

REVIEW

Palliative care in COPD: an unmet area for quality improvement

Julia H Vermylen¹ Eytan Szmuilowicz² Ravi Kalhan³

¹Department of Medicine, ²Section of Palliative Medicine, Department of Medicine, ³Asthma and COPD Program, Department of Medicine, Northwestern University Feinberg School of Medicine, Chicago, IL, USA **Abstract:** COPD is a leading cause of morbidity and mortality worldwide. Patients suffer from refractory breathlessness, unrecognized anxiety and depression, and decreased quality of life. Palliative care improves symptom management, patient reported health-related quality of life, cost savings, and mortality though the majority of patients with COPD die without access to palliative care. There are many barriers to providing palliative care to patients with COPD including the difficulty in prognosticating a patient's course causing referrals to occur late in a patient's disease. Additionally, physicians avoid conversations about advance care planning due to unique communication barriers present with patients with COPD. Lastly, many health systems are not set up to provide trained palliative care physicians to patients with chronic disease including COPD. This review analyzes the above challenges, the available data regarding palliative care applied to the COPD population, and proposes an alternative approach to address the unmet needs of patients with COPD with proactive primary palliative care.

Keywords: advance care planning, primary palliative care, end-of-life care, advanced lung disease, quality of life, prognosis

Background

COPD is a complex, chronic disease defined by lung function limitation on spirometric testing that despite several recent breakthroughs in treatments remains an incurable and life-altering disease. 1,2 Worldwide, 210 million people suffer from COPD,3 and in 2002, death from COPD was estimated to be the fifth leading cause of death comprising approximately 7.5% of overall deaths. Unlike ischemic heart disease and cerebrovascular disease that are decreasing in prevalence, death from COPD is increasing. In the United States, for instance, age-standard death rates from COPD increased from 21.4 to 43.4 per 100,000, a change of over 100%, from 1970 to 2002. Conversely, age-standard death rates from stroke and heart disease decreased by 63.1% and 52.1% respectively over this same time frame. 5

Despite this large burden of disease, patients with COPD frequently have unmet needs at the end of life. ⁶⁻⁹ Although palliative care has been shown to improve symptom burden, quality of life, and patient satisfaction for patients across many diseases, ^{10–15} the majority of patients with COPD die without access to palliative care. ⁷ This review focuses on evaluating the current burden of disease, reviews what is known about palliative care and COPD, and argues that incorporation of palliative care into COPD management could improve overall quality of care delivered to people living with COPD.

Palliative care: what is it and what can it do?

Palliative care began in the 1960s with the beginning of the hospice movement and was focused on symptom management at the end of life. ¹⁶ Today, however, the World Health Organization (WHO) defines palliative care as:

Correspondence: Julia H Vermylen Northwestern University, 251 E Huron Street, Galter Pavilion Suite 3-150, Chicago, IL 60611, USA Tel +1 312 926 0008 Fax +1 312 926 4588 Email j-vermylen@northwestern.edu An approach that improves the quality of life of patients and their families facing the problem associated with lifethreatening illness, through the prevention and relief of suffering by means of early identification and impeccable assessment and treatment of pain and other problems, physical, psychosocial and spiritual.17

Using this new framework, palliative care now extends throughout the course of a patient's illness from disease onset through the terminal phase of illness.

Proactive approaches to palliative care involve a variety of care models. While some models include direct interactions with a trained palliative care physician, other models employ a multidisciplinary team of social workers, physicians, and nurses to provide intensive outpatient resources. All care models focus on improving the symptoms and psychosocial needs of the patient and also actively engage the family and caregivers in training and support as shown in Figure 1.

Outpatient palliative care improves patient quality of life, decreases symptom burden, and in some cases even improves mortality. 12,15,18,19 There are also potential significant financial benefits to the proactive use of palliative care. Kaiser Permanente, a US-based vertically organized health maintenance organization, found that the care for patients living with serious illness who received a multidisciplinary palliative intervention cost US\$6,580 less per patient in the last months of life, the equivalent of a 45% cost reduction in the intervention cohort compared to the baseline cohort.²⁰ Among patients with metastatic non-small-cell lung cancer (NSCLC), proactive palliative care has been shown to not only decrease costs but also improve mortality.¹² The findings within NSCLC are so profound that it is now considered standard of care to include early introduction of palliative care along-side more aggressive treatment.²¹ The American Thoracic Society²² and the recent Global Initiative for Obstructive Lung Disease statement² have called for increased palliative care services for patients living with COPD but at this time proactive palliative care for patients with COPD is rare and far from standard of care.

The burden of COPD

Not only is COPD a leading case of mortality, it is also a leading cause of disability throughout the world. In 2002 it was ranked as the eleventh leading cause of disability life adjusted years and is projected to increase to the seventh worldwide leading cause by 2030.4 In 2014, a systematic review of the indirect costs of COPD found that 13%-18% of individuals with COPD in the United States reported limitations in their ability to work.²¹

Across many studies of patients with COPD at the end of life, severe breathlessness is profoundly distressing to patients and is often poorly controlled and under-addressed by care providers. 6,12,23,24 In a cross-sectional study evaluating symptom burden in Sweden, approximately 90% of patients

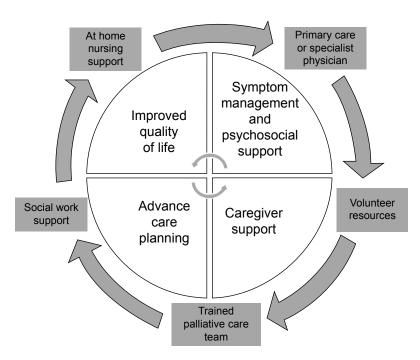


Figure 1 Structure and goals of outpatient palliative care services.

with stable lung function report shortness of breath.²³ Shortness of breath is not only a distressing symptom to patients but is also associated with decreased health-related quality of life (HRQoL) and a risk factor for further COPD exacerbations and functional decline.^{12,25}

Patients with COPD have a large burden of symptoms beyond breathlessness including fatigue, cough, and pain. Even among patients with minimal decrements in lung function, many report symptoms of cough, dry mouth, and fatigue.8 In the Swedish cross-sectional study, patients with moderate to severe airflow limitations were found to have on average 7.9 symptoms at any given time and patients with severe and very severe airflow limitations typically reported between nine and 14 symptoms.²³ In an international survey of 12 countries, 50% of countries reported that the average patient had severe to very severe burden of symptoms when scored with the COPD Assessment Test.²⁴ This burden of symptoms is similar to or worse than that reported in many diseases including NSCLC and heart failure though referral to palliative care subspecialty services is significantly reduced compared to these two conditions. 7,8,26,27

Unrecognized and uncontrolled anxiety and depression are major symptoms that impact quality of life in COPD. The prevalence of anxiety and depression among the COPD population varies greatly between studies though is consistently found to be increased compared to other chronic illnesses. ^{26,28} Depression is estimated to be as high as 40% among all stages of COPD and as high as 62% among patients who are prescribed domiciliary oxygen. ^{28,29} Anxiety is similarly found to be as high as 19% across patients with stable COPD though as high as 75% among patients with severe airflow limitation. ²⁶ Anxiety and depression are both associated with poor health outcomes including increased frequency of hospital admission, decreased quality of life, and early death. ²⁸

The burden of significant symptoms and comorbid health conditions in COPD are comparable or worse than other chronic illnesses including heart failure, HIV, and metastatic cancer. Patients with COPD, however, are less likely to have adequate treatment of symptoms at the end of their life, are more likely to have a decreased HRQoL, and are significantly less likely to receive specialist palliative care referral. Patients with COPD are more likely to have aggressive care at the end of life including mechanical ventilation and stays in the intensive care unit compared to those with lung cancer. COPD patients are also more likely to receive cardiopulmonary resuscitation (CPR) at the end of life though evidence from the Study to Understand Prognosis

and Preferences for Outcomes and Treatments (SUPPORT) showed no difference in preferences for mechanical ventilation or wishes to receive CPR between COPD patients and those with lung cancer.³⁰ Patients with COPD, therefore, are frequently under-supported with significant symptom burden, disability, quality of life impairment, and aggressive use of health care resources at the end of their life.

The current state of COPD and palliative care

Despite refractory symptoms and recurrent hospitalizations, patients with COPD die without access to palliative care. Many physicians view palliative care only as care given to actively dying individuals rather than a multidisciplinary approach that focuses on a patient's quality of life and burden of symptoms as well as a means to provide psychosocial support for patients and their caregivers. Physicians have difficulty predicting which patients are at highest risk of dying and have difficulty discussing these concerns with their patients. Additionally, the current health systems are not designed to adequately reimburse palliative care treatments for patients with COPD. Consequently, patients with COPD are often not referred to palliative care recourses and have unmet needs at the end of life. ^{6,9}

This review focuses on the many barriers to providing palliative care including the difficulties of prognostication and communication as well as the obstacles within the current health systems. Although specific pharmacotherapy options for managing symptoms of COPD are outside the scope of this article, this review lays out a new framework for delivery of palliative care. Additionally, this review specifically discusses how to identify patients appropriate for palliative care as well as how to screen for and address the unmet needs of patients with COPD.

Barriers to providing palliative care to patients with COPD

Difficulties of prognosis

The variable and prolonged course of COPD patients makes prognostication difficult for both physicians, patients, and their caregivers, and makes addressing end-of-life goals difficult. Unlike the typical patient with metastatic cancer who classically has a predictable, steady decline after diagnosis, the natural history of COPD is typically protracted and heterogeneous. Often, patients are first diagnosed with COPD after developing breathlessness in the outpatient setting. The diagnosis is made with lung function testing though the

degree of lung function impairment is often not correlated to the patient's symptom burden or disease course. 2,31 Patients experience a continuous slow decline in lung function accompanied by a slow accumulation of symptoms punctuated by episodes of acute exacerbation and decline with only partial recovery.32

While patients often experience a slow decline in functional status they also suffer episodes of acute exacerbation that may require hospitalization. 33,34 Although patients are at high risk of death during and just following these exacerbations, patients will often survive many acute episodes before their time of death.³⁵ The episodic course of COPD makes the exact time of death often feel unexpected and abrupt for both families and physicians since the patient had survived seemingly similar episodes in the past. The SUPPORT study reinforced this finding as it showed that 5 days before death, physicians estimated that their patients with COPD were more than 50% likely to be alive in 6 months.³⁶ Patients with lung cancer, in the same study however, were predicted to be less than 10% likely to be alive in 6 months.³⁶

The difficulty in prognostication causes both patients and physicians to struggle to recognize the terminal phase of illness. For this reason, approaches that limit palliative care resources to the final stages of life overlook many dying patients as their terminal phase of illness goes unrecognized. For this reason, an approach whereby COPD health care providers adopt proactive palliative care that does not rely on the precise prognostication and identification of imminent death in COPD would likely result in greater access to palliative care services for people living with COPD.

Difficulties with communication

Communication is often compromised between patients with COPD and their physicians. Due to the difficulty of prognostication, physicians are often hesitant to discuss their patient's disease course and expected course of illness.^{37,38} Physicians also report concern about removing hope for their patient and fear that this will have detrimental effects on their patient's quality of life (QoL) and mortality.³⁹ Even among physicians who feel their patients would benefit from goals of care conversations, many report inadequate training and insufficient time to approach these conversations. 40-42

Patients are also reluctant to discuss the end of their life. Commonly cited barriers include not knowing which physician they should speak to about their wishes, not knowing the extent of their own wishes, as well as wanting to focus on curative treatments instead of thinking about COPD as a life-limiting illness. In a qualitative study of patients with

COPD, many were quoted saying "I would rather concentrate on staying alive than talk about death". 39 In review of the Phase II intervention of the SUPPORT trial, only 23% of patients had discussed their wishes regarding CPR before hospitalization. Among patients who had not discussed their preferences, only 42% were interested in doing so leaving a majority of patients, approximately 58%, uninterested in having a conversation regarding CPR preferences. Interestingly, however, 87% of patients who had not and did not want to discuss their preferences wanted to avoid prolonged mechanical ventilation and 25% wanted to avoid resuscitation completely. Additionally, 80% of patients who did not want to discuss their end-of-life wishes also had not completed advanced directives.⁴³ Although patients may not initially be interested in discussing advanced directives with their physicians, this study suggests that many patients still have unexpressed wishes that may not be respected if the conversation is not broached delicately.9,43

Despite the many barriers to communication from both the physician and patient perspective, patients express the desire to have emotional support and discuss end-of-life care with their physicians. 44 In particular COPD patients report particular deficits in discussing prognosis as well as the dying process with their physicians.³⁷

Health-systems issues

There is great variability in access to palliative care worldwide. The UK for instance, the founder of the Hospice movement, is rather progressive with one of the highest services to population ratios of 15 services (defined as hospital units, palliative care teams, home care teams etc) per million inhabitants.⁴⁵ In other parts of the world, however, such as Sub-Saharan Africa, palliative care is thought to be a luxury and not a basic right of an individual and there are no national palliative care services available. 46 Even in areas of the world that have well-defined palliative care resources, many are limited to individuals in the terminal phase of illness or limited to particular diagnoses such as cancer, 47 making a proactive palliative care approach in a chronic medical condition like COPD challenging. In the USA, current reimbursement structures are set up to fund curative treatment or terminal illness. There are not, however, consistent systems in place to reimburse providers for palliative treatment throughout the course of a patient's illness or to support the holistic and psychosocial aspects of a patient's care. Consequently, even if patients and physicians are discussing symptom management, improved home recourses, support for primary caregiver, and long-term care planning, they are

often unable to be reimbursed for these extensive recourses and thus are unable to address these unmet needs.⁴⁸

Palliative care and COPD

Identifying patients appropriate for referral to palliative care: the surprise question

Prognostication of the exact time of death will always be difficult for the clinician and the patient with COPD. When palliative care is included not only during the terminal phase of illness but also during the last few stages of disease, the difficulties of prognostication are often alleviated. One proposed method for identifying patients appropriate for proactive palliative care is the use of "the surprise question". This method, encourages physicians to ask "would I be surprised if my patient were to die in the next 12 months?"49 If the answer is no, proactive palliative care should be considered. 49,50 The "surprise question" has been adopted by the UK national policies to be used as a supplementary screening tool for primary care physicians to identify patients who would benefit from early palliative care.51 It has also been studied in other diseases including end-stage renal disease and has been shown to aid in the prediction of short-term mortality risk in hemodialysis patients.52

Twelve-month predictors of mortality

Many studies have looked into predictors of 12-month mortality as well as predictors of mortality post-hospitalization for an acute exacerbation of COPD. Consistently authors report that patients with advanced age, cardiac comorbidity, low forced expiratory volume in 1 second, low body mass index, poor functional status, and poor HRQoL were at increased risk of dying within the following year. 35,53-56 Although each individual's mortality will vary, Table 1 identifies flags to recognize patients that would benefit from proactive supportive palliative care interventions.

Refractory symptoms

In addition to those patients with high risk of death within the next year of life, patients with symptoms that are refractory to initial therapy would benefit from an early palliative care approach. As discussed earlier, patients with COPD have a multitude of symptoms ranging from dyspnea fatigue, cachexia, anxiety, and depression. Although some patients may respond quickly to pharmacotherapy, opioids, and oxygen therapy, others continue to suffer after initial steps

Table I Triggers to begin or intensify proactive palliative resources for patients with COPD

Age $\geq 75^{35,54-56,61}$

Comorbidities^{a,35,54,55,61–66}

Change 6-minute walk by 50 m⁵³

Functional dependence and patient reported minimal physical bactivity 35,53,65,67

Poor health-related quality of life^{c,35,57,65,68}

 $\text{FEV}_{_{1}} < 30\%^{9,54,66}$

BMI < 20%61,69

≥ I hospitalization within last year^{55,65}

Notes: a Comorbidities include diabetes, cardiovascular disease (ischemic heart disease, congestive heart failure), end-stage chronic renal disease. Patient reported minimal physical activity defined as spending more time in a recliner chair or bed than not during the day. Methods to assess health-related quality of life include the St George's Respiratory Questionnaire, Chronic Respiratory Disease Questionnaire, and the Breathing Problems Questionnaire.

Abbreviations: FEV,, forced expiratory volume in 1 second; BMI, body mass index.

are taken. In these patients, intensive outpatient support that can be provided by a palliative care team should be considered.

New models of care

Though the focus of both inpatient and outpatient palliative care has been on oncology patients, there have been a few studies evaluating palliative care and holistic interventions within the COPD population.

A recent study published in December 2014 was a singleblind randomized controlled trial for patients with advanced disease and refractory breathlessness. In this study both patients with cancer and advanced chronic illness, including COPD, were enrolled and followed for a 6-week intervention. The support services involved a multidisciplinary team including pulmonologists, palliative care specialists, nurses, physical therapists, respiratory therapists, and occupational therapists. Patients were screened at routine clinic appointments and given a breathlessness pack including medications, a crisis plan, as well as a poem. Patients then received a home assessment 2–3 weeks after enrollment by a physical therapist and occupational therapist and a follow-up appointment with the pulmonologist and palliative care physician at 4 weeks after enrollment. Not surprisingly, patients in the breathlessness support arm of the trial had a significant reduction in their burden of symptoms. Surprisingly, unlike the cancer patients involved in the intervention who had no change in mortality, patients with COPD had a significant mortality benefit upon 6-month follow-up; 100% of COPD patients in the breathlessness support arm were alive at 6 months whereas only 79% were found to be alive in the control cohort.⁵⁷ Similar mortality benefits were found for all non-oncologic patients in this cohort. This study is the first randomized controlled trial addressing breathlessness through a proactive palliative method in the outpatient setting and suggests that this will not only treat the symptoms of advanced illness but potentially also improve the mortality of patients with COPD.⁵⁷

Another effective tool has been the use pulmonary rehabilitation (PR). PR has been studied in the stable COPD population as well as in the post-hospitalization of acute exacerbation management where it has been found to be effective at reducing symptoms and re-hospitalization as well as improving HRQoL.58 In many studies PR has also been found to reduce symptoms of anxiety and depression by increasing the patient's functional status and allowing them to engage in their community.⁵⁹ Although PR is not typically classified as a palliative care measure, it is already addressing many of the unmet needs of COPD patients. 46 It is a model of care that already involves a multidisciplinary approach after a vulnerable time in a patient's illness. PR programs, however, do not typically assess patients for concurrent depression and anxiety and do not routinely engage patients in end-of-life discussions at this time. 48 With further resources such as scheduled palliative care visits during PR, this could be an ideal time and place to introduce palliative care to COPD patients.

Conclusion: where to go from here

In order to improve the suffering of end-stage COPD patients, palliative care needs to be incorporated earlier into the trajectory of illness. Currently patients and physicians are waiting until identification of the terminal phase of illness that is often indistinguishable from the slow chronic decline with repeated acute exacerbations characteristic of this disabling disease. Opportunities are missed to decrease disease associated disability and suffering for patients. Instead, patients are left with uncontrolled symptoms, disability, anxiety and depression, and repeated hospitalizations at the end of their life. Proactive palliative care must be incorporated to address the needs of patients during the last few years of life in order to prevent suffering and excess spending at the end of life.

Proactive palliative care should be incorporated early into the patient's illness as shown in Figure 2 to address complex refractory symptoms, advance care planning, and increase outpatient resources for patients and caregivers. Not all patients who would benefit from proactive palliative support need to be referred to palliative care physicians. Unfortunately given the health systems barriers discussed previously, this will not be feasible in the current system of care in many nations. Instead, primary care providers as well as specialty pulmonologists should be trained and capable of screening patients who would benefit from proactive outpatient supportive care and provide primary palliative care as outlined in Table 2. These frontline providers should provide increased focus on symptom management, advance care planning, and support for patients as part of comprehensive COPD care. Interdisciplinary teams including nursing, social work, respiratory, and physical therapists as well as chaplains or pastoral care should be engaged to meet the complex needs of their patients.

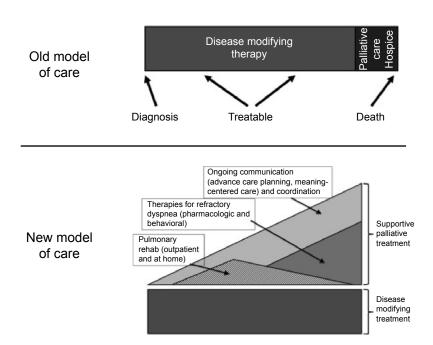


Figure 2 Proactive palliative care in COPD.

Table 2 Primary palliative care

Step 1: build a team, share the load Nurses Medical triage and support Social workers Psychosocial support, connecting patients to needed community resources Respiratory and physical therapists Exercise training and non-pharmacologic approaches to dyspnea relief Chaplains/pastoral care palliative care clinicians Spiritual support, legacy work complex symptom management, and additional support Step 2: screen for and address unmet needs Symptom management Consider opioids for refractory dyspnea and train patients/caregivers to use non-pharmacologic techniques to help decrease breathlessness (eg, positioning and use of personal fans) Depression and anxiety Routine screen given high prevalence Consider pharmacotherapy Early and regular discussions about future care "Given how things are going, what is most important to you?" and concerns "What are you most concerned about?" "Would it help to talk about what to expect or what to prepare for as your illness worsens?" "Hope for the best, but prepare for the worst" Advance care planning: identifying decision-makers "Who would speak for you and share your wishes with regards to medical decisions and clarifying limits around potential interventions if you were not able to speak for yourself?" "Are there treatments or interventions that you would not want as you approached the end of life?"

Further research is needed to assess the improvement in patients' symptoms, QoL and mortality as well as the proposed financial benefits of proactive palliative care in the treatment of COPD. Although we know patients with COPD are suffering and proactive palliative care has been used in other diseases, few research has been performed to prove the benefits of proactive palliative care in COPD. The PROLONG study is an ongoing prospective cluster controlled trial involving six hospitals in the Netherlands using proactive palliative care. This will be the first study of its kind to use a proposed set of clinical indicators to prompt proactive palliative care and will assess the effects on mortality and QoL of study participants. This is an exciting area of research that will provide new insight into the power of proactive palliative care in COPD.

Disclosure

JH Vermylen and E Szmuilowicz have no disclosures to report in this work. R Kalhan reports personal fees from Forest Laboratories, grants and personal fees from Boerhinger Ingelheim, grants from GlaxoSmithKline and from PneumRx, personal fees from Merck and from Quantia Communications, and grants from Spiration, outside the submitted work.

References

- Petty TL. The history of COPD. Int J Chron Obstruct Pulmon Dis. 2006; 1(1):3–14.
- goldcopd.org [homepage on the Internet]. Global strategy for the diagnosis, management and prevention of COPD. Global Initiative for Chronic Obstructive Lung Disease (GOLD); 2015. Available from: http://www.goldcopd.org/guidelines-global-strategy-for-diagnosis-management. html. Accessed July 12, 2015.

- World Health Organization. Prevention and Control of Noncommunicable Diseases: Guidelines for Primary Health Care in Low Resource Settings. Geneva: World Health Organization; 2012. Available from: http://apps.who.int/iris/bitstream/10665/76173/1/9789241548397_eng. pdf. Accessed July 12, 2015.
- Mathers CD, Loncar D. Projections of global mortality and burden of disease from 2002 to 2030. PLoS Med. 2006;3(11):e442.
- Jemal A, Ward E, Hao Y, Thun M. Trends in the leading causes of death in the United States, 1970–2002. JAMA. 2005;294(10):1255–1259.
- Schroedl CJ, Yount SE, Szmuilowicz E, Hutchison PJ, Rosenberg SR, Kalhan R. A qualitative study of unmet healthcare needs in chronic obstructive pulmonary disease. A potential role for specialist palliative care? *Ann Am Thorac Soc.* 2014;11(9):1433–1438.
- Gore JM, Brophy CJ, Greenstone MA. How well do we care for patients with end stage chronic obstructive pulmonary disease (COPD)?
 A comparison of palliative care and quality of life in COPD and lung cancer. *Thorax*. 2000;55(12):1000–1006.
- Walke LM, Byers AL, Tinetti ME, Dubin JA, McCorkle R, Fried TR. Range and severity of symptoms over time among older adults with chronic obstructive pulmonary disease and heart failure. *Arch Intern Med.* 2007;167(22):2503–2508.
- Curtis JR. Palliative and end-of-life care for patients with severe COPD. Eur Respir J. 2008;32(3):796–803.
- Brumley R, Enguidanos S, Jamison P, et al. Increased satisfaction with care and lower costs: results of a randomized trial of in-home palliative care. J Am Geriatr Soc. 2007;55(7):993–1000.
- 11. Teno JM, Clarridge BR, Casey V, et al. Family perspectives on end-of-life care at the last place of care. *JAMA*. 2004;291(1):88–93.
- 12. Temel JS, Greer JA, Muzikansky A, et al. Early palliative care for patients with metastatic non-small-cell lung cancer. *N Engl J Med*. 2010;363(8):733–742.
- Lautrette A, Darmon M, Megarbane B, et al. A communication strategy and brochure for relatives of patients dying in the ICU. N Engl J Med. 2007;356(5):469–478.
- Casarett D, Pickard A, Bailey FA, et al. Do palliative consultations improve patient outcomes? *J Am Geriatr Soc.* 2008;56(4):593–599.
- Bakitas M, Lyons KD, Hegel MT, et al. Effects of a palliative care intervention on clinical outcomes in patients with advanced cancer: the Project ENABLE II randomized controlled trial. *JAMA*. 2009; 302(7):741–749.

Vermylen et al Dovepress

 Saunders C. The evolution of palliative care. JR Soc Med. 2001;94(9): 430–432

- World Health Organization [homepage on the Internet]. WHO Definition of Palliative Care. World Health Organization; 2015. Available from: http://www.who.int/cancer/palliative/definition/en/. Accessed July 12, 2015
- Rabow M, Kvale E, Barbour L, et al. Moving upstream: a review of the evidence of the impact of outpatient palliative care. *J Palliat Med*. 2013; 16(12):1540–1549.
- Rabow MW, Dibble SL, Pantilat SZ, McPhee SJ. The comprehensive care team: a controlled trial of outpatient palliative medicine consultation. *Arch Intern Med.* 2004;164(1):83–91.
- Brumley RD, Enguidanos S, Cherin DA. Effectiveness of a homebased palliative care program for end-of-life. *J Palliat Med*. 2003;6(5): 715–724.
- Smith TJ, Temin S, Alesi ER, et al. American Society of Clinical Oncology provisional clinical opinion: the integration of palliative care into standard oncology care. *J Clin Oncol.* 2012;30(8):880–887.
- Lanken PN, Terry PB, Delisser HM, et al. An official American Thoracic Society clinical policy statement: palliative care for patients with respiratory diseases and critical illnesses. *Am J Respir Crit Care Med.* 2008;177(8):912–927.
- Greer JA, Pirl WF, Jackson VA, et al. Effect of early palliative care on chemotherapy use and end-of-life care in patients with metastatic non-small-cell lung cancer. *J Clin Oncol*. 2012;30(4):394–400.
- Rabow MW, Petersen J, Schanche K, Dibble SL, McPhee SJ. The comprehensive care team: a description of a controlled trial of care at the beginning of the end of life. *J Palliat Med*. 2003;6(3):489–499.
- Blinderman CD, Homel P, Billings JA, Tennstedt S, Portenoy RK. Symptom distress and quality of life in patients with advanced chronic obstructive pulmonary disease. *J Pain Symptom Manage*. 2009; 38(1):115–123.
- Solano JP, Gomes B, Higginson IJ. A comparison of symptom prevalence in far advanced cancer, AIDS, heart disease, chronic obstructive pulmonary disease and renal disease. *J Pain Symptom Manage*. 2006; 31(1):58–69.
- Edmonds P, Karlsen S, Khan S, Addington-Hall J. A comparison of the palliative care needs of patients dying from chronic respiratory diseases and lung cancer. *Palliat Med.* 2001;15(4):287–295.
- Maurer J, Rebbapragada V, Borson S, et al. Anxiety and depression in COPD: current understanding, unanswered questions, and research needs. *Chest.* 2008;134(4 Suppl):43S–56S.
- Lacasse Y, Rousseau L, Maltais F. Prevalence of depressive symptoms and depression in patients with severe oxygen-dependent chronic obstructive pulmonary disease. *J Cardiopulm Rehabil*. 2001;21(2):80–86.
- Claessens MT, Lynn J, Zhong Z, et al. Dying with lung cancer or chronic obstructive pulmonary disease: insights from SUPPORT. Study to Understand Prognoses and Preferences for Outcomes and Risks of Treatments. J Am Geriatr Soc. 2000;48(5 Suppl):S146–S153.
- Casanova C, de Torres JP, Aguirre-Jaime A, et al. The progression of chronic obstructive pulmonary disease is heterogeneous: the experience of the BODE cohort. Am J Respir Crit Care Med. 2011;184(9): 1015–1021.
- Murray SA, Kendall M, Boyd K, Sheikh A. Illness trajectories and palliative care. BMJ. 2005;330(7498):1007–1011.
- 33. Fletcher C, Peto R. The natural history of chronic airflow obstruction. *Br Med J.* 1977;1(6077):1645–1648.
- Sanchez-Salcedo P, Divo M, Casanova C, et al. Disease progression in young patients with COPD: rethinking the Fletcher and Peto model. Eur Respir J. 2014;44(2):324–331.
- Steer J, Gibson GJ, Bourke SC. Predicting outcomes following hospitalization for acute exacerbations of COPD. QJM. 2010;103(11): 817–829.
- No authors listed. A controlled trial to improve care for seriously ill
 hospitalized patients. The study to understand prognoses and preferences for outcomes and risks of treatments (SUPPORT). The SUPPORT
 Principal Investigators. *JAMA*. 1995;274(20):1591–1598.

 Curtis JR, Engelberg RA, Nielsen EL, Au DH, Patrick DL. Patientphysician communication about end-of-life care for patients with severe COPD. Eur Respir J. 2004;24(2):200–205.

- Christakis NA, Iwashyna TJ. Attitude and self-reported practice regarding prognostication in a national sample of internists. *Arch Intern Med.* 1998;158(21):2389–2395.
- Knauft E, Nielsen EL, Engelberg RA, Patrick DL, Curtis JR. Barriers and facilitators to end-of-life care communication for patients with COPD. *Chest*. 2005;127(6):2188–2196.
- Sullivan AM, Lakoma MD, Block SD. The status of medical education in end-of-life care: a national report. *J Gen Intern Med.* 2003;18(9): 685–695.
- 41. Elkington H, White P, Higgs R, Pettinari CJ. GPs' views of discussions of prognosis in severe COPD. *Fam Pract*. 2001;18(4):440–444.
- 42. Mulcahy P, Buetow S, Osman L, et al. GPs' attitudes to discussing prognosis in severe COPD: an Auckland (NZ) to London (UK) comparison. *Fam Pract*. 2005;22(5):538–540.
- 43. Hofmann JC, Wenger NS, Davis RB, et al. Patient preferences for communication with physicians about end-of-life decisions. SUP-PORT Investigators. Study to Understand Prognoses and Preference for Outcomes and Risks of Treatment. *Ann Intern Med.* 1997;127(1): 1–12.
- 44. Curtis JR, Wenrich MD, Carline JD, Shannon SE, Ambrozy DM, Ramsey PG. Patients' perspectives on physician skill in end-of-life care: differences between patients with COPD, cancer, and AIDS. *Chest*. 2002;122(1):356–362.
- Centeno C, Clark D, Lynch T, et al. Facts and indicators on palliative care development in 52 countries of the WHO European region: results of an EAPC Task Force. *Palliat Med*. 2007;21(6):463–471.
- Mwangi-Powell FN, Powell RA, Harding R. Models of delivering palliative and end-of-life care in sub-Saharan Africa: a narrative review of the evidence. *Curr Opin Support Palliat Care*. 2013;7(2):223–228.
- 47. Van Beek K, Woitha K, Ahmed N, et al. Comparison of legislation, regulations and national health strategies for palliative care in seven European countries (Results from the Europall Research Group): a descriptive study. BMC Health Serv Res. 2013;13:275.
- 48. Hardin KA, Meyers F, Louie S. Integrating palliative care in severe chronic obstructive lung disease. *COPD*. 2008;5(4):207–220.
- Murray SA, Boyd K, Sheikh A. Palliative care in chronic illness. *BMJ*. 2005;330(7492):611–612.
- Seamark DA, Seamark CJ, Halpin DM. Palliative care in chronic obstructive pulmonary disease: a review for clinicians. *J R Soc Med*. 2007;100(5):225–233.
- Murray S, Boyd K. Using the 'surprise question' can identify people with advanced heart failure and COPD who would benefit from a palliative care approach. *Palliat Med.* 2011;25(4):382.
- Da Silva Gane M, Braun A, Stott D, Wellsted D, Farrington K. How robust is the 'surprise question' in predicting short-term mortality risk in haemodialysis patients? *Nephron Clin Pract*. 2013;123(3–4): 185–193.
- 53. Benzo R, Siemion W, Novotny P, et al. Factors to inform clinicians about the end of life in severe chronic obstructive pulmonary disease. *J Pain Symptom Manage*. 2013;46(4):491–499.e4.
- Groenewegen KH, Schols AM, Wouters EF. Mortality and mortalityrelated factors after hospitalization for acute exacerbation of COPD. *Chest.* 2003;124(2):459–467.
- McGhan R, Radcliff T, Fish R, Sutherland ER, Welsh C, Make B. Predictors of rehospitalization and death after a severe exacerbation of COPD. *Chest.* 2007;132(6):1748–1755.
- Bustamante-Fermosel A, De Miguel-Yanes JM, Duffort-Falco M, Munoz J. Mortality-related factors after hospitalization for acute exacerbation of chronic obstructive pulmonary disease: the burden of clinical features. *Am J Emerg Med.* 2007;25(5):515–522.
- Higginson IJ, Bausewein C, Reilly CC, et al. An integrated palliative and respiratory care service for patients with advanced disease and refractory breathlessness: a randomised controlled trial. *Lancet Respir Med*. 2014;2(12):979–987.

- Puhan MA, Gimeno-Santos E, Scharplatz M, Troosters T, Walters EH, Steurer J. Pulmonary rehabilitation following exacerbations of chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2011;(10): CD005305.
- Paz-Diaz H, Montes de Oca M, Lopez JM, Celli BR. Pulmonary rehabilitation improves depression, anxiety, dyspnea and health status in patients with COPD. Am J Phys Med Rehabil. 2007;86(1):30–36.
- Duenk RG, Heijdra Y, Verhagen SC, Dekhuijzen RP, Vissers KC, Engels Y. PROLONG: a cluster controlled trial to examine identification of patients with COPD with poor prognosis and implementation of proactive palliative care. *BMC Pulm Med.* 2014;14:54.
- 61. Connors AF Jr, Dawson NV, Thomas C, et al. Outcomes following acute exacerbation of severe chronic obstructive lung disease. The SUPPORT investigators (Study to Understand Prognoses and Preferences for Outcomes and Risks of Treatments). Am J Respir Crit Care Med. 1996;154(4 Pt 1):959–967.
- Fruchter O, Yigla M. Cardiac troponin-I predicts long-term mortality in chronic obstructive pulmonary disease. COPD. 2009;6(3):155–161.
- 63. Ho TW, Tsai YJ, Ruan SY, et al. In-hospital and one-year mortality and their predictors in patients hospitalized for first-ever chronic obstructive pulmonary disease exacerbations: a nationwide population-based study. *PloS One*. 2014;9(12):e114866.

- Maters GA, de Voogd JN, Sanderman R, Wempe JB. Predictors of allcause mortality in patients with stable COPD: medical co-morbid conditions or high depressive symptoms. COPD. 2014;11(4):468–474.
- Almagro P, Calbo E, Ochoa de Echaguen A, et al. Mortality after hospitalization for COPD. Chest. 2002;121(5):1441–1448.
- Gudmundsson G, Gislason T, Lindberg E, et al. Mortality in COPD patients discharged from hospital: the role of treatment and co-morbidity. Respir Res. 2006;7:109.
- Waschki B, Kirsten A, Holz O, et al. Physical activity is the strongest predictor of all-cause mortality in patients with COPD: a prospective cohort study. *Chest*. 2011;140(2):331–342.
- Yohannes AM, Baldwin RC, Connolly MJ. Predictors of 1-year mortality in patients discharged from hospital following acute exacerbation of chronic obstructive pulmonary disease. *Age Ageing*. 2005;34(5): 491–496.
- Hallin R, Gudmundsson G, Suppli Ulrik C, et al. Nutritional status and long-term mortality in hospitalised patients with chronic obstructive pulmonary disease (COPD). Respir Med. 2007;101(9):1954–1960.

International Journal of COPD

Publish your work in this journal

The International Journal of COPD is an international, peer-reviewed journal of therapeutics and pharmacology focusing on concise rapid reporting of clinical studies and reviews in COPD. Special focus is given to the pathophysiological processes underlying the disease, intervention programs, patient focused education, and self management protocols.

This journal is indexed on PubMed Central, MedLine and CAS. The manuscript management system is completely online and includes a very quick and fair peer-review system, which is all easy to use. Visit http://www.dovepress.com/testimonials.php to read real quotes from published authors.

Submit your manuscript here: http://www.dovepress.com/international-journal-of-chronic-obstructive-pulmonary-disease-journal

