

Health-related quality of life and chronic obstructive pulmonary disease in North Carolina

¹David W. Brown, DSc, ^{2,4}Roy Pleasants, PharmD, ³Jill A. Ohar, MD, ⁴Monica Kraft, MD, ⁵James F. Donohue, MD.
⁶David M. Mannino, MD, ⁷Winston Liao, MSc, ⁸Harry Herrick, MSc.

¹Center for Disease Control and Prevention, Atlanta, Georgia, USA.

²Cambell University School of Pharmacy, Buies Creek, NC, USA.

³Wake Forest University School of Medicine, Bethesda, MD, USA.

⁴Duke University School of Medicine, Durham, North Carolina, USA.

⁵University of North Carolina at Chapel Hill School of Medicine, Chapel Hill, NC, USA.

⁶University of Kentucky College of Public Health, Lexington, Kentucky, USA.

⁷North Carolina Division of Public Health Asthma Program, Raleigh, NC, USA.

⁸North Carolina State Center for Health Statistics, Raleigh, NC, USA

Citation: Brown DW, Pleasants R, Ohar JA, Kraft M, Donohue JF, Mannino DM, Liao W, Herrick H. Health-related quality of life and chronic obstructive pulmonary disease in North Carolina. *North Am J Med Sci* 2010; 2: 60-65.

Doi: 10.4297/najms.2010.260

Availability: www.najms.org

ISSN: 1947 – 2714

Abstract

Background: Comparisons of health-related quality of life (HRQOL) between persons with chronic obstructive pulmonary disease (COPD) and adults in the general population are not well described. **Aims:** To examine associations between COPD and four measures of HRQOL in a population-based sample. **Patients & Methods:** These relationships were examined using data from 13,887 adults aged ≥ 18 years who participated in the 2007 Behavioral Risk Factor Surveillance System (BRFSS) conducted in North Carolina (NC). Logistic regression was used to obtain adjusted relative odds (aOR). **Results:** The age-adjusted prevalence of COPD among NC adults was 5.4% (standard error 0.27). Nearly half of adults with COPD reported fair/poor health compared with 15% of those without the condition (age-aOR, 5.5; 95% confidence interval [CI], 4.4 to 6.8). On average, adults with COPD reported twice as many unhealthy days (physical/mental) as those without the condition. The age-adjusted prevalence of ≥ 14 unhealthy days during the prior 30 days was 45% for adults with COPD and 17% for those without. The aOR of ≥ 14 unhealthy days was 1.7 (95% CI, 1.4 to 2.2) times greater among adults with COPD compared with those without. **Conclusions:** These results suggest COPD is independently associated with lower levels of HRQOL and reinforce the importance of preventing COPD and its complications through health education messages stressing efforts to reduce total personal exposure to tobacco smoke, occupational dusts and chemicals, and other indoor and outdoor air pollutants linked to COPD and early disease recognition. Our findings represent one of the few statewide efforts in the US and provide guidance for disease management and policy decision making.

Keywords: Health-related quality of life, chronic obstructive pulmonary disease, North Carolina, behavioral risk factor surveillance system, age-adjusted prevalence, adults, pollutants, tobacco, occupational dusts, chemicals, policy decision making.

Correspondence to: David W Brown, Centers for Disease Control and Prevention, 4770 Buford Highway NE (MS K67), Atlanta, Georgia, 30341 USA. Tel.: (770) 4885269, Fax: (770) 4885965. Email: dbrown6@cdc.gov.

Introduction

The Global Initiative for Chronic Obstructive Lung Disease (GOLD) defines COPD as “a disease state

characterized by airflow limitation that is not fully reversible. The airflow limitation is usually both progressive and associated with an abnormal inflammatory response of the lungs to noxious particles or gases” [1]. In

2006, COPD affected approximately 12 million US adults with nearly 24 million Americans having evidence of impaired lung function based on GOLD criteria [2, 3]. In 2005, approximately 125,000 persons aged ≥ 25 years in the United States died with COPD as the underlying cause, an increase of 8% from COPD deaths in 2000, with considerable variations in mortality rates across states [4].

Health-related quality of life (HRQOL) has evolved to include aspects of life that affect perceived physical, emotional, and social aspects of health and well-being, and it is a fundamental measure used to understand the health status of a population [5]. HRQOL is increasingly used as an outcome measure in clinical populations with COPD, but few studies have compared the HRQOL of persons with COPD with health of adults in the general population (i.e., population-based samples) to monitor the burden of COPD. In this study, we examined associations between COPD and HRQOL among a large sample of adults aged 18 years or older residing in North Carolina during 2007 using data from the Behavioral Risk Factor Surveillance System (BRFSS).

Patients and Methods

The BRFSS is a state-based surveillance system which collects data on many of the behaviors and conditions that place adults (aged ≥ 18 years) at risk for chronic disease. Trained interviewers collect data on a monthly basis using an independent probability sample of households with landline telephones from the noninstitutionalized North Carolina population. In 2007, complete survey data were collected for 13,887 persons in North Carolina (partial complete surveys totaled 890). The Council for American Survey Research Organizations (CASRO) response rate for North Carolina in 2007 was 55%, and the cooperation rate (i.e., the proportion of all respondents interviewed of all eligible units in which a respondent was selected and actually contacted) was 75% [6]. Although response rates have declined for BRFSS, as well as for other telephone and personal interview surveys globally, research suggests little bias as a result of the nonresponse rate at this point-in-time with BRFSS estimates paralleling those of other national surveys in the US [7,8,9]. A detailed description of the survey's design and random sampling procedures is available [10]. The BRFSS has been approved as exempt research by the Centers for Disease Control and Prevention's institutional review board.

COPD was defined by an affirmative response to the question, "Have you ever been told by a doctor or health professional that you have chronic obstructive pulmonary disease (COPD), emphysema or chronic bronchitis?" The crude prevalence of COPD was greater among those excluded from the analysis (9.4% [standard error, 1.03] vs. 5.6% (0.27)).

All survey respondents were also asked four questions related to their health status or HRQOL: (1) "Would you say that in general your health is excellent, very good, good, fair, or poor?"; (2) "Now thinking about your

physical health, which includes physical illness and injury, for how many days during the past 30 days was your physical health not good?"; (3) "Now thinking about your mental health, which includes stress, depression, and problems with emotions, for how many days during the past 30 days was your mental health not good?"; and (4) "During the past 30 days, for about how many days did poor physical or mental health keep you from doing your usual activities, such as self-care, work, or recreation?" Respondents were not asked for specific underlying reasons of any reported unhealthy days. These questions and their validity and reliability are described elsewhere [5, 11-14]. We calculated overall unhealthy days as the sum of physically and mentally unhealthy days, not to exceed 30 days. We defined a dichotomous HRQOL variable as < 14 , ≥ 14 unhealthy days. A total of 14 unhealthy days is a meaningful cut point for those reporting substantially impaired HRQOL and corresponds to the upper 10% to 15% of the distribution for each healthy day's measure in the BRFSS [15]. With the exception of unhealthy mental days and activity limitation days, measures of HRQOL were lower among those excluded from the analysis (data not shown).

Prevalences were age-standardized to the 2000 US standard population [16]. We used logistic regression to obtain odds ratios (ORs) and 95% confidence intervals (CIs) adjusted for age (< 45 , 45 to 54, 55 to 64, 65 to 74, ≥ 75), gender, race/ethnicity (white, black, other, Hispanic), veteran status (yes/no), education ($<$ high school, high school, some college, college graduate), employment status (employed, self-employed, unemployed, not able to work, other), income, health insurance (yes/no), time since last routine checkup (< 12 months, 12 to 23 months, 24 to 59 months, ≥ 60 months, never), smoking status (current, former, never), body mass index (< 18.5 , 18.5 to 24.9, 25.0 to 29.9, ≥ 30.0 kg/m²), diabetes (yes/no), high blood cholesterol (yes/no), hypertension (yes/no), heart disease or stroke (yes/no), and asthma (yes/no). Confidence interval functions [17] are provided for associations between measures of health-related quality of life and COPD. The data were weighted to account for the age, race, and gender distribution in the state. We used SUDAAN 9.0 (RTI International, Research Triangle Park, NC) to account for the survey's complex sampling design.

Results

For this analysis, data were available for 11,878 persons aged 18 years or older who had complete information for study variables. Overall, 11% of adults were 18 to 24 years of age; 19%, 25 to 34; 20%, 35 to 44; 19%, 45 to 54; 15%, 55 to 64; 9%, 65 to 74; and 7%, 75 or older. Half (51%) of the sample were women; and 74% were white, 18% black, 8% other, and 7% Hispanic. Nearly 59% of adults had more than a high school education, nearly one-quarter were current smokers (23%), 29% were obese [body mass index (BMI) ≥ 30 kg/m²], 9% reported diabetes, and 9% had cardiovascular diseases. Persons excluded from the analysis were slightly younger and less likely to be men, of white race, non-Hispanic ethnicity, or a college graduate

compared with those included in the study.

The age-standardized prevalence of self-reported, physician-diagnosed COPD among adults aged ≥ 18 years was 5.4% (standard error, 0.27) (n=1198). The age-adjusted prevalence of COPD is shown in Table 1 by respondent characteristics including sociodemographics, access to care, and comorbid conditions. As expected, we observed an increased prevalence with increasing age. The slightly lower prevalence among those aged 75 years or older is not surprising since BRFSS is a point-in-time survey and those who survive to age 75 and are able to complete the survey are more likely to be healthier overall (and therefore less likely to have COPD). The prevalence of COPD was greater among those with lower education levels (vs. college graduates), the unemployed or those unable to work (vs. employed persons), persons with lower income levels (vs. those with incomes $\geq \$75,000$), as well as among those who reported never visiting a doctor for a routine checkup (vs. those with a routine doctor visit within the past 12 months). Finally, the prevalence of COPD was greater among persons with selected comorbid conditions including asthma, diabetes, hypertension, hypercholesterolemia, and cardiovascular diseases than for those without these conditions (Table 1).

The age-standardized prevalences of 14 or more unhealthy days during the previous 30-day period and fair or poor health status by COPD status are shown in Figure 1. For each HRQOL measure, the prevalence of 14 or more unhealthy days and fair or poor health was greater among persons with COPD compared with those without.

We examined the relationship between the number of unhealthy days during the previous 30-day period and COPD. Overall, the mean (standard error) number of unhealthy days (physical or mental) for all adults was 6.1 (0.13); 51% of respondents reported no unhealthy days (physical or mental). On average, adults with COPD reported more than twice as many physically or mentally impaired days (13.7 [0.68] vs. 5.7 [0.14]). After age adjustment, adults with COPD had 8 (95% CI, 6.6 to 9.3) more unhealthy days (physical or mental) on average than adults without COPD.

We estimated the relative odds of reporting 14 or more unhealthy days comparing adults with COPD with those without after multivariable adjustment (Fig. 2). Adults with COPD were more likely to have lower levels of HRQOL for each of the 4 unhealthy day measures compared with adults without COPD. For example, the relative odds of 14 or more unhealthy (physical or mental) days were 1.7 (95% CI, 1.4 to 2.2) times greater among adults with COPD than among those without after multivariable adjustment. Compared with those without the condition, persons with COPD were 2.8 (95% CI, 2.1 to 3.7) times more likely to report fair or poor health after multivariable adjustment (age-adjusted OR, 5.5; 95% CI, 4.4 to 6.8).

Table 1 Prevalence of self-reported, physician-diagnosed COPD by respondent characteristics, North Carolina, 2007, Behavioral Risk Factor Surveillance System.

Respondent Characteristic	Age-Adjusted* Prevalence of COPD % (standard error)	aOR (95% CI)†
Overall	5.4 (0.27)	
Age, years		
< 45	2.5 (0.40)	1.0 (referent)
45–54	5.5 (0.55)	2.5 (1.7–3.6)
55–64	8.5 (0.66)	4.0 (2.8–5.7)
65–74	13.1 (0.94)	5.7 (4.0–8.1)
≥ 75	11.1 (1.03)	4.3 (3.0–6.4)
Sex		
Men	5.3 (0.46)	1.0 (referent)
Women	5.5 (0.29)	1.1 (0.9–1.3)
Race		
White	5.5 (0.31)	1.0 (referent)
Black	4.2 (0.55)	0.6 (0.4–0.8)
Other	7.8 (1.90)	1.1 (0.6–1.9)
Hispanic ethnicity		
No	5.2 (0.26)	1.0 (referent)
Yes	8.6 (1.93)	1.2 (0.7–2.7)
Education		
Less than high school	10.9 (1.11)	4.0 (2.9–5.6)
High school	6.1 (0.51)	2.2 (1.7–2.9)
Some college	5.3 (0.61)	1.9 (1.4–2.6)
College graduate	3.1 (0.30)	1.0 (referent)
Veteran		
No	5.5 (0.30)	1.0 (referent)
Yes	4.5 (0.55)	1.0 (0.8–1.4)

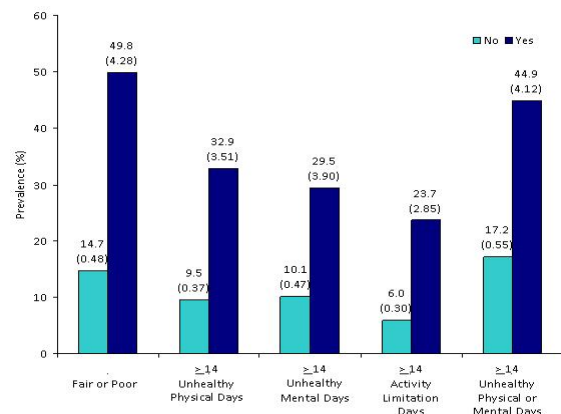
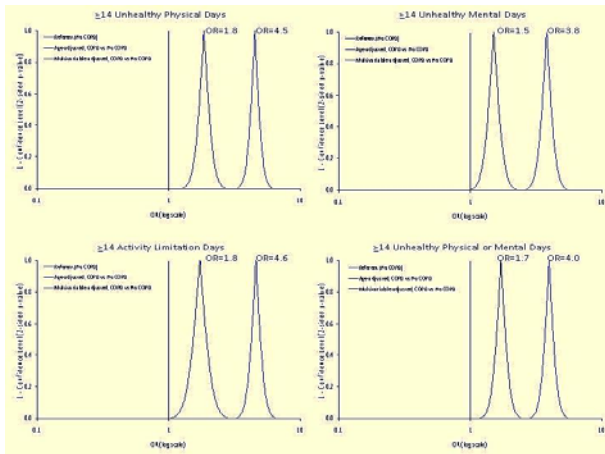


Fig. 1 Age-adjusted prevalence of fair or poor health and unhealthy days by COPD status, North Carolina, 2007, Behavioral Risk Factor Surveillance System.



COPD: Chronic obstructive pulmonary disease, OR: Odds ratio, CI: Confidence interval, *Multivariable-adjusted models adjusted for all variables shown in Table 1.

Fig. 2 Confidence Interval Functions for Associations Between Measures of Health- Related Quality of Life and COPD, North Carolina, 2007, Behavioral Risk Factor Surveillance System.

Discussion

COPD is a growing public health problem. According to projections, COPD is predicted to become the third leading cause of death worldwide by 2030 [18]. In this population-based, cross-sectional study, we observed that adults with COPD have lower levels of HRQOL than those without the condition. After age adjustment, adults with COPD reported 8 additional days of impaired physical or mental health during the previous 30 days than adults without COPD. Furthermore, adults with COPD were 70% more likely to report 14 or more unhealthy days (physical or mental) during the previous 30 days.

The results of this analysis are subject to some limitations. These data are cross-sectional; therefore, determinations of cause-and-effect are not possible. BRFSS is a telephone-based survey; therefore, persons of low socioeconomic status or those who are institutionalized are less likely to have a telephone and be included in BRFSS. Recent research also identifies differences between persons who only maintain a cell phone, and therefore are not included in the BRFSS, compared with persons who maintain a household landline [19]. Nonresponse is always a concern in survey research with regards to the possible introduction of bias; findings of others suggest that low response rates in the BRFSS do not appear to bias estimates at the national level [7-9].

Severity of COPD is not taken into account in this analysis. Also, because BRFSS interviews only noninstitutionalized persons, persons with COPD in this study may have less severe disease and/or comorbid conditions than the total COPD population in North Carolina. Data are self-reported; it is unclear how well self-reported COPD reflects true presence of disease. Also, the HRQOL measures used in this study are global measures; research comparing global with disease-specific HRQOL measures may provide different results.

Considering these limitations, the results of this study are consistent with prospective population-based studies. The Third US National Health and Nutritional Examination Survey reported a 7% prevalence of diagnosed COPD [20], and a meta-analysis of COPD epidemiological studies showed the overall prevalence of COPD to be 7.6% [21]. We also found a higher prevalence of COPD in women than men, consistent with other studies [22].

The relationship between COPD and HRQOL is complex and not well understood. Our findings of poor HRQOL among adults with COPD are consistent with previous studies, although we were able to identify only a few studies comparing HRQOL between persons with and without COPD or in population-based samples. In a study of adults aged >65 years, Peruzza and associates observed substantial impairment in quality of life measured by the Saint George Respiratory Questionnaire among 60 men with COPD (diagnosed based on European Respiratory Society criteria for respiratory functional impairment) compared with 58 men without COPD who were recruited from patients seen for a routine clinical examination [23]. In a population based sample of 2300 adults from the Hordaland County Cohort Study in Norway, Voll-Aanerud and colleagues found strong inverse associations between physical and mental quality of life (measured by the SF-12[®] Health Survey [SF-12]) and the number of respiratory symptoms as well as with presence of COPD or impaired lung function (measured by spirometry and classified according to GOLD criteria) [24]. Most studies have examined measures of HRQOL among patients with COPD and examined predictors for poor HRQOL levels, such as presence of acute exacerbations, level of dyspnea, and select medications [25-48].

Associations between COPD and lower levels of HRQOL are not surprising, as proper management of COPD often requires individuals to make extensive lifestyle changes. These changes may involve physical or behavioral adjustments, such as modifications in smoking behavior, physical activity, or prescription therapy, and may be accompanied by psychological consequences including depression and treatment-related frustration or emotional distress [49]. At the same time, persons with COPD who are better able to manage their disease may report higher levels of HRQOL due to fewer acute exacerbations or COPD-related complications. Also, whether COPD has a greater impact on either physical or emotional dimensions of HRQOL is unclear. Population-based studies with prospective designs will be helpful to assess the COPD outcomes. In our study, COPD was associated with perceived general health status and measures of impaired physical health or functioning as well as with impaired mental health.

Conclusion

We observed that self-reported, physician-diagnosed COPD was associated with lower levels of HRQOL compared with persons without COPD in a population-based sample of adults. COPD is a serious lung

disease that is treatable. Early diagnosis is important. Persons at risk for COPD who have cough, sputum production, or shortness of breath should talk with their physicians and be tested for the disease using spirometry, a simple breathing test for assessing lung function.

* This material was presented in part at the 2009 American Thoracic Society International Conference, San Diego, California.

Acknowledgement

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention. There is no conflict of interest.

References

- World Health Organization. Global strategy for diagnosis, management, and prevention of COPD. Geneva: World Health Organization, 2006. (Accessed January 13, 2010, at <http://www.goldcopd.com/GuidelinesResources.asp?l1=2&l2=0>)
- American Lung Association. COPD Fact Sheet. (Accessed January 13, 2010, at <http://www.lungusa.org>)
- Mannino DM, Braman S. The epidemiology and economics of chronic obstructive pulmonary disease. *Proc Am Thorac Soc* 2007; 4:502–506.
- Centers for Disease Control and Prevention. Deaths from chronic obstructive pulmonary disease—United States, 2000–2005. *MMWR Morb Mortal Wkly Rep* 2008; 57:1229–1232.
- Centers for Disease Control and Prevention. Measuring healthy days—population assessment of health-related quality of life. Atlanta, GA: US Centers for Disease Control and Prevention, 2000
- Centers for Disease Control and Prevention 2007 Behavioral Risk Factor Surveillance System Summary Data Quality Report. (Accessed January 13, 2010, at http://www.cdc.gov/brfss/technical_infodata/quality)
- Nelson DE, Powell-Griner E, Town M, Kovar MG. A comparison of national estimates from the National Health Interview Survey and the Behavioral Risk Factor Surveillance System. *Am J Public Health* 2003; 93:1335–1341.
- Fahimi M, Link M, Mokdad A, Schwartz DA, Levy P. Tracking chronic disease and risk behavior prevalence as survey participation declines: statistics from the behavioral risk factor surveillance system and other national surveys. *Prev Chronic Dis* 2008; 5:A80
- Mokdad AH. The Behavioral Risk Factors Surveillance System: past, present, and future. *Annu Rev Public Health* 2009; 30:43–54.
- Centers for Disease Control and Prevention. Behavioral Risk Factor Surveillance System User's Guide. Atlanta, GA: US Centers for Disease Control and Prevention, 1998, 30–43.
- Ôunpuu S, Chambers LW, Chan D, et al. Validity of the US Behavioral Risk Factor Surveillance System's health related quality of life survey tool in a group of older Canadians. *Chronic Dis Canada* 2001; 22:931–1001.
- Moriarty DG, Zack MM, Kobau R. The Centers for Disease Control and Prevention's Healthy Days Measures—Population tracking of perceived physical and mental health over time. *Health Qual Life Outcomes* 2003; 1:37.
- Andresen EM, Catlin TK, Wyrwich KW, et al. Retest reliability of surveillance questions on health related quality of life. *J Epidemiol Community Health* 2003; 57:339–343.
- Ford ES, Moriarty DG, Zack MM, et al. Self-reported body mass index and health-related quality of life: findings from the Behavioral Risk Factor Surveillance System. *Obes Res* 2001; 9:21–31.
- Brown DW, Balluz LS, Heath GW, et al. Associations between recommended levels of physical activity and health-related quality of life. Findings from the 2001 Behavioral Risk Factor Surveillance System. *Prev Med* 2003; 37:520–528.
- Klein RJ, Schoenborn CA. Age adjustment using the 2000 projected US population. *Healthy People 2000. Stat Notes* 2001: 1–9.
- Sullivan KM, Foster DA. Use of the confidence interval function. *Epidemiology* 1990; 1:39–42.
- World Health Organization World health statistics 2008. Geneva, Switzerland: WHO Press, 2008
- Link MW, Battaglia MP, Frankel MR, et al. Reaching the US cell phone generation: comparison of cell phone survey results with an ongoing landline telephone survey. *Public Opinion Quarterly* 2007; 71:814–839.
- Mannino DM, Gagnon RC, Petty TL, et al. Obstructive lung disease and lung function in adults in the United States. Data from the National Health and Nutrition Examination Survey. 1988–1994. *Arch Intern Med* 2000; 160:1683–1689.
- Halbert RJ, Natoli JL, Gano A, et al. Global burden of COPD. Systematic review and meta-analysis. *Eur Resp J* 2006; 28:523–532.
- Buist AS, McBurnie MA, Vollmer WM, et al. International variation in the prevalence of COPD; The BOLD Study. A population-based prevalence study. *Lancet* 2007; 370:741–750.
- Peruzza S, Sergi G, Vianello A, et al. Chronic obstructive pulmonary disease (COPD) in elderly subjects: impact on functional status and quality of life. *Respir Med* 2003; 97:612–617.
- Voll-Aanerud M, Eagan TM, Wentzel-Larsen T, et al. Respiratory symptoms, COPD severity, and health related quality of life in a general population sample. *Respir Med* 2008; 102:399–406.
- Wyrwich KW, Fihn SD, Tierney WM, et al. Clinically important changes in health-related quality of life for patients with chronic obstructive pulmonary disease: an expert consensus panel report.

- J Gen Intern Med 2003; 18:196–202.
26. Wijnhoven HA, Kriegsman DM, Hesselink AE, et al. The influence of co-morbidity on health-related quality of life in asthma and COPD patients. *Respir Med* 2003; 97:468–475.
 27. Katsura H, Yamada K, Kida K. Both generic and disease specific health-related quality of life are deteriorated in patients with underweight COPD. *Respir Med* 2005; 99:624–630.
 28. Arnardóttir RH, Sörensen S, Ringqvist I, et al. Two different training programmes for patients with COPD: a randomised study with 1-year follow-up. *Respir Med* 2006; 100:130–139.
 29. Rodrigue JR, Baz MA, Kanasky WF Jr, MacNaughton KL. Does lung transplantation improve health-related quality of life? The University of Florida experience. *J Heart Lung Transplant* 2005; 24:755–763.
 30. Milstone A, Patsimas J, Farzan D, Castaldo R, Singh H, Feurer I, Harnett J, Luke DR; the PROPeR Use Study Group. Prospective observational study of patient-reported outcomes for azithromycin versus usual care in the treatment of bacterial acute exacerbation of chronic bronchitis. *Clin Ther* 2005; 27:926–939.
 31. Katsura H, Yamada K, Wakabayashi R, et al. The impact of dyspnoea and leg fatigue during exercise on health-related quality of life in patients with COPD. *Respirology* 2005; 10:485–490.
 32. Hu J, Meek P. Health-related quality of life in individuals with chronic obstructive pulmonary disease. *Heart Lung* 2005; 34:415–422.
 33. Nishiyama O, Taniguchi H, Kondoh Y, et al. Factors in maintaining long-term improvements in health-related quality of life after pulmonary rehabilitation for COPD. *Qual Life Res* 2005; 14:2315–2321.
 34. Carrasco Garrido P, de Miguel Díez J, Rejas Gutiérrez J, et al. Negative impact of chronic obstructive pulmonary disease on the health-related quality of life of patients. Results of the EPIDEPOC study. *Health Qual Life Outcomes* 2006; 4:31.
 35. Skumlien S, Haave E, Morland L, et al. Gender differences in the performance of activities of daily living among patients with chronic obstructive pulmonary disease. *Chron Respir Dis* 2006; 3:141–148.
 36. Ries AL. Impact of chronic obstructive pulmonary disease on quality of life: the role of dyspnea. *Am J Med* 2006; 119:12–20.
 37. Reardon JZ, Lareau SC, ZuWallack R. Functional status and quality of life in chronic obstructive pulmonary disease. *Am J Med* 2006; 119:32–37.
 38. Omata M, Wakabayashi R, Kudoh S, et al. Correlation between bronchodilator responsiveness and quality of life in chronic obstructive pulmonary disease. *Allergol Int* 2007; 56:15–22.
 39. Katsura H, Yamada K, Wakabayashi R, et al. Gender-associated differences in dyspnoea and health-related quality of life in patients with chronic obstructive pulmonary disease. *Respirology* 2007; 12:427–432.
 40. Medinas-Amorós M, Alorda C, Renom F, et al. Quality of life in patients with chronic obstructive pulmonary disease: the predictive validity of the BODE index. *Chron Respir Dis* 2008; 5:7–11.
 41. Shioya T, Satake M, Sato K, et al. Long-term effect of the beta2-receptor agonist procaterol on daily life performance and exercise capacity in patients with stable chronic obstructive pulmonary disease. Clinical study with special reference to health-related quality of life and activities of daily living. *Arzneimittelforschung* 2008; 58:24–28.
 42. Duiverman ML, Wempe JB, Bladder G, et al. Health-related quality of life in COPD patients with chronic respiratory failure. *Eur Respir J* 2008; 32:379–386.
 43. Ozalevli S, Karaali H, Cankurtaran F, et al. Comparison of Short Form-36 Health Survey and Nottingham Health Profile in moderate to severe patients with COPD. *J Eval Clin Pract* 2008; 14:493–499.
 44. Tonnel AB, Perez T, Grosbois JM, et al. Effect of tiotropium on health-related quality of life as a primary efficacy endpoint in COPD. *Int J Chron Obstruct Pulmon Dis* 2008; 3:301–310.
 45. Donesky-Cuenco D, Nguyen HQ, Paul S, et al. Yoga therapy decreases dyspnea-related distress and improves functional performance in people with chronic obstructive pulmonary disease: a pilot study. *J Altern Complement Med* 2009; 15:225–234.
 46. Talley CH, Wicks MN. A pilot study of the self-reported quality of life for patients with chronic obstructive pulmonary disease. *Heart Lung* 2009; 38:141–150.
 47. Antoniu SA. UPLIFT Study: the effects of long-term therapy with inhaled tiotropium in chronic obstructive pulmonary disease. Evaluation of: Tashkin DP, Celli B, Senn S et al.: a 4-year trial of tiotropium in chronic obstructive pulmonary disease. *N Engl J Med* 2008; 359(15):1543–1554. *Expert Opin Pharmacother* 2009; 10:719–722.
 48. Habraken JM, Ter Riet G, Gore JM, et al. Health-Related Quality of Life in End-Stage COPD and Lung Cancer Patients. *J Pain Symptom Manage* 2009; 37:973–981.
 49. Weinger K, Jacobson AM. Psychosocial and quality of life correlates of glycemic control during intensive treatment of type 1 diabetes, *Patient Educ Couns* 2001; 42:123–131.