and “divorced,” and it was grouped as “living with a spouse” and “others.” The educational level was reported as either high school or < high school. Household income was reported as lower (within the first and second quartiles) and higher (within the third and fourth quartiles). Perceived health was reported as “very good,” “good,” “moderate,” “bad,” or “very bad,” and then it was grouped as “very good/good/moderate” and “bad/very bad.” As for the activities of daily living (ADL), no disturbances in everyday life were defined when subjects answered “n” to the question “Do you have any disturbances in everyday life?” Chronic diseases included stroke, hypertension, diabetes mellitus, ischemic heart disease, liver cirrhosis, arthritis, asthma and chronic renal failure, which were selected as main diseases in the 2005 and 2008 KNHANES. Subjects with one or more cancers in the stomach, liver, lung, colon, cervix or breast were defined as ‘major cancer’ subjects, which were based on the answer “y” to the question “Were you diagnosed with cancer(s) by a physician?” in order to avoid bias generated by subjective assessment. Subjects with a body mass index (BMI) of <18.5 were classified as underweight, those with a BMI of 18.5–24.9 as normal weight, and those with a BMI of ≥ 25
as obese.

3. Statistical Analysis

Statistical analyses were performed using SPSS ver. 17.0 (SPSS Inc., Chicago, IL, USA). A P-value of <0.05 was considered statistically significant. The distribution of variables in subject groups with and without depressive symptom was assessed using the chi-square test, and differences in variables were examined using the chi-square test and Fischer’s exact test. Each frequency was calculated by adding weighted values (variable with weighted values, wt_itv). In multivariate analysis, factors associated with depressive symptom were examined. Independent risk factors were investigated by logistic regression analysis. In model 1, variables were analyzed without any adjustment, while in model 2, variables were analyzed after adjustment for sex, age, household income, educational level and current smoking status. All analyses were performed by reflecting weighted values at the stratification of the samples applied to the KNHANEs.

RESULTS

1. Socio-Demographic Characteristics of Subjects (Table 1)

Of the total 180 subjects, 45 (25%) had depressive symptom. There were 102 males (55%) and 78 females (45%) with slight predilection for males. There were 116 subjects (66.3%) in the ≤64 age group, 40 subjects (20.1%) in the 65–74 age group, and 24 (13.6%) in the ≥75 age group. Subjects with a spouse accounted for 116 (68.4%) of all subjects. Subjects with < middle school education accounted for 116 (58.1%) of all subjects. Subjects with lower income accounted for 113 (60.7%) of all subjects. There were no significant differences in perceived health between subjects in good health status and those who were not. Disturbances in everyday life were more common in the independent group than in the dependent group. The number of associated chronic disease was none in 71 subjects (42.2%), 1 in 60 subjects (30.3%), 2 in 26 subjects (17.0%), and ≥3 in 23 subjects (10.5%). Main cancers were observed in 7 subjects (3.0%). As assessed by BMI, 36 subjects (16.2%) were obese, 105 subjects (75.3%) were normal weight, and 8 subjects (8.5%) were underweight. The number of current smokers was 47 (22.8%).

| Characteristics                          | Total (n = 180) | Standard error (%) |
|-----------------------------------------|----------------|-------------------|
| Sex                                     |                |                   |
| Male                                    | 102 (55.0)    | 3.5               |
| Female                                  | 78 (45.0)     | 3.5               |
| Age                                     |                |                   |
| ≤64                                     | 116 (66.3)    | 3.1               |
| 65–74                                   | 40 (20.1)     | 2.6               |
| ≥75                                     | 24 (13.6)     | 2.6               |
| Living with a spouse                    |                |                   |
| Yes                                     | 135 (68.4)    | 3.3               |
| No                                      | 45 (31.6)     | 3.3               |
| Education                               |                |                   |
| ≥High school                            | 64 (41.9)     | 4.4               |
| ≤Middle school                          | 116 (58.1)    | 4.4               |
| Income                                  |                |                   |
| Lower                                   | 113 (60.7)    | 4.3               |
| Higher                                  | 63 (39.3)     | 4.3               |
| Perceived health                        |                |                   |
| Not available                           | 4 (2.2)       |                   |
| Good                                    | 92 (50.3)     | 3.4               |
| Not good                                | 88 (49.7)     | 3.4               |
| Activities of daily living              |                |                   |
| Independent                             | 139 (76.7)    | 3.2               |
| Dependent                               | 41 (23.3)     | 3.2               |
| No. of chronic disease                  |                |                   |
| 0                                       | 71 (42.2)     | 3.5               |
| 1                                       | 60 (30.3)     | 2.9               |
| 2                                       | 26 (17.0)     | 1.9               |
| ≥3                                      | 23 (10.5)     | 2.8               |
| Major cancer                            |                |                   |
| No                                      | 173 (97.0)    | 1.0               |
| Yes                                     | 7 (3.0)       | 1.0               |
| Body mass index                         |                |                   |
| Obese                                   | 36 (16.2)     | 4.1               |
| Normal weight                           | 105 (75.3)    | 4.4               |
| Underweight                             | 8 (8.5)       | 2.9               |
| Current smoking status                  |                |                   |
| Not available                           | 31 (17.2)     |                   |
| Current smoker                          | 47 (22.8)     | 2.6               |
| No current smoker                       | 133 (72.2)    | 2.6               |

*Weighted percent. †Chronic disease: stroke, hypertension, diabetes, ischemic heart disease, liver cirrhosis, arthritis, asthma and chronic renal failure. ‡Major cancer: stomach, liver, lung, cervix, breast and colon cancers. §BMI: obese, ≥25; normal weight, 18.5–24.9; underweight, <18.5.
Table 2. Depressive symptom according to socio-demographic characteristics, chronic disease and smoking status in study subjects.

| Variable                  | Total no. | Depressive symptom | P-value |
|---------------------------|-----------|--------------------|---------|
|                           |           | Yes                | No      |
|                           |           | No. (%)*           | SE (%)  | No. (%)* | SE (%)  |
| Sociodemographic          |           |                    |         |
| Sex                       | 180       |                    |         |
| Male                      | 20 (14.4) | 3.6                | 82 (85.6) | 3.6     |
| Female                    | 25 (36.0) | 8.1                | 53 (64.0) | 8.1     |
| Age                       | 180       |                    |         |
| ≤64                       | 31 (24.3) | 4.2                | 85 (75.7) | 4.2     |
| 65–74                     | 10 (22.6) | 8.2                | 30 (77.4) | 8.2     |
| ≥75                       | 4 (27.9)  | 17.4               | 20 (72.1) | 17.4    |
| Married                   | 180       |                    |         |
| Yes                       | 31 (19.4) | 3.2                | 104 (80.6) | 3.2  |
| No                        | 14 (38.2) | 13.0               | 31 (61.8) | 13.0    |
| Education                 | 180       |                    |         |
| ≥High school              | 16 (20.6) | 3.9                | 48 (79.4) | 3.9     |
| ≤Middle school            | 29 (27.2) | 5.9                | 87 (72.8) | 5.9     |
| Income                    | 176       |                    |         |
| Lower                     | 31 (23.4) | 3.5                | 82 (76.6) | 3.5     |
| Higher                    | 12 (19.3) | 5.7                | 51 (80.7) | 5.7     |
| Perceived health          | 180       |                    |         |
| Good                      | 20 (20.6) | 5.4                | 72 (79.4) | 5.4     |
| Not good                  | 25 (29.0) | 8.1                | 63 (71.0) | 8.1     |
| Activities of daily living| 180       |                    |         |
| Independent               | 27 (16.4) | 3.6                | 112 (83.6) | 3.6 |
| Dependent                 | 18 (52.0) | 12.1               | 23 (48.0) | 12.1    |
| No. of chronic disease†   | 180       |                    |         |
| 0                         | 16 (21.7) | 5.8                | 55 (78.3) | 5.8     |
| 1                         | 13 (20.9) | 5.0                | 47 (79.1) | 5.0     |
| 2                         | 4 (11.0)  | 6.7                | 22 (89.0) | 6.7     |
| ≥3                        | 12 (60.4) | 17.2               | 11 (39.6) | 17.2    |
| Major cancer‡             | 180       |                    |         |
| No                        | 44 (25.2) | 4.1                | 129 (74.8) | 4.1  |
| Yes                       | 1 (6.1)   | 6.3                | 6 (93.9)  | 6.3     |
| Body mass index§           | 149       |                    |         |
| Obese                     | 8 (20.9)  | 6.8                | 28 (79.1) | 6.8     |
| Normal weight             | 27 (23.8) | 5.7                | 78 (76.2) | 5.7     |
| Underweight               | 3 (53.1)  | 22.8               | 5 (46.9)  | 22.8    |
| Chronic disease           |           |                    |         |
| Stroke                    | 180       |                    |         |
| No                        | 43 (13.2) | 13.3               | 133 (86.8) | 13.3 |
| Yes                       | 2 (21.9)  | 2.9                | 2 (78.1)  | 2.9     |
| Hypertension              | 180       |                    |         |
| No                        | 31 (30.6) | 12.7               | 102 (69.4) | 12.7  |
| Yes                       | 14 (23.0) | 4.3                | 33 (77.0) | 4.3     |
Table 2. Continued.

| Variable                        | Total no. | Depressive symptom | P-value |
|---------------------------------|-----------|--------------------|---------|
|                                |           | Yes                |         |
|                                 |           | No. (%)* | SE (%) | Yes. (%)* | SE (%) |
| Diabetes                        | 180       | 38 (19.7) | 8.9 | 125 (80.3) | 8.9 | 0.64 |
| No                              | 38 (19.7) | 8.9 | 125 (80.3) | 8.9 | 0.64 |
| Yes                             | 7 (25.1)  | 4.7 | 10 (74.9)  | 4.7 | 0.64 |
| Ischemic heart disease          | 180       | 39 (58.6) | 48.5 | 129 (41.4) | 48.5 | 0.11 |
| No                              | 39 (58.6) | 48.5 | 129 (41.4) | 48.5 | 0.11 |
| Yes                             | 6 (54.9)  | 22.5 | 6 (45.1)   | 22.5 | 0.11 |
| Liver cirrhosis                 | 180       | 45 (24.7) | 4.1 | 135 (75.3) | 4.1 | NA   |
| No                              | 45 (24.7) | 4.1 | 135 (75.3) | 4.1 | NA   |
| Yes                             | 0         | 0    | 0         | 0    | NA   |
| Arthritis                       | 180       | 26 (6.8)  | 7.2 | 101 (93.2) | 7.2 | 0.004|
| No                              | 26 (6.8)  | 7.2 | 101 (93.2) | 7.2 | 0.004|
| Yes                             | 19 (53.1) | 11.3 | 34 (46.9)  | 11.3 | 0.004|
| Asthma                          | 180       | 38 (25.9) | 4.8 | 109 (74.1) | 4.8 | 0.36 |
| No                              | 38 (25.9) | 4.8 | 109 (74.1) | 4.8 | 0.36 |
| Yes                             | 7 (17.0)  | 7.0 | 26 (83.0)  | 7.0 | 0.36 |
| Chronic renal failure           | 180       | 43 (24.6) | 4.2 | 130 (75.4) | 4.2 | 0.90 |
| No                              | 43 (24.6) | 4.2 | 130 (75.4) | 4.2 | 0.90 |
| Yes                             | 2 (23.0)  | 10.9 | 5 (77.0)   | 10.9 | 0.90 |
| Smoking status                  |           |         |         |         |         | 0.36 |
| Smoking                         | 180       |         |         |         |         | 0.36 |
| Current smoking                 | 12 (32.6) | 9.6 | 35 (67.4)  | 9.6 | 0.36 |
| No current smoking              | 33 (22.2) | 5.2 | 100 (77.8) | 5.2 | 0.36 |
| Mean smoking per day in current smokers | 47       |         |         |         |         | 0.13 |
| ≥10 cigarettes                  | 10 (25.2) | 7.2 | 31 (74.8)  | 7.2 | 0.13 |
| <10 cigarettes                  | 2 (36.9)  | 23.7 | 4 (63.1)   | 23.7 | 0.13 |
| Intention of smoking cessation in current smokers | 47       |         |         |         |         | 0.42 |
| In 1 mo                         | 3 (42.9)  | 29.3 | 4 (47.1)   | 29.3 | 0.42 |
| In 6 mo                         | 0 (0.0)   | 0    | 6 (100.0)  | 0    | 0.42 |
| At some time                    | 5 (26.3)  | 18.1 | 14 (73.7)  | 18.1 | 0.42 |
| No intention                    | 4 (26.7)  | 13.2 | 11 (73.3)  | 13.2 | 0.42 |
| Mean smoking amount per day in past smokers | 49       |         |         |         |         | 0.74 |
| ≥10 cigarettes                  | 7 (16.7)  | 8.4 | 35 (83.3)  | 8.4 | 0.74 |
| <10 cigarettes                  | 2 (28.6)  | 14.3 | 5 (71.4)   | 14.3 | 0.74 |
| Experience of smoking cessation campaigns | 166     |         |         |         |         | 0.65 |
| Yes                             | 41 (23.6) | 3.4 | 119 (76.4) | 3.4 | 0.65 |
| No                              | 1 (15.3)  | 14.7 | 5 (84.7)   | 14.7 | 0.65 |

NA: not applicable.
*Weighted percent. †Chronic disease: stroke, hypertension, diabetes, ischemic heart disease, liver cirrhosis, arthritis, asthma and chronic renal failure. ‡Major cancer: stomach, liver, lung, cervix, breast, colon cancer. §BMI: obese, ≥25; normal weight, 18.5–24.9; underweight, <18.5.
2. Comparison of Characteristics according to Depressive Symptom (Table 2)

Depressive symptom tended to occur more frequently in females with statistical significance. Depressive symptom was not significantly associated with age, the presence of a spouse, educational level, household income level or perceived health. The prevalence of depressive symptom was significantly higher in subjects with dependent ADL than in those with independent ADL. Depressive symptom occurred more frequently in subjects with associated chronic diseases. The presence of ≥3 main cancers was not significantly associated with the risk for depressive symptom. BMI tended to be more frequently associated with depressive symptom, but without any statistical significance. As for chronic disease, ischemic heart disease, hypertension, diabetes mellitus, asthma and chronic renal failure were not significantly associated with depressive symptom, whereas stroke and arthritis were significantly associated with depressive symptom. Depressive symptom was not associated with current smoking, mean smoking amount, intention of smoking cessation or experience with smoking cessation campaigns.

3. Predictors of Depressive Symptoms (Table 3)

In multivariate analysis, dependent ADL (OR, 2.42; 95% CI, 2.06 to 2.84) and association with the number of chronic diseases (OR of one, two, and three, 1.40, 1.72, 2.60; 95% CI of one, two, and three, 1.20—1.63,1.41—2.10,1.99—3.39) were still significant predictors even after adjustment.

**DISCUSSION**

In this study of COPD patients in the general Korean population, dependent ADL and association with chronic diseases were the significant predictors of depressive symptom, which corresponds well with the result of the earlier study conducted by Ng et al.\textsuperscript{12} The odds ratio continued to be significant after adjustment for sex, age, household income level and current smoking. Depressive symptom was significantly associated with the number of comorbid chronic diseases, and the odds ratio continued to be significant after adjustment for variables. Jung et al.\textsuperscript{23} have reported that there is a positive relationship between the number of comorbid chronic diseases and dependent ADL. Based on these results and ours, it is conceivable that dependent ADL may be indirectly associated with depressive symptom.

| Table 3. Riskfactors for depressive symptom: adjusted odds ratios from a multivariate logistic regression model. |
|---|---|---|---|
| Factors | Model 1* | Model 2† |
| | OR‡ | 95% CI‡ | OR‡ | 95% CI‡ |
| Activities of daily living | | | | |
| Independent | Reference | Reference | | |
| Dependent | 2.94 | 2.54—3.42 | 2.42 | 2.06—2.84 |
| No. of chronic disease§ | | | | |
| 0 | Reference | Reference | | |
| 1 | 1.48 | 1.28—1.71 | 1.40 | 1.20—1.63 |
| 2 | 2.13 | 1.77—2.57 | 1.72 | 1.41—2.10 |
| ≥ 3 | 3.35 | 2.61—4.30 | 2.60 | 1.99—3.39 |
| Stroke | No | Reference | Reference | |
| Yes | 0.36 | 0.08—1.64 | 0.47 | 0.11—2.03 |
| Arthritis | No | Reference | Reference | |
| Yes | 0.98 | 0.74—1.30 | 0.84 | 0.64—1.11 |

*Model 1: Crude odds ratio. †Model 2: Adjustment for sex, age, economic status and current smoking status. ‡Odds ratio (OR) and 95% confidence interval (CI) were estimated after adjustment using sample weights and survey year. §Chronic disease: stroke, hypertension, diabetes, ischemic heart disease, liver cirrhosis, arthritis, asthma and chronic renal failure.
Some studies have demonstrated that there are significant associations of depressive symptoms with sex, age, educational level, household income level or BMI, whereas others have not.\(^1\text{-}^4,13,14,24,25\) Compared to previous studies, depressive symptom in our study tended to increase without any statistical significance, probably due to a relatively small sample size as well as to differences in study subjects and assessment tools between previous studies and ours.

It is well known that chronic diseases are closely associated with depressive symptoms. In our study, comorbid stroke and arthritis were significantly associated with depressive symptoms in univariate analysis but not in multivariate analysis after adjustment for variables. Schane et al.\(^13\) have indicated that depressive symptoms are significantly associated with diabetes mellitus and arthritis, whereas Chavannes et al.\(^14\) have reported that the reverse is true. Further studies are needed to confirm these results.

In our study, it was found that depressive symptom was not significantly associated with current smoking status. Although it is unclear whether smoking induces depressive symptoms or whether depressive symptoms induce smoking, a positive association between smoking and depressive symptoms is well known.\(^13,26,27\) In addition, it is important to determine whether there is a positive association between depressive symptoms and smoking amounts in COPD patients.\(^28\) Some previous studies have pointed out that depressive symptoms are not significantly associated with current smoking.\(^12,14\) Based on the results of these studies and ours, it is thought that depressive symptoms are less strongly associated with smoking in COPD patients than in the general population. The differences in such associations between the previous studies and ours may be because 1) the subjects of our study were restricted to COPD patients, 2) the number of study subjects was relatively small, and 3) the definition of depressive symptoms of previous studies was different. Depressive symptom was not significantly associated with smoking amounts, intention of smoking cessation or experience with smoking cessation campaigns.

The results of this study are subjected to some limitations. First, because this was a cross-sectional study, there is a limit to interpretations of a cause-and-effect relationship between depressive symptoms and related variables. Second, because we analyzed data obtained from the 2005 and 2008 surveys which were conducted according to different criteria, the prevalence or incidence of COPD in either 2005 or 2008 may not have been accurately investigated. However, in multivariate analysis, we attempted to minimize the difference in prevalence or incidence of COPD by using dummy variables. Third, depressive symptoms were assessed by using the answer to a single question “Have you felt sad or desperate continuously for ≥ 2 weeks so as to disturb your everyday life?” Watkins et al.\(^29\) investigated the accuracy of a single question “Do you often sad or depressed?” in screening for depression in a cohort of patients after strokes and reported a sensitivity of 86%, a specificity of 78%, a positive predictability of 82% and a negative predictability of 82%. Fourth, the incidence of physician-diagnosed chronic disease adopted in our study seemed to be lower than the actual incidence. Furthermore, because we did not determine whether the chronic diseases were in an active state or not, their association with depressive symptoms seemed to be insignificant. Fifth limitation is that these results might not be representative of Korean COPD patients. In this study two different sample weights were used for the reflection of each yearly stratified sampling. Because integrative sample weight of two-year data set was not developed. Finally, the number of current smokers was only 47, which was insufficient to investigate the differences in depressive symptoms according to current smoking status or intention of smoking cessation.

This study provides the basis for managing COPD patients in a clinical setting by understanding the number and characteristics of COPD patients with depressive symptoms. The results of this study suggest that primary physicians should manage COPD patients in consideration of risk factors for depressive symptoms and recommend that they stop smoking, regardless of the presence of depressive symptoms. Additional controlled studies are warranted.

ACKNOWLEDGEMENTS

This work was supported by Grant from Inje University, 2009.

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