# **BMJ Open** Population-based analysis of patients with COPD in Catalonia: a cohort study with implications for clinical management

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#### ABSTRACT

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Correspondence to Professor Josep Roca; jroca@clinic.cat **Background** Clinical management of patients with chronic obstructive pulmonary disease (COPD) shows potential for improvement provided that patients' heterogeneities are better understood. The study addresses the impact of comorbidities and its role in health risk assessment.

**Objective** To explore the potential of health registry information to enhance clinical risk assessment and stratification.

**Design** Fixed cohort study including all registered patients with COPD in Catalonia (Spain) (7.5 million citizens) at 31 December 2014 with 1-year (2015) follow-up.

**Methods** A total of 264 830 patients with COPD diagnosis, based on the International Classification of Diseases (Ninth Revision) coding, were assessed. Performance of multiple logistic regression models for the six main dependent variables of the study: mortality, hospitalisations (patients with one or more admissions; all cases and COPDrelated), multiple hospitalisations (patients with at least two admissions; all causes and COPD-related) and users with high healthcare costs. Neither clinical nor forced spirometry data were available.

**Results** Multimorbidity, assessed with the adjusted morbidity grouper, was the covariate with the highest impact in the predictive models, which in turn showed high performance measured by the C-statistics: (1) mortality (0.83), (2 and 3) hospitalisations (all causes: 0.77; COPD-related: 0.81), (4 and 5) multiple hospitalisations (all causes: 0.80; COPD-related: 0.87) and (6) users with high healthcare costs (0.76). Fifteen per cent of individuals with highest healthcare costs to year ratio represented 59% of the overall costs of patients with COPD.

**Conclusions** The results stress the impact of assessing multimorbidity with the adjusted morbidity grouper on considered health indicators, which has implications for enhanced COPD staging and clinical management. **Trial registration number** NCT02956395.

## **INTRODUCTION**

Chronic obstructive pulmonary disease (COPD) is one of the major disorders included in the WHO programme addressing non-communicable diseases.<sup>1</sup> It is estimated that COPD will become the third leading cause of death by 2020.<sup>2</sup> Moreover, projections on COPD prevalence and costs over

# Strengths and limitations of this study

- The main strength of the study is that it contributes to risk prediction of relevant clinical events in patients with chronic obstructive pulmonary disease (COPD).
- The study shows high potential to assess health risk factors at population level indicating the high impact of comorbidities. It can contribute to define innovative strategies aiming at reducing the healthcare impact of patients with COPD.
- Full potential of the approach should be proven by integrating registry information and electronic medical records.
- Lack of clinical information, spirometric data and history of tobacco smoking reduces the potential for standardised risk characterisation of patients with COPD.

the next 15 years indicate a rapidly escalating burden, mainly due to population ageing, on both health and social support systems.<sup>34</sup>

While acknowledging the progress made in terms of standard of care recommendations,<sup>5</sup> it is accepted that a better understanding of patients' heterogeneities constitutes a key challenge to further enhance both prevention and management of patients with COPD aiming at healthcare value generation.<sup>67</sup>Recent studies indicate a high impact of comorbidities on use of healthcare resources in patients with COPD prompting the need for assessing novel integrated care strategies with a patient-oriented approach.<sup>89</sup> It is well accepted that several prevalent chronic conditions often occur as clusters of comorbidities in patients with COPD<sup>10-12</sup> and potential explanatory mechanisms for the phenomenon have been proposed.<sup>1314</sup>

The current study addresses comorbidities in patients with COPD based on the hypothesis that assessment of all patients with COPD living in a given geographical area, a population-based analysis of the patients with the disease, can provide valuable insights to better understand COPD heterogeneity. Specifically, the research objective was to explore the potential of the health registry information contained in the Health Surveillance System of the region of Catalonia (Spain) (7.5 million inhabitants) to enhance health risk assessment and stratification in the clinical arena.<sup>15</sup><sup>16</sup> To this end, we analysed a total of 264830 patients with COPD from all healthcare layers, registered in 31 December 2014 and followed up in 2015, to elaborate predictive models for six key health indicators: (1) mortality, (2 and 3) hospitalisations (all causes and COPD-related), (4 and 5) multiple hospitalisations (all causes and COPD-related) and (6) users with high healthcare costs. Ultimately, this study aimed to consider the weight of the different covariates included in the predictive models to provide useful information to enhance clinical management.

The study is a relevant component of the programme for collaborative management of complex chronic patients being deployed in the region during the period 2016–2020.<sup>17</sup> An ancillary aim of the research was to assess factors determining the economic impact of the patients with the disease, as well as to identify areas of action to increase healthcare efficiencies in the management of these patients.

#### **METHODS**

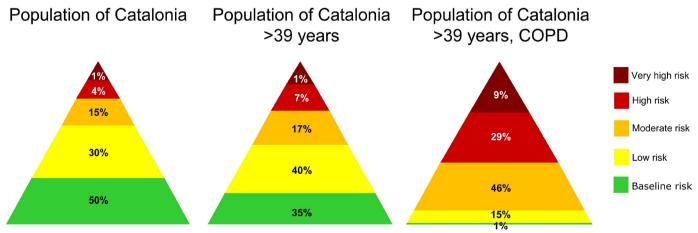
## Population-based risk assessment: adjusted morbidity grouper

The Catalan Health Surveillance System (CHSS) includes updated registries of the region of Catalonia (Spain) (7.5 million inhabitants) from Primary Care, Hospital-related events (hospitalisations, emergency room consultations and specialised outpatient visits), Pharmacy, Mental Health, Socio-sanitary services and other items (homebased respiratory therapies, dialysis, outpatient rehabilitation and non-urgent healthcare transportation) since 2011.<sup>18 19</sup> It allows analyses on use of healthcare resources, pharmacy consumption, prevalence of key disorders and population-based health risk assessment.<sup>15 16</sup> It is of note that although integration of CHSS registry data with electronic medical records is not yet in place, it constitutes the main goal of the PADRIS programme,<sup>20</sup> officially launched on January 2017.

The regional population-based health risk assessment tool, named GMA (Adjusted Morbidity Groups), is used to elaborate the health risk strata pyramid of the general population of Catalonia (figure 1, left triangle).<sup>15 16</sup> The GMA tool predicts individual patient risk, periodically updated on a 6-month basis, based on multimorbidity information gathered from CHSS registry data. The rationale behind the use of GMA, against alternative health risk assessment tools, is that it complies with four main recommended criteria,<sup>15</sup> that is, (1) a population health approach (uses the entire population of 7.5 million inhabitants of the region), (2) publicly owned without licensing constraints, (3) open source computational algorithms and (4) the adjusted morbidity grouper relies mostly on statistical criteria, as opposed to other tools that include expert-based coefficients, thus facilitating quick transferability to other territories. Detailed descriptions of the GMA, as well as its evaluation, have been reported elsewhere.<sup>15</sup><sup>16</sup> Methodological details of the GMA algorithm are described in the online supplementary figure 1S.

## Study dataset and design

The current study design is a fixed cohort analysis of the entire population of 264830 patients, alive and above 39 years of age, included in the CHSS on 31 December 2014 with the clinical diagnosis of COPD. New patients registered during 2015 were not included in the study. For the study purposes, the diagnosis of COPD was based on the International Classification of Diseases (Ninth Edition) coding<sup>21</sup> (online supplementary table 1S) declared by the patient's responsible physician, either a



**Figure 1** GMA health risk grades. The left triangle depicts the distribution of all Catalan citizens, expressed as percentage, in five arbitrary health risk layers defined using the GMA (Adjusted Morbidity Groups)<sup>15 16</sup> as a population-based health risk stratification tool. The central triangle indicates the distribution in these five health risk layers of the subset of Catalan citizens older than 39 years. The right triangle displays the distribution of the study group of patients with chronic obstructive pulmonary disease (COPD) (264 830 patients) across the GMA health risk grades: baseline, low, moderate, high and very high.

primary care professional or a specialist. The study did not take into account background clinical information, nor forced spirometry data.

The analyses were carried out for the entire population of patients with COPD, but the study considered different subsets of patients based on the GMA health risk grading: (1) baseline, (2) low, (3) moderate, (4) high and (5) veryhigh-risk patients. The thresholds defining these subsets of patients correspond to the percentiles 50, 80, 95 and 99 of the GMA grading for the general population (figure 1, left triangle). For the purposes of the study, patients with COPD falling into the baseline health risk group (1%) were merged with the low-risk group such that only four GMA grades were considered in the analysis (figure 1, right triangle).

Health risk predictive modelling was elaborated with registry data from year 2015 for the six main dependent variables: (1) mortality, (2 and 3) hospitalisation (all causes and COPD-related), (4 and 5) multiple hospitalisations (all causes and COPD-related) and (6) users with high healthcare costs, as described below. In the study, dependent variables were defined as binary variables. Mortality was defined as true if a patient died during the period from 1 January to 31 December 2015, regardless of whether it occurred in the hospital, at the patient's home or in other settings such as skilled nursing facilities. The category all causes hospitalisations was true if a patient had one or more hospital admissions due to any cause during 2015. COPD-related hospitalisations refer only to events triggered by acute exacerbations of COPD. Multiple hospitalisations were defined as two or more hospital admissions. Users with high healthcare costs refer to subjects above percentile 85 (PCT85) in terms of yearly healthcare costs during 2015.

The study used retrospective deidentified data from administrative databases (CHSS). Therefore, neither informed consent nor ethical committee approval was required according to the current legislation in Catalonia. The analyses were developed under the umbrella of the Nextcare project (http://www.nextcarecat.cat/) (https://clinicaltrials.gov/: NCT02956395).

#### Assessment of economic burden

Allocation of healthcare expenditure to each patient, including pharmacy, was done through the Personal Health Identification Number since it allows each billing invoice to be attributed to a given patient. The healthcare expenditure includes hospitalisation, primary care, pharmacy, health transport, respiratory home care therapies, outpatient visits and skilled nursing facilities. The key outcome variable of the analysis was 1-year healthcare resource use and expenditure by patient.<sup>19</sup> Calculation of individual healthcare costs was done by addition of costs for each item included in the CHSS registries alluded to above. This methodology allows calculation of the total healthcare expenditure in patients with COPD. Consequently, it allows to perform a holistic analysis of healthcare expenditure.

#### **Statistical analysis**

The study outcomes are described for the entire population of patients with COPD. Comparisons among the four subgroups defined by GMA health risk grades were done. The results are summarised in the main manuscript and complementary information is reported in detail in the online supplementary material. In this population-based analysis (table 1), age and number of chronic comorbidities are summarised as mean and SD, the proportion of women and the morbidities are expressed as percentages, while mortality rate, hospitalisation rate and COPD-related hospitalisation rate are expressed per 100 patients with COPD among patient groups. Comparisons among groups were done using analysis of variance for continuous variables, and  $\chi^2$  test for binary and nominal variables.

Statistical analyses were performed using SPSS software V.18.0. All statistical tests and confidence intervals were constructed with a type I error (alpha) level of 5%, and P values lower than 0.05 were considered statistically significant.

#### **Predictive modelling**

Multiple logistic regression analyses were used to generate health risk predictive models for the six main outcome variables of the study, namely, (1) mortality, (2 and 3) hospitalisation (all causes and COPD-related), (4 and 5) multiple hospitalisations (all causes and COPD-related) and (6) users with high healthcare costs. The following model independent variables were considered: age, sex, GMA health risk grades, previous history of hospital admissions, emergency room consultations and use of social support services. Each independent variable was included in the model as a categorical variable to allow for possible non-linearity in the relationship between variables and the relevant outcomes. For each measure, we collapsed the uppermost categories to ensure there were enough individuals in each cell to allow estimation of the parameters. All covariates were entered in the model one by one and retained when they showed a significant contribution to the predictive accuracy (P < 0.10). The predictive role of each covariate into the model was assessed with a log-likelihood ratio test. To evaluate the performance of the resulting predictive models, we calculated the C-statistics (ie, the area under the receiver operating characteristic curve).<sup>22 23</sup>

#### RESULTS

#### **Population-based analysis**

The distribution of the entire population of the region in the health risk strata pyramid is depicted in figure 1, left triangle, wherein citizens are distributed in five health risk grades defined by the GMA percentiles, namely, (1) 50% of the population with baseline health risk (green), (2) 30% of individuals with low health risk (yellow), (3) 15% with moderate health risk (orange), (4) four percent with high risk and (5) 1% with very high health risk (brown).

Table 1 Main characteristics of the study group by GMA health risk grades									
	Low risk	Moderate risk	High risk	Very high risk	Total				
Patients (n)	41809	121918	76237	24866	264830				
Age (years)*	60.8±11.8	69.4±11.8	75.1±10.9	77.4±10.1	70.5±12.5				
Women (%)*	36.0	36.2	35.9	35.3	36.0				
Morbidity									
No of chronic comorbidities*	2.9±1.1	5.2±1.4	7.0±1.6	8.4±1.7	5.6±2.1				
Diabetes (%)*	5.2	22.7	39.4	52.6	27.5				
Heart failure (%)*	0.2	5.2	30.7	63.7	17.2				
Hypertension (%)*	24.0	60.4	81.1	88.9	63.3				
Renal failure (%)*	0.3	5.8	23.4	47.8	13.9				
Dementia (%)*	0.3	2.6	7.8	13.6	4.7				
Cirrhosis (%)	0.5	1.6	3.0	4.8	2.1				
Depression (%)	8.8	20.1	28.6	34.2	22.1				
Stroke (%)*	0.5	5.9	19.2	31.8	11.3				
Ischaemic coronary disease (%)*	0.9	9.0	28.0	42.7	16.4				
Malignancy (%)*	4.2	17.0	32.1	40.9	21.6				
Locomotor system (%)*	29.3	56.3	67.9	72.0	56.8				
Osteoporosis (%)*	3.9	10.0	14.3	17.9	11.0				
Arthrosis (%)*	9.8	32.8	47.6	51.0	35.1				
2015 events	n (%)	n (%)	n (%)	n (%)	n (%)				
Mortality*	391 (0.9)	3213 (2.6)	6792 (8.9)	5955 (23.9)	16351 (6.2)				
Patients with hospitalisations (all causes)*	1785 (4.3)	12451 (10.2)	19417 (25.5)	12 496 (50.3)	46149 (17.4)				
Patients with hospitalisations (COPD related)*	404 (1.0)	2769 (2.3)	4697 (6.2)	3600 (14.5)	11470 (4.3)				
Patients with multiple hospitalisations (all causes)*	366 (0.9)	3026 (2.5)	6687 (8.8)	6035 (24. 0)	16114 (6.1)				
Patients with multiple hospitalisations (COPD related)*	71 (0.2)	524 (0.4)	1169 (1.5)	1210 (4.9)	2974 (1.1)				
Users with high healthcare costs*	1119 (2.7)	10064 (8.3)	17333 (22.7)	11609 (46.7)	40 125 (15.2)				

Age and number of chronic comorbidities expressed as mean±SD; gender and morbidities are expressed as percentages; comparisons among risk grades were done using analysis of variance for continuous variables and  $\chi^2$  test for binary and nominal variables. \*P<0.01.

COPD, chronic obstructive pulmonary disease; GMA, Adjusted Morbidity Groups.

These risk strata show strong associations with mortality, hospital admissions, use of healthcare resources and expenditures, as reported in detail elsewhere.<sup>15 16</sup>

The central triangle (figure 1) indicates the distribution of the general population older than 39 years in the five health risk grades, described above. It constitutes the reference risk strata pyramid to be compared with the population-based COPD analysis. Finally, the distribution of the study group of patients with COPD is depicted in the right triangle that indicates the effects of the chronic pulmonary disease on health risk stratification. As expected, most of the patients with COPD (84%) were distributed between moderate (46%), high (29%) and very high (9%) health risk layers. Only 15% of the patients fell in the low risk level (yellow) and 1% were allocated in the baseline health risk level (green).

The study showed a COPD prevalence of 6.6% of all subjects above 39 years of age. The mean age was 70

(SD 12.5) years. Women represented 36% of the study group. On average, these patients presented 5.6 (SD 2.1) comorbid conditions. The mortality rate was 6.2%. The hospitalisation rate, all causes, was 17.4% (n=46149), whereas the rate of COPD-related hospitalisations was 4.3% (n=11470), which represents 24.9% of the total number of hospitalisations registered in these patients. As expected, the rate of comorbidities showed a consistent increase with GMA grading, which was the most apparent for cardiovascular disorders, type 2 diabetes mellitus–metabolic syndrome and/or anxiety–depression, as displayed in table 1.

## **Predictive modelling**

The summary information of the predictive modelling for each of the six outcome variables—mortality, hospitalisations (all causes and COPD-related hospitalisations), multiple hospitalisations (all causes and COPD-related

Table 2 Summary description of the six predictive models									
		Hospitalisations		Multiple hospitalisations		Users with high healthcare			
	Mortality	All causes	COPD related	All causes	COPD related	costs (PCT85)			
C-statistics (AUC)	0.829	0.766	0.807	0.803	0.865	0.763			
Covariates	OR								
Sex									
Male	1	1	1	1	1	1			
Female	0.701	0.825	0.885	0.818	0.823	0.84			
Age (years)									
40–54	1	1	1	1	1	1			
55–64	1.693	1.254	1.435	1.308	1.428	0.884			
65–74	2.103	1.506	1.675	1.576	1.7	0.91			
75–84	3.749	2.054	2.017	1.983	1.634	0.898			
>84	9.911	2.827	2.049	2.335	1.272	0.681			
Admissions									
Group A	1	1	1	1	1	1			
Group B	1.394	1.903	4.699	1.952	6.952	1.59			
Group C	1.368	2.039	8.592	2.266	15.214	1.577			
GMA grade									
Low	1	1	1	1	1	1			
Moderate	1.826	1.896	1.644	2.128	1.649	3.125			
High	4.316	3.998	2.646	5.419	2.852	8.858			
Very high	8.919	7.851	3.548	11.042	3.891	20.285			
Emergency room vi	sits								
0	1	1	1	1	1	1			
1–2	1.106	1.405	1.37	1.526	1.505	1.353			
3–5	1.276	1.865	1.684	2.179	2.013	1.771			
>5	1.643	2.615	2.235	3.354	3.032	2.428			
Social support									
No	1	1	1	1	1	1			
Yes	2.289	1.008	0.892	0.917	0.922	0.877			

The intensities of grey background colour reflect the magnitude of ORs, being white when value is one and stronger grey when they are closer to 0 or have a higher positive value.

Admissions:

Group A corresponds to patients with no registries of hospital admissions within the period 2011–2014; Group B includes patients with history of admissions before 2014, but without admissions in that year; Group C includes patients with hospital admissions during 2014. Graphical representation and details of the six predictive models with the corresponding 95% CIs of the ORs are shown in the online supplementary figures 2S–7S and tables 2S–7S.

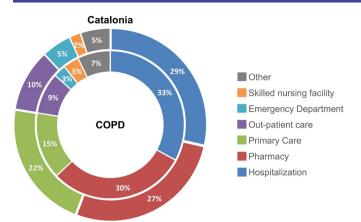
AUC, area under the receiver operating characteristic curve; COPD, chronic obstructive pulmonary disease; GMA, Adjusted Morbidity Groups.

repeated admissions) and users with high healthcare costs—are depicted in table 2, wherein significant covariates, the corresponding ORs and the C-statistics are indicated for each predictive model.

It is of note that age, closely followed by GMA grading, showed the two highest independent associations in the mortality model. Likewise, GMA grading depicted the highest predictive role in three out of the six models: hospitalisations (all causes), multiple hospitalisations (all causes) and users with high healthcare costs. Interestingly, for COPD-related events (hospitalisations and multiple hospitalisations), the covariate with highest predictive role was history of hospitalisations, whereas the ORs of the covariate GMA grades were markedly lower than in the other predictive models.

## Economic impact of patients with COPD

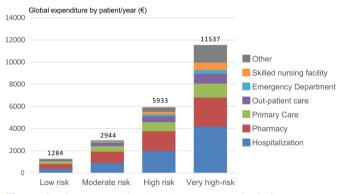
Figure 2 compares the costs of the different items, expressed as percentages, for the general population of Catalonia (outer circle) and those generated by the study

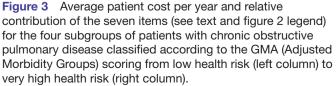


**Figure 2** Indicates the distribution of costs of main items, expressed as percentages. The outer circle corresponds to overall cost for the Catalan Health System, whereas the inner circle indicates the corresponding relative costs ascribed to patients with chronic obstructive pulmonary disease (COPD). The absolute values are (1) Hospitalisation (€2291.8 million and €356.6 million, respectively), (2) Pharmacy (€2193.4 million and €325.8 million), (3) Primary care (€1745.0 million and €158.9 million), (4) Outpatient specialised care (€842.9 million and €98.1 million), (5) Emergency department (€401.5 million and €29.7 million), (6) Skilled nursing facility (Catalonia €155.1 million; COPD €37.5 million), and (7) Other (€404.0 million and €78.0 million). The last item, Others, includes home-based respiratory therapies, dialysis, outpatient rehabilitation and non-urgent healthcare transportation.

group of patients with COPD (inner circle). Briefly, COPD hospitalisations, pharmacy, skilled nursing care and other costs (respiratory therapies) are above the mean cost of the corresponding items in the general population of the region.

The average healthcare costs of patients with COPD per year was  $\in$ 4238 as compared with a mean of  $\in$ 987 a year per citizen in the region. It is of note that the cost generated by patients with COPD represents 13.5% of the overall healthcare costs in the region.





The cost analysis by GMA grading (figure 3) clearly showed a steady increase of costs per patient/year with considered GMA risk grades: (1) low,  $\in 1284$ ; (2) moderate,  $\in 2944$ ; (3) high,  $\in 5933$ ; and (4) very high,  $\in 11537$ . Hospitalisation and pharmacy are the two items with highest impact on costs associated to GMA grading. It is worth to mention that we observed huge differences (7.41 times) between the users with high healthcare costs per year (PCT85:  $\in 16\ 131$ ), which represents 59% of the overall costs of patients with COPD, and patients with COPD below PCT85 ( $\in 2177$ ). Online supplementary table 8S indicates use of healthcare resources by GMA category.

## DISCUSSION

## Interpretation of the main findings

The main finding of the current study was the identification of comorbidities, expressed as GMA grades, as the covariate with highest discriminative impact on target events (mortality, hospitalisations) (table 2) and on yearly healthcare costs per patient (figure 3). These results can be relevant for commissioning of innovative healthcare services aiming at preventing materialisation of recent predictions on increases of healthcare impact of patients with COPD over the next 15 years.<sup>3 4</sup> However, they may also foster enhanced clinical management of individual cases with COPD.

It is of note that the performance of these models (table 2 and online supplementary figure S2–S7) shows quite acceptable goodness of fit (ie, C-statistics), indicating the potential of exploring synergies between population-health risk assessment (ie, GMA grading system) and clinical information to enhance health risk assessment and stratification in the clinical arena, as reported in Dueñas-Espín *et al.*<sup>15</sup> Ultimately, integration between registry data and electronic medical records of healthcare providers emerge as a high priority goal to properly pave the way towards personalised medicine for patients with chronic disorders.<sup>24</sup>

As expected, the covariate GMA showed a higher contribution in all causes hospital-related events than in only COPD-related admissions (table 2). It is of note that the latter represented only 22% of all admissions (table 1). While current GOLD recommendations address pulmonary events explaining frequent COPD exacerbations, the results of the current study suggest the need for further analyses aiming at identifying specific management needs for complex patients with COPD and comorbid conditions.

## Two complementary healthcare strategies

The distribution of patients with COPD in the regional health risk stratification pyramid (figure 1, right triangle) allows identification of two different scenarios with well-defined associated challenges.

### Patients with high health risk

The distribution of yearly healthcare costs per patient (figure 3) provides a strong rationale for targeting

patients close to the tip of the risk stratification pyramid as candidates for large-scale deployment of innovative services based on care coordination. However, two main limiting factors should be overcome in order to achieve proper designs of integrated care services.<sup>25</sup> First, poor comparability among interventions assessing effects on integrated care management for patients with COPD indicates an urgent need for standardisation of service workflows. Second is the limited healthcare impact of standard interventions addressed to patients at the tip of the pyramid,<sup>26</sup> that is, the low ratio between magnitude of the interventional effects and the resources devoted to achieve them, which may imply little healthcare value generation.<sup>67</sup> These two factors strengthen the need for further evidence on cost-effectiveness of well-defined integrated care interventions for complex chronic patients.<sup>27</sup>

#### Patients with low and moderate health risk

A better understanding of underlying mechanisms of comorbidity clustering in these patients emerges as a central need to effectively slow down patients' progress towards the tip of the pyramid. A natural consequence should be the development of efficient preventive interventions<sup>28</sup> aiming at delaying patients' worsening in terms of health risk scoring, which is recognised as a central unmet need for enhanced COPD management.

## Strengths and limitations of the study

The uniqueness of the current study is that it was carried out using registry data that allow population-based analyses of all patients with a given condition(s) in the region. It is of note that complexities involved in implementation and optimisation of large-scale health information technology systems often impede health assessment of the entire population in a real-world setting.<sup>29</sup>

We acknowledge, however, that the use of registry information alone reflects underdiagnosis of COPD and constitutes a significant limitation that may explain a rather low figure for COPD prevalence in the region.<sup>5 30</sup> The lack of clinical information, spirometric data and history of tobacco smoking reduces the potential for a proper characterisation of patients with COPD. As indicated above, the barriers associated to the existence of health information silos further strengthens the need for speeding up the current efforts to achieve real integration between clinical and registry data<sup>5</sup> that will open new avenues to enhance both medical knowledge and clinical practice.

Episodes of inpatient and outpatient care carried out in private hospitals were not available for analysis because private providers do not use the Personal Health Identification Number. Nevertheless, the vast majority (approximately 97.5% in 2015) of COPD hospitalisations are done in public hospitals.

#### Clinical impact and perspectives generated by the study

The current study confirms that prevalent chronic conditions such as cardiovascular disorders, type 2 diabetes mellitus-metabolic syndrome and/or anxiety-depression often occur in high-risk patients with COPD.<sup>10–12</sup> We believe that the study provides a strong rationale for further research on subject-specific health risk prediction and stratification aiming at early identification of patients with low to moderate health risk who are prone to develop comorbid conditions in order to enhance preventive management in a cost-effective manner.<sup>31 32</sup> The current research may contribute to foster future developments of GOLD recommendations<sup>5</sup> addressing non-pulmonary manifestations of COPD that should have a positive impact on both staging and management of complex chronic patients.

Moreover, the study reinforces the ongoing strategies aiming at speeding up the evolution of the current health surveillance system in Catalonia towards a Digital Health Framework conceptually formulated in Cano et al<sup>33 34</sup> with potential to articulate three categories of data: (1) outcomes from population-based health-risk predictive models; (2) healthcare and biomedical research knowledge resulting from integration of clinical, physiological and biological/molecular information; and (3) informal care information from in-place personal health folders<sup>15 17</sup> encompassing information on lifestyle, adherence profile, socioeconomic status, social support and environmental factors. It is envisaged that inclusion of all these covariates influencing patient health should markedly increase the predictive accuracy and facilitate clinical decision-making based on sound estimates of the prognosis of an individual.

## **CONCLUSIONS**

The current study provides a population-based analysis of 264830 patients with COPD based on administrative health registries in Catalonia. The results illustrate the high impact of comorbidities on undesirable clinical events. We believe that the results highly encourage further developments fostering interoperability between health registries and electronic medical records to enhance clinical risk prediction.

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