



**ORIGINAL ARTICLE** COPD

# Understanding variation in length of hospital stay for COPD exacerbation: **European COPD audit**

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ABSTRACT Chronic obstructive pulmonary disease (COPD) care across Europe has high heterogeneity with respect to cost and the services available. Variations in length of stay (LOS) may be attributed to patient characteristics, resource and organisational characteristics, and/or the so-called hospital cluster effect.

The European COPD Audit in 13 countries included data from 16018 hospitalised patients. The recorded variables included information on patient and disease characteristics, and resources available. Variables associated with LOS were evaluated by a multivariate, multilevel analysis.

Mean±sD LOS was 8.7±8.3 days (median 7 days, interquartile range 4-11 days). Crude variability between countries was reduced after accounting for clinical factors and the clustering effect. The main factors associated with LOS being longer than the median were related to disease or exacerbation severity, including GOLD class IV (OR 1.77) and use of mechanical ventilation (OR 2.15). Few individual resource variables were associated with LOS after accounting for the hospital cluster effect.

This study emphasises the importance of the patients' clinical severity at presentation in predicting LOS. Identifying patients at risk of a long hospital stay at admission and providing targeted interventions offers the potential to reduce LOS for these individuals. The complex interactions between factors and systems were more important that any single resource or organisational factor in determining differences in LOS between hospitals or countries.



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LOS determined by COPD exacerbation severity and interactions within hospitals constituting effective patient care http://ow.ly/XD9hy







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# Introduction

Chronic obstructive pulmonary disease (COPD) is a condition with high morbidity and cost to health services. Hospitalisations due to COPD negatively affect the well-being of patients with an impact on length and quality of life. Studies in different European countries have highlighted the cost of the disease and that of admissions. In a survey conducted in 1998–1999, annual direct expenditure for COPD treatment per patient in Europe ranged from  $\mathfrak{E}530$  in France to  $\mathfrak{E}3238$  in Spain [1] with a significant proportion of this cost due to inpatient treatment of exacerbations [1, 2]. Accordingly, several initiatives have been proposed to reduce the number of hospitalisations or to reduce the length of stay (LOS) from a cost-effectiveness and clinical perspective [3–5].

Various factors have been implicated in increasing LOS of hospital admissions, both clinical [6–10] and social [11]. European health systems have a large inter-hospital variability in terms of resources and the range of services offered [12, 13], and some previous reports have found a relationship between LOS and such factors [4, 14, 15], whilst others have not [16, 17]. To our knowledge, there has been no comprehensive analysis studying the effects of clinical, organisational and resource factors on LOS in European patients admitted with exacerbations of COPD across a range of European countries and healthcare systems.

The European COPD Audit was a European Respiratory Society (ERS)-funded project designed as a pilot study, performed in 2010–2011 to evaluate clinical practice variability as well as clinical and organisational factors related to outcomes for COPD hospital admissions across Europe, in which 432 hospitals from 13 European countries participated including 16018 cases of COPD exacerbations admitted to hospital.

In the present analysis, we aimed to determine the mean LOS across Europe and determine the degree of variability to test the hypothesis that these variations in LOS may be attributed to either patient characteristics or the organisational and resource characteristics, and determine the relative importance of these factors. We also hypothesised that LOS may be influenced by the hospital cluster effect, which indicates that patients with similar characteristics may experience different processes of care and outcomes, depending on the hospital to which they are admitted. These include a wide range of complex factors not easily measured that might influence clinical care including local guidelines and policies, practices of individual clinicians, the admission policy, cultural practices that may influence the decision whether to admit or discharge patients, or clinical characteristics that may be observed within geographical clusters. These include factors common to the local population, such as socio-demographics, family and social support networks, cultural beliefs and attitudes, and environmental susceptibilities. Such effects on patient outcomes, although not measured individually, can be combined in a theoretical statistical model as the cluster effect to explain residual variation once the factors that have been measured have been accounted for.

# Methods

The study design and the methodology of the European COPD Audit have been extensively described elsewhere [18]. Briefly, this ERS-funded COPD audit was designed as a pilot study to evaluate clinical practice variability as well as clinical, organisational and resource factors related to outcomes for COPD hospital admissions in 13 countries across Europe (Austria, Belgium, Croatia, Greece, Ireland, Malta, Poland, Romania, Slovakia, Spain, Switzerland, Turkey and the UK). It was designed as a prospective, observational, non-interventional cohort trial.

The study comprised of a first 8-week phase during which all consecutive cases admitted to hospital for greater than 12 h due to an exacerbation of COPD were identified. These were considered potential cases, which were then confirmed as an exacerbation upon discharge, according to the diagnosis on the discharge report. Information on clinical practice was gathered for these patients. Comorbidities were recorded as a Charlson Comorbidity Index score, a validated measure of comorbidity status [19]. During this period, participating hospitals were also asked to complete a cross-sectional survey of the resources devoted to COPD acute care. Cases included had their medical history regarding the admission audited, and they were followed-up for 90 days after the admission to evaluate readmissions and survival.

The ERS managed the audit and named a steering committee, which reported to the ERS Executive Committee, to oversee the process. Items for the organisational and resource database were selected by the steering committee, based upon those that had previously been used and validated in the Spanish and UK national audits [20, 21]. The audit included questions related to the hospital and to the respiratory unit resources separately. An item list of the final survey is available [18].

The study period comprised hospital admissions from October 25, 2010, to December 19, 2010, and from January 3, 2011, until February 27, 2011, according to the seasonal COPD exacerbation peak of the participating countries. As there is no European ethics committee for audits; national societies ensured compliance with European and national ethical requirements, and all participants followed the European ethical requirements for scientific studies. All partners of the project accepted the general ethical rules of the ERS.

A software company (IDCode, Lausanne, Switzerland) was commissioned to design a web-based collection tool encompassing both the resource and organisational as well as clinical databases. The anonymised data was entered remotely at each participating site to a centrally controlled server. The web tool was established as a multilingual database, to allow the data to be documented in the language of each participating country. The web tool provided a "help facility" with explanatory text to assist with interpretation of the survey questions.

#### Data analysis

Statistical analysis was performed by a data-analysis team in Spain using Stata 12 (StataCorp LP, College Station, TX, USA). Descriptive variables were summarised by using the mean±sp for quantitative variables and the absolute (relative) frequencies for categorical ones.

Since LOS did not follow a normal distribution, the variables were characterised and those patients with a LOS beyond the median were further studied. Associations of variables potentially associated with LOS were tested by using a multivariable, multilevel logistic regression analysis for LOS longer than the median as the dependent variable. Continuous variables were categorised in different strata in which a missing value was considered one independent category. As part of an audit, missing information is also relevant. By categorising missing values, it allowed us to control the effect of one missing piece of information in the clinical performance. The only exceptions were age and Charlson index, which were not categorised since there was no missing information in these variables. In this multilevel model, data are analysed considering the hierarchical distribution of data, taking into consideration if the variables represent hospital characteristics (level 1) or patient characteristics (level 2). These analyses were performed with the following assumptions: 1) the multilevel model assumes a hierarchical data structure with explanatory variables measured at both hospital and patient levels; and 2) inter-hospital variation in the dependent variable was random.

In this analysis, we acknowledged that patients within the same hospital may receive similar care, which might be different from the care they may receive in other hospitals for the same clinical situation. This is so because patients within one hospital are subjected to distinct common contextual, hospital-linked influences. Our hypothesis is that this different behaviour in the different participant hospitals, the so-called hospital cluster effect, may influence LOS. Multivariable, multilevel regression analysis enables quantification of this hospital cluster effect and provides unbiased estimates of the regression coefficients. It can also concurrently include the predictor variables at both the patient level and hospital level. The coefficients of the predictor variables were transformed into odds ratios with 95% confidence intervals.

In addition, the hospital cluster effect was evaluated and quantified by two indicators: 1) the intra-cluster correlation coefficient (ICC) adapted to the logistic regression [22] and 2) the median odds ratio (MOR) with 95% confidence intervals [23, 24]. The ICC quantifies the fraction of the total variability in the outcomes attributable to the clustering effect. The ICC ranges from 0 to 1, and a lower ICC indicates a smaller cluster effect. The MOR is a measure of cluster heterogeneity and is defined as the median value of the distribution of odds ratios obtained when two individuals from different hospitals (*i.e.* one individual from the highest LOS hospital and one individual from the lowest LOS hospital) were randomly selected and compared. Simplified, the MOR can be interpreted as the median increased odds of reaching the outcome if an individual was admitted to another hospital with a greater risk of that outcome. The measure is always ≥1, where the MOR equals 1 when there is no between-cluster variation and the MOR increases as the variation becomes larger. Interestingly, the MOR is statistically independent of the prevalence of the outcome, can be easily computed, and is directly comparable with the coefficients of the individual- and hospital-level predictors. To plot the reduction in the between-hospital outcome variations after accounting for the hospital cluster effect and the predictor variables, the crude estimates of probability were transformed to the logit scale for comparisons, and cases with no LOS recorded were excluded.

For the multilevel analysis, we first fitted an "empty model" that only considered the clustering effect on the variability of LOS, expressed as ICC and MOR. Backward selection was attempted; however, the data did not converge in the multilevel analysis because of the high number of variables included so we opted for the forward selection procedure, which provided a better fit. The "final model", therefore, used a forward selection procedure based on Wald tests results, added as covariates to the model those "explanatory variables" associated with the outcome of interest in the bivariate analysis.

Various approaches to handle missing data were considered including analysis of the data without considering the missing information. This was rejected as in the multivariable analysis, a large number of cases would have been deleted due to missing values for different variables, which would have led to a shorter and biased database. Imputation was also considered; however, again, it was felt that this would result in a dataset that would again be artificial and not representative of our sample. Therefore, we considered the "missing values" as an additional category of those variables.

# **Results**

During the study, 19150 cases were initially considered upon admission as potential cases. Of these, the diagnosis of COPD exacerbation was confirmed by the lead investigator in that centre in 16018 cases upon discharge. However, in 37 cases, the LOS was not possible to calculate due to lack of information. Thus, the sample for the present analysis was 15981 patients. Mean±sd LOS was 8.7±8.3 days (median 7 days, interquartile range 4–11 days). The crude distribution of the LOS between countries is presented in table 1 in the supplementary material. Of the analysed population, 6955 (43.4%) cases had a LOS longer than the median. The description of the patient- and disease-related variables are summarised in table 1. The sample of patients had high pack-year smoking histories (mean 47.2 pack-years). Forced expiratory volume in 1 s was skewed towards more severe disease (mean 44% predicted) and a large proportion had features of infection (50% had sputum colour change) or co-existent lung consolidation (18.5%). The organisational variables within the sample varied significantly. Only 31.3% of units had an early supported discharge service and 48.8% had access to a pulmonary rehabilitation service. The hospital size and the numbers of respiratory specialist consultants, nurses, lung function technicians and physiotherapists varied greatly.

The bivariate analysis is summarised in the supplementary material (tables 1S and 2S). Resource and organisational variables associated with an increased LOS beyond the median were size-related (hospital size by number of beds or the number of pulmonologists), staff-related (having a respiratory team or the number of lung function technicians) and performance-related (the percentage of patients seen by a physiotherapist or the percentage of eligible patients included in a rehabilitation programme). Patient-related variables associated with an increased LOS beyond the median were related to comorbidities, the severity of the exacerbations and interventions given.

Male sex       10865 (67.8)         Age years       70.7±10.7         Current smokers       5012 (31.3)         Tobacco history pack-years       47.2±31.2         Comorbidity	Intercountry range
Current smokers       5012 [31.3]         Tobacco history pack-years       47.2±31.2         Comorbidity       2.4±1.6         Charlson Comorbidity Index score       2.4±1.6         Cardiovascular disease       6491 [40.5]         Diabetes       3181 [19.9]         Neoplasms       1950 [12.2]         Body mass index kg·m²       26.6±6.4         Admissions in previous 12 months       1.2 [1.8]         Spirometry       FVC % predicted         FV1 % predicted       44.0±17.4         Previous GOLD spirometric classification       No spirometry available         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15409 [96.2]         Sputum increase       10373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Aao <sub>2</sub> kPa       6.4±2.1         Aao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-tike findings </td <td>47.3-86.0</td>	47.3-86.0
Tobacco history pack-years         47.2±31.2           Comorbidity         2.4±1.6           Charlson Comorbidity Index score         2.4±1.6           Cardiovascular disease         6491 (40.5)           Diabetes         3181 (19.9)           Neoplasms         1950 (12.2)           Body mass index kg·m²         26.6±6.4           Admissions in previous 12 months         1.2 (1.8)           Spirometry         FVC % predicted           FEV1 % predicted         65.4±20.3           FEV1 % predicted         44.0±17.4           Previous GOLD spirometric classification         No spirometry available           No spirometry available         6512±40.7           Spirometry with no obstruction         1226±7.7           Mild         195±1.2           Moderate         2175±13.6           Severe         3737±23.3           Very severe         2098±13.1           Missing         75±0.5           Dyspnoea increase         15409 (96.2)           Sputum colour change         8134 (50.8)           pH         7.39±0.07           Paco <sub>2</sub> kPa         6.4±2.1           Ano <sub>2</sub> kPa         6.4±2.1           Ano <sub>2</sub> kPa         8.6±3.4           Chest r	63.4-72.7
Comorbidity Charlson Comorbidity Index score Cardiovascular disease Diabetes Diabetes Neoplasms Sepirometry FVC % predicted FEV1 % predicted Spirometry available Spirometry with no obstruction Mild Moderate Severe Moderate Severe Missing Very severe Missing Spynome increase Sputum colour change PH Road Sputum COPPD-like findings  C44+1.6 C49-14-1.6 C49-14-1.6 C49-14-1.6 C49-14-1.9 C49-14-18-18-18-18 C49-14-18-18 C49-14-18 C49-14-	15.4-46.8
Charlson Comorbidity Index score       2.4±1.6         Cardiovascular disease       6491 [40.5]         Diabetes       3181 [19.9]         Neoplasms       1950 [12.2]         Body mass index kg·m²       26.6±6.4         Admissions in previous 12 months       1.2 [1.8]         Spirometry       FVC % predicted         FV1 % predicted       65.4±20.3         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification       No spirometry available         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Aco <sub>2</sub> kPa       6.4±2.1         Ano <sub>2</sub> kPa       6.4±2.1         Chest radiography       8051 [50.3]	29.8-70.8
Cardiovascular disease       6491 [40.5]         Diabetes       3181 [19.9]         Neoplasms       1950 [12.2]         Body mass index kg·m⁻²       26.6±6.4         Admissions in previous 12 months       1.2 [1.8]         Spirometry       FVC % predicted         FV1 % predicted       44.0±17.4         Previous GOLD spirometric classification       5512±40.7         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Aco₂ kPa       6.4±2.1         Aco₂ kPa       6.4±2.1         Aco₂ kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 [50.3]	
Diabetes       3181 [19.9]         Neoplasms       1950 [12.2]         Body mass index kg·m⁻²       26.6±6.4         Admissions in previous 12 months       1.2 [1.8]         Spirometry       FVC % predicted         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification       FEV1 % predicted         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 [50.3]	1.7–3.1
Neoplasms       1950 (12.2)         Body mass index kg·m⁻²       26.6±6.4         Admissions in previous 12 months       1.2 (1.8)         Spirometry       FVC % predicted         FV1 % predicted       44.0±17.4         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 (96.2)         Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 (50.3)	32.4-71.4
Body mass index kg·m⁻²       26.6±6.4         Admissions in previous 12 months       1.2 {1.8}         Spirometry       FVC % predicted       65.4±20.3         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 {96.2}         Sputum increase       10 373 {64.8}         Sputum colour change       8134 {50.8}         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 {50.3}	11.9–27.5
Admissions in previous 12 months       1.2 [1.8]         Spirometry       FVC % predicted       65.4±20.3         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 [50.3]	2.6-18.9
Spirometry         FVC % predicted       65.4±20.3         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15409 [96.2]         Sputum increase       10373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 [50.3]	25.0-28.5
FVC % predicted       65.4±20.3         FEV1 % predicted       44.0±17.4         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15409 [96.2]         Sputum increase       10373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 [50.3]	0.4-1.8
FEV1 % predicted         Previous GOLD spirometric classification         No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase         Sputum increase       15 409 (96.2)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings       8051 (50.3)	
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No spirometry available       6512±40.7         Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 (96.2)         Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings	38.9-56.1
Spirometry with no obstruction       1226±7.7         Mild       195±1.2         Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 (96.2)         Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       8051 (50.3)	
Mild     195±1.2       Moderate     2175±13.6       Severe     3737±23.3       Very severe     2098±13.1       Missing     75±0.5       Dyspnoea increase     15409 [96.2]       Sputum increase     10373 [64.8]       Sputum colour change     8134 [50.8]       pH     7.39±0.07       Paco <sub>2</sub> kPa     6.4±2.1       Pao <sub>2</sub> kPa     8.6±3.4       Chest radiography       Normal or COPD-like findings     8051 [50.3]	9.9-53.6
Moderate       2175±13.6         Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography       Normal or COPD-like findings	3.2-34.0
Severe       3737±23.3         Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 [96.2]         Sputum increase       10 373 [64.8]         Sputum colour change       8134 [50.8]         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography         Normal or COPD-like findings       8051 [50.3]	0-3.0
Very severe       2098±13.1         Missing       75±0.5         Dyspnoea increase       15 409 (96.2)         Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco <sub>2</sub> kPa       6.4±2.1         Pao <sub>2</sub> kPa       8.6±3.4         Chest radiography         Normal or COPD-like findings       8051 (50.3)	2.7-31.0
Missing     75±0.5       Dyspnoea increase     15 409 (96.2)       Sputum increase     10 373 (64.8)       Sputum colour change     8134 (50.8)       pH     7.39±0.07       Paco₂ kPa     6.4±2.1       Pao₂ kPa     8.6±3.4       Chest radiography       Normal or COPD-like findings     8051 (50.3)	11.6–34.9
Dyspnoea increase       15 409 (96.2)         Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography         Normal or COPD-like findings       8051 (50.3)	4.8-23.8
Sputum increase       10 373 (64.8)         Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography         Normal or COPD-like findings       8051 (50.3)	0-2.7
Sputum colour change       8134 (50.8)         pH       7.39±0.07         Paco₂ kPa       6.4±2.1         Pao₂ kPa       8.6±3.4         Chest radiography         Normal or COPD-like findings       8051 (50.3)	92.5–100
pH 7.39±0.07  Paco₂ kPa 6.4±2.1  Pao₂ kPa 8.6±3.4  Chest radiography  Normal or COPD-like findings 8051 [50.3]	49.2-79.0
Paco <sub>2</sub> kPa 6.4±2.1 Pao <sub>2</sub> kPa 8.6±3.4  Chest radiography  Normal or COPD-like findings 8051 [50.3]	36.9-63.3
Pa02 kPa8.6±3.4Chest radiography8051 (50.3)	7.37–7.41
Chest radiography Normal or COPD-like findings 8051 (50.3)	5.7–7.0
Normal or COPD-like findings 8051 (50.3)	7.3–10.8
<b>3</b>	
	33.0-55.5
Consolidation         2969 (18.5)           Other findings         4752 (29.7)	8.9-32.6 16.7-58.0

Data are presented as n (%) or mean $\pm$ sD, unless otherwise stated. FVC: forced vital capacity; FEV1: forced expiratory volume in 1 s; GOLD: Global Initiative for Chronic Obstructive Lung Disease;  $P_0$ CO<sub>2</sub>: arterial carbon dioxide tension;  $P_0$ CO<sub>2</sub>: arterial oxygen tension; COPD: chronic obstructive pulmonary disease.

The empty model exhibited a significant hospital cluster effect (ICC 0.18) and cluster heterogeneity (MOR 2.30). In essence, 18% of the variability was explained by clustering resulting in a 2.3-fold increased association with LOS. The adjusted model retained a number of variables as outcome predictors, and these were split into the five categories: 1) patient-related variables, 2) disease-related variables, 3) treatment-related variables, 4) exacerbation severity-related variables, and 5) resource-related variables; and the odds ratios depicting the likelihood of the association are in table 2. The strongest relationships were seen between advanced Global Initiative for Chronic Obstructive Lung Disease (GOLD) status (OR 1.77 for GOLD class IV) and the use of mechanical ventilation (OR 2.15). The use of antibiotics and the presence of consolidation on the chest radiography were also strongly associated with LOS (OR 1.46 and 1.27, respectively). Although there was a crude variability of LOS between the different countries, this was reduced after accounting for these clinical factors and the clustering effect (figure 1). The graphical representation of significant variables is depicted in figure 2.

# **Discussion**

This study is a comprehensive analysis of factors associated with LOS for COPD patients admitted to hospital with an exacerbation across Europe. Other clinically relevant outcomes including death and readmissions have been previously published elsewhere [25]. The mean and median LOS across Europe were higher than expected from previous studies in individual countries around Europe [10, 26] and showed great variability. This is in keeping with a recent systematic review that showed consistently high variability of COPD LOS across countries in Europe, North America and Australasia [27]. The variability observed in the present study was, however, reduced when the hospital cluster effect (an important consideration in reducing bias from intrinsic institutional factors not measured in the analysis [24]) was accounted for. Clinical and patient factors were associated with LOS but resource and organisational factors had less influence individually than expected from previous studies.

#### Patient-related factors

In contrast to previous studies [9, 28], the variables of an increase in Charlson index by 1 point, increasing age and body mass index (BMI) classified as overweight or obese were not strongly associated with longer LOS. However, there was a suggestion within the trend that those who had a low BMI (OR 1.13, 95% CI 0.97–1.33) were more likely to have a longer LOS. This is in accordance with previous studies demonstrating that nutritional status is of importance in quality of life, respiratory muscle strength and exercise capacity [29, 30], all factors which would affect recovery from an exacerbation. More recently, an American study reported a reduction in LOS, readmission rate and hospitalisation cost associated with oral nutritional supplementation and further studies evaluating this are needed [31].

Less anticipated was that current smoking status was also associated with a shorter LOS. This may be attributed to the "healthy smoker effect" whereby smokers have less perception or greater tolerance of their symptoms [32, 33]. Further to this, the desire for discharge may be enhanced by additional motivational factors, such as a desire to smoke and therefore to mobilise earlier than ex-smokers. Though the present study did not assess psychosocial factors which have been previously noted to impact LOS [11, 34], this observed effect with smoking status may be related to such factors and should be further evaluated.

# Disease and exacerbation severity-, and treatment-related factors

Disease and exacerbation severity-, and treatment-related factors were more strongly associated with a LOS longer than the median, and of those, the biggest impact was noted with GOLD classification IV. This finding is in keeping with previous reports [8, 35, 36]. A history of previous admissions was also associated with longer LOS. This effect may be due to disease severity but is also likely to be impacted by psychosocial factors. Certainly, an association between LOS, depression and early readmission has been reported [37].

Although severity scores for COPD exacerbation exist, such as BODE index [38], these were not recorded as a variable in the present study. Therefore, other clinical features that occur in more severe episodes have been evaluated and interpreted as surrogate markers of exacerbation severity.

Patients who were acidotic or had sputum purulence were also more likely to have an increased LOS (OR 1.25 and 1.11, respectively) as reported by previous studies in different populations [6–9].

The use of invasive ventilation was associated with increased LOS. In keeping with this is a report by Limsuwat *et al.* [39], who also found intubation and mechanical ventilation was associated with increased LOS. The use of antibiotics, theophylline and diuretics were also associated with an increased risk of a LOS longer than the median. This may be attributed to a more severe episode, as such factors may be considered surrogate markers of a more severe episode. However, this interpretation is subject to appropriate treatment selection, and does not consider the potential increase in LOS caused by over-treatment that may result in complications such as antibiotic-associated diarrhoea or oxygen-induced hypercapnic respiratory failure, which may impact LOS.

ariable ariable	Category		OR (95% CI)	
And and and a second discount		<u> </u>		
Patient-related variables			1 01 (1 01	1 1 02)
Age years Smoking status	Ex-smoker		1.01 (1.01–1.02) 1	
Smoking status	Active		•	0 00)
	Never-smoker		0.85 (0.78–0.93) 0.94 (0.80–1.12)	
	Miss		1.08 (0.92	
Charles Camarhidity Index core	MIS	sing	1.04 (1.0	
Charlson Comorbidity Index score Body mass index kg·m <sup>-2</sup>	<1'	D F	1.13 (0.97	
Body Mass Maex kg·m	19.5–		1.13 (0.77	7-1.55)
	25-2		0.89 (0.79	2 1 00)
	>29		0.88 (0.78	
	Miss		0.83 (0.74	
isease-related variables	MIS	sing	0.63 (0.72	1-0.73)
Previous admissions	N		1	
Previous admissions			1.15 (1.07	7 1 2/)
	Yes Missing		1.15 (1.07	
GOLD stage	MIS	•	1.11 (0.96	
GOLD stage	2		1.10 (0.79	
	3		1.20 (0.8 <i>6</i> 1.77 (1.25	
		•	•	•
	Miss	sing	1.36 (0.97	7-1.90)
cacerbation severity-related variables	7	٥٦	1.25 (1.12	2 1 (0)
рН		<7.35		2-1.40)
	≥7		1 0 / 10 0	2.4.4.1
D. I	Miss	5	1.04 (0.93	3-1.16)
Purulent sputum		0	1 11 (1 2)	2 4 00)
		es	1.11 (1.03	
	Miss		1.05 (0.91	
Consolidation	N		1 05 (1 1	
	Ye	es	1.27 (1.16	5-1.40)
eatment-related variables	N.		4	
Invasive mechanical ventilation	N		1	- 0.770)
1. 1	Yes		2.15 (1.25	0-3.72)
In-hospital antibiotics	No Yes		1 1.46 (1.30–1.64)	
				J-1.64J
In-hospital methylxanthines	N		1	7 4 (0)
	Ye		1.22 (1.07	/-1.4UJ
In-hospital inhaled corticosteroids	N		1 10 (1 0	1 4 00)
	Ye		1.10 (1.01	1-1.20)
In-hospital diuretics	N		1 0 / (4 0)	- 4 (0)
D. I	Ye		1.36 (1.25	0-1.491
Radiography performed		0	1 (0 (1 0	1.05)
	Ye	es	1.42 (1.03	3–1.95J
esources-related variables	.,			
Respiratory ward	Yes		1 1.33 (1.22–1.45)	
	No			
	Missing		0.38 (0.11	1-1.32J
Number of respiratory specialists in unit	<6 ≥6		1 10 (1.0)	2 4 (0)
			1.19 (1.00	
	Miss	sing	0.68 (0.38–1.24)	
_evel	Empty model		Final model	
	Country	Hospital	Country	Hospit
	<u> </u>	<u> </u>		
tra-class correlation coefficient	0.11	0.19	0.12	0.21
ledian odds ratio	0.19	2.31	1.94	2.42

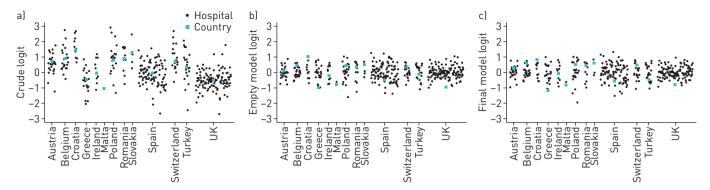


FIGURE 1 The variability of probability of that the length of stay is longer than the median by country. The black dots represent the logit of each hospital a, b accounting only for cluster effects without covariates and c accounting for cluster effects and covariates. The green dots represent the logit of each country. Logit is defined as the probability determined by logistic regression: logit(p) = log(p/1-p).

#### Resource and organisational factors

Of the resource and organisational factors that were found to be associated with a LOS longer than the median in the initial bivariate analysis, only two were retained in the final model. This was despite taking a more conservative value for significance (p>0.1). In this analysis, no association was noted between LOS in hospitals where early supported discharge service was available, where there was a respiratory consultant available out of hours or where there were more than six respiratory consultants in the organisation, contrasting with previous reports [4, 14]. Whilst in the UK, 75.2% of centres had an early supported discharge service, in the rest of Europe, this figure declined to 15.4% and is likely to include a number of different service models under that umbrella term.

Data from the UK COPD audit data from 2003 and 2008 [14, 26], very much support the implementation of services such as early supported discharge programmes due to an observed reduction in LOS and these reports suggested that units with more respiratory consultants and better quality organised care had lower mortality and reduced LOS. Furthermore, a previous randomised controlled trial [4] also noted a dramatic reduction in LOS from 6.1 to 3.2 days in a group of patients receiving care from an early supported discharge service *versus* standard inpatient care. The explanation for our findings is unclear but may relate to the relatively high LOS reported in this audit by the UK centres and the low level of this form of service across Europe as a whole. Other studies have supported use of early supported discharge services in the absence of a reduction in LOS [40–42] due to patient preference or positive impacts on patients' well-being with use of such services.

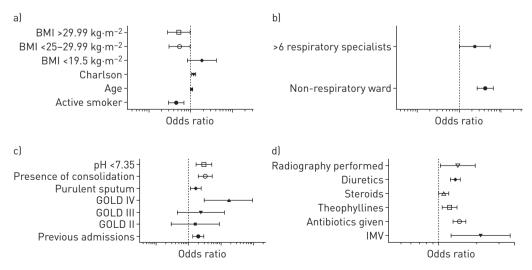


FIGURE 2 Odds ratios for a length of stay longer than the median of the variables retained in the final model. Vertical dashed lines represent OR 1. BMI: body mass index; GOLD: Global Initiative for Chronic Obstructive Lung Disease; IMV: invasive mechanical ventilation.

This study has found that when resource and organisational factors are considered within a cluster model their individual impact on length of stay is greatly diminished. In real-life situations, it is the complex manner in which these various factors interact that determines outcomes for patients and that is reflected in the cluster analysis. Further studies are needed to evaluate this complex interaction in more detail in order to tease out the important influencing factors and to provide appropriate targets for improving outcomes with relation to length of stay.

# Strengths and limitations

The design of the study and statistical methods underwent a great deal of consideration. Other methods of data analysis, including logit, Poisson, negative and polynomial regression, were rejected as it was felt that the chosen methodology, which enabled accounting for factors that were not measured as outcomes, was superior. Factors such as hospital admission policies, cultural practices, presence of family at home to care for the discharged patients, weather, availability of beds, clinical practice styles, physician thresholds for discharge and socio-demographic factors vary greatly across the countries in Europe included within this audit, and all have an impact on LOS. As none of these was recorded in this study and many are difficult to measure, the multi-level approach was considered ideal, as these factors would be incorporated within the hospital cluster effect.

Use of such a model has obvious limitations in keeping with other mathematical models; for example, it may over-account for factors, making some associations difficult to find. The aim, however, is to highlight the most important and significant associations. Further studies are vital to evaluate the relative importance of the factors found to be associated with LOS in this study and other factors that may be encompassed within the hospital cluster effect that have not been measured.

Participation was on a voluntary basis, so there was no intention of representative sampling. The heterogeneity of the study participants across multiple countries and the interpretation of study variables by different investigators are likely additional limiting factors.

The diagnosis of exacerbation of COPD may be subject to interpretation, and by excluding patients who were admitted for less than 12 h, we perhaps imposed a degree of selection bias into our population, and perhaps it would be also interesting to evaluate those who presented to hospital but were discharged more quickly.

Missing data were a significant problem, as is often the case with retrospective audits. Various methods of dealing this were considered, all of which posed their own problems. We acknowledge that the approach utilised was not flawless, but we considered it the least problematic and, in most cases, the more amenable to clinical interpretation.

Despite these limitations, the strengths of the present study, such as the wide European sampling and the comprehensive statistical analysis of organisational and clinical variables by using a multilevel approach, is highly informative and serves to eliminate biases associated with the clustering effect.

### Conclusions

This study has shown that there is a wide observed variation in LOS across Europe. Some of this is likely to be attributable to the different service models used in those countries. Patient and care process factors of highest impact that were associated with increased LOS in this study were reduced BMI, advanced GOLD classification, presence of respiratory acidosis, consolidation and the use of diuretics. Resource and organisational factors such as size of hospital and the presence of early supported discharge service were not key determinants of LOS after accounting for the clustering effect, whilst admission to a respiratory ward remained an important factor. Identifying patients at risk of a long hospital stay at admission and providing targeted interventions such as early mobilisation offers the potential to reduce LOS for these individuals.

Whilst there are a number of individual factors that affect the LOS of any one patient, it is not just the presence of resources or services but the complex interactions of how a hospital functions to use those resources and services effectively, or not, that is most important in determining the average LOS of a cohort of admitted COPD patients. We suggest that future research should be targeted at identifying those factors that optimise the organisational effectiveness of hospitals irrespective of the health service system within which they operate.

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