Socio-environmental correlates of physical activity in patients with chronic obstructive pulmonary disease (COPD)

Ane Arbillaga-Etxarri,1,2,3,4 Elena Gimeno-Santos,1,2,3 Anaël Barberan-García,5,6 Marta Benet,1,2,3 Eulàlia Borrell,7 Payam Dadvand,1,2,3 Maria Foraster,8,9 Alicia Marín,10 Mónica Monteagudo,11,12 Robert Rodríguez-Roisín,6 Pere Vall-Casas,13 Jordi Vilaró,14 Judith Garcia-Aymerich,1,2,3 on behalf of the Urban Training Study Group

ABSTRACT
Background Study of the causes of the reduced levels of physical activity in patients with COPD has been scarce and limited to biological factors.

Aim To assess the relationship between novel socio-environmental factors, namely dog walking, grandparenting, neighbourhood deprivation, residential surrounding greenness and residential proximity to green or blue spaces, and amount and intensity of physical activity in COPD patients.

Methods This cross-sectional study recruited 410 COPD patients from five Catalan municipalities. Dog walking and grandparenting were assessed by questionnaire. Neighbourhood deprivation was assessed using the census Urban Vulnerability Index, residential surrounding greenness by the satellite-derived Normalized Difference Vegetation Index, and residential proximity to green or blue spaces as living within 300 m of such a space. Physical activity was measured during 1 week by accelerometer to assess time spent on moderate-to-vigorous physical activity (MVPA) and vector magnitude units (VMU) per minute.

Findings Patients were 85% male, had a mean (SD) age of 69 (9) years, and post-bronchodilator FEV1 of 56 (17) %pred. After adjusting for age, sex, socio-economic status, dyspnoea, exercise capacity and anxiety in a linear regression model, both dog walking and grandparenting were significantly associated with an increase both in time in MVPA (18 min/day (p<0.01) and 9 min/day (p<0.05), respectively) and in physical activity intensity (76 VMU/min (p=0.05) and 59 VMUs/min (p<0.05), respectively). Neighbourhood deprivation, surrounding greenness and proximity to green or blue spaces were not associated with physical activity.

Conclusions Dog walking and grandparenting are associated with a higher amount and intensity of physical activity in COPD patients.

Trial registration number Pre-results, NCT01897298.

INTRODUCTION
Patients with COPD are substantially less active than their healthy peers1 due to progressive airflow limitation, abnormal gas exchange, skeletal muscle dysfunction, comorbidities and behavioural changes.2 However, maintaining a sufficient amount of physical activity is critical for the patients’ prognosis, quality of life and mortality.3 Thus, increasing physical activity has been recently included as a relevant health outcome of a wide range of interventions, including pharmacological treatment,4 pulmonary rehabilitation5 and interventions based on behavioural strategies such as self-monitoring, self-efficacy, motivation, goal setting and feedback.6 Of note, only one study so far has shown a sustained long term increase in physical activity, exclusively in a subgroup of patients who were not sufficiently active at baseline.7

It has been argued that this absence of effectiveness could be due to lack of knowledge concerning...
the causes of physical activity behaviour.8 Remarkably, the number of studies on physical activity determinants in COPD patients is lower than the number of studies assessing interventions to modulate this behaviour.3 Moreover, study of physical activity determinants in COPD patients has been limited to individual factors, mostly in the psychological and biological areas. This is in contrast with the commonly accepted socio-ecological models guiding most research on physical activity determinants in the general population and in patients with other chronic diseases.5

Socio-ecological models explain physical activity as being affected by psychological, biological and socio-environmental characteristics, and their interactions, both at the individual and population levels. To facilitate the study of socio-environmental characteristics and physical activity, three main dimensions have been described: social inequality, interpersonal relationships and neighbourhood characteristics.10 In patients with COPD, only social inequality has been investigated as a potential determinant of physical activity through the application of common measures of socio-economic status (SES), such as educational or occupational categories, and has shown inconsistent associations with physical activity.7 However, interpersonal relationships such as walking the dog or grandparenting have not previously been investigated despite the potential of interpersonal relationships to affect physical activity behaviour.10 Likewise, neighbourhood characteristics, such as the proximity of parks within walking distance or lower neighbourhood deprivation levels, have been associated with higher levels of physical activity in the general population,11 although they have not previously been tested in patients with COPD.

We aimed to assess the relationship of novel socio-environmental factors (dog walking, grandparenting, neighbourhood deprivation, residential surrounding greenness, and residential proximity to green or blue spaces) with the physical activity level of stable mild-to-very severe patients with COPD from five Mediterranean municipalities.

METHODS
Study design and population
This cross-sectional analysis used baseline data from the ongoing multicentre trial ‘Effectiveness of an intervention of urban training in patients with chronic obstructive pulmonary disease (COPD): a randomized controlled trial’ (NCT01897298).12 We first identified all patients with a COPD diagnosis (according to American Thoracic Society and European Respiratory Society (ATS/ERS) criteria)13 from 33 primary care centres and five tertiary hospitals of five seaside municipalities in Catalonia: Barcelona, Badalona, Mataró, Viladecans and Gavà (see online supplementary table S1 for their population and area characteristics). Exclusion criteria were: spending >3 months/year away from their home address, mental disability, severe psychiatric disease, comorbidity limiting survival to 1 year, or any other severe comorbidity. Between October 2013 and February 2015, all candidate patients were approached in random order within each municipality (Viladecans and Gavà were grouped together because they are adjacent municipalities). Patients were consecutively included until the end of the recruitment period specified for each geographical area based on the primary objective of the study. A total of 410 clinically stable (defined as at least 4 weeks without antibiotics or oral corticosteroids) patients with COPD were enrolled: 189 patients from Barcelona, 30 from Badalona, 73 from Mataró and 119 from Viladecans/Gavà. The study was approved by the Ethics Committees of all participating institutions and written informed consent was obtained from each participant.

Variables and instruments
All patients answered an interviewer-administered questionnaire about dog ownership (question: ‘Do you have a dog at home?’), dog walking (if positive to having a dog, ‘Do you walk the dog?’), having grandchildren (‘Do you have grandchildren?’) and grandparenting (if positive to having grandchildren, ‘Do you take care of them or collaborate in their care?’). The questionnaire also included data on education, marital status, working status, occupation, number of people living at home, smoking status, dyspnoea (assessed using the modified Medical Research Council scale (mMRC)), health-related quality of life (assessed using the COPD Assessment Test (CAT)), and anxiety and depression symptoms (assessed using the Hospital Anxiety and Depression scale (HAD)). SES was obtained from questions on employment characteristics, and classified using the National Statistics Socio-economic Classification into six categories (I, professional; II, managerial and technical; IIIN, skilled non-manual; III, skilled manual, IV, partly skilled; and V, unskilled occupations). Comorbidities were assessed from medical records. Body mass index (BMI) was obtained by physical examination. FEV1 and FVC before and after bronchodilator use, and distance walked in the 6-min walking test (6MWD) were determined following standardised methodology.

We geocoded patients’ addresses to assess individual measures of neighbourhood deprivation, residential surrounding greenness, and residential proximity to green or blue spaces. Neighbourhood deprivation was assessed using the Urban Vulnerability Index14 which is a measure of neighbourhood SES at the census tract level (median area of 0.26 km²) ranging from 0 (lowest) to 1 (highest level of neighbourhood deprivation).

To measure residential surrounding greenness, we used the Normalized Difference Vegetation Index (NDVI) which is an indicator of green (ie, photosynthetically active) vegetation density based on the difference between visible red and near-infrared surface reflectance. It was derived from Landsat 8 satellite images with a 30 m x 30 m spatial resolution during the greenness season in 2014. NDVI has a range from −1 to +1, with higher values indicating a high density of green vegetation.15 Residential surrounding greenness was abstracted as the average NDVI in 100, 300, 500 and 1000 m buffers from patients’ addresses calculated for both network and Euclidean distances. Average NDVI in a network distance of 300 m was used as the main measure of residential surrounding greenness; the remaining variables were included in sensitivity analyses.

We identified outdoor green (ie, green urban areas, agricultural land and pastures, and forests or country parks) and blue (ie, inland and non-inland water bodies) spaces of ≥0.25 ha. We calculated the network distance between participants’ home addresses and the closest green or blue space. Following previous studies16 and European Commission recommendations,17 we defined residential proximity as living within 300 m of a green or blue space (equivalent to a 15 min walk).

Physical activity was objectively measured using the Dynaport accelerometer (McRoberts, The Hague, The Netherlands), which has been shown to be a valid tool for measuring physical activity in COPD subjects.18 Patients were instructed to wear it attached to the centre of their lower back with an elastic strap for 1 week. A valid physical activity measurement was previously defined as a minimum of 3 days with at least 8 hours of wearing time per day.19 The main outcomes of the amount and intensity of physical activity were, respectively, time spent on...
moderate-to-vigorous physical activity (min/day, defined as activity of 3–6 Metabolic Equivalent Tasks (METs)) and vector magnitude units (VMU) per minute. We used three secondary outcomes of physical activity amount: steps per day, walking time (min/day) and time spent on physical activities (defined as ≥1.5 METs).

Statistical analysis

Sample size power estimation is detailed in the online supplementary file. To assess the association between socio-environmental determinants and physical activity, we first tested the bivariate relationship between each socio-environmental determinant (dog walking, grandparenting, neighbourhood deprivation, residential surrounding greenness, and residential proximity to green or blue spaces) and each physical activity variable (time spent on moderate-to-vigorous physical activity, VMU per minute, steps, walking time and time spent on physical activities) using Student’s t-test, ANOVA, Kruskal-Wallis, and Wilcoxon rank-sum tests, as appropriate. Second, a multivariable regression model was built for each physical activity variable. As the physical activity variables were normally distributed, we used linear regression. Each model included as exposure variables those socio-environmental characteristics that had exhibited a p value <0.2 in the corresponding bivariate analysis. Age, sex, education, marital status, socio-economic status, working status, smoking history, BMI, dyspnoea, anxiety, depression, FEV1, 6MWD, FEV1/FVC ratio, health-related quality of life (CAT score), diabetes, coronary heart disease, hypertension, musculoskeletal disorders, comorbidities that could affect walking, and municipality were tested as potential confounders. Model building combined step-forward and backward algorithms, and covariates were included in the final model if (i) they related to both the exposure and the outcome in bivariate analysis, (ii) they modified (>10% change in regression coefficient) the estimates of the remaining variables in the multivariable models, or (iii) there was consistent evidence in the literature of their association with physical activity. Goodness of fit was assessed by means of normality of residuals, heteroscedasticity, linearity, co-linearity, and identification of influential data.

Potential effect modification on the studied associations was evaluated by stratification of the final models according to sex, COPD severity stages (mild-to-moderate and severe-to-very severe), municipality and neighbourhood deprivation (categorised by its median value), and statistical testing of the interaction term. Sensitivity analysis was conducted (i) excluding subjects with extreme values (>99th percentile) in the accelerometer measures, (ii) using a mixed effects linear model to account for the potential correlation between subjects within a municipality as a random intercept, and (iii) using alternative distance measures to calculate residential surrounding greenness as detailed in the Methods section. Analysis was conducted using a complete case approach in Stata V14.0 (StataCorp, College Station, Texas, USA).

RESULTS

Patients were mostly male (85%) and had a mean (SD) age of 69 (9) years (table 1). According to the ATS/ERS classification, 9% of patients had mild, 53% moderate, 31% severe and 7% very severe COPD. According to the GOLD ABCD classification, 23% of patients were in group A, 31% in B, 13% in C and 33% in D. Patients walked a mean of 486 (95) m in the 6MWT. Only 12% and 38% of patients reported walking the dog or grandparenting, respectively. Half of the patients had green or blue spaces within 300 m of their homes. All patients fulfilled the criterion of wearing time per day (median wearing days 7, range 3–7; median recording time 14.9 hours, range 11.1–15 of 15 hours maximum from 7:00 to 22:00). Patients spent a mean of almost 3 hours/day on physical activity and half of this activity was at least of moderate intensity (table 1).

Figure 1 shows that dog walking and grandparenting were significantly associated with both higher amount and higher intensity of physical activity. However, neighbourhood deprivation, residential surrounding greenness, or residential proximity to green or blue spaces were not related to physical activity parameters. Online supplementary tables S2 and S3 show the bivariate associations between potential confounders and both exposures and outcomes. After adjusting for age, sex, SES,
smoking, dyspnoea, exercise capacity and anxiety, dog walking was significantly associated with an increase in moderate-to-vigorous physical activity of 18 min/day (p<0.01), and an increase in physical activity intensity of 76 VMUs/min (p=0.05), as compared to not walking a dog (table 2). Grandparenting was also related to both higher amount and higher intensity of physical activity after adjusting for potential confounders (table 2).

Very similar associations were obtained for steps and times engaged in walking and physical activity of any intensity (see online supplementary table S4). There was no interaction between dog walking and grandparenting in their association with physical activity amount or intensity. Goodness of fit tests did not reveal any abnormality.

After stratification by sex, the association between grandparenting and time in moderate-to-vigorous physical activity was observed only in men (p-for-interaction=0.02) (see tables 3 and online supplementary table S5). No differences were observed after stratification according to COPD severity stage (mild-to-moderate and severe-to-very severe), municipality and neighbourhood deprivation. Sensitivity analysis excluding subjects with extreme physical activity values (see online supplementary table S6) or using a mixed effects linear model (see online supplementary table S7) showed very similar estimates. Alternative buffers to calculate residential surrounding greenness did not provide additional information (see online supplementary figure S1).

**DISCUSSION**

We assessed the relationship between socio-environmental factors and the physical activity level of clinically stable mild-to-very severe patients with COPD and found, for the first time, that dog walking and grandparenting are both associated with a higher amount and intensity of physical activity in COPD patients. However, the characteristics of green or blue spaces surrounding patients’ homes were not associated with physical activity practice.

These results are consistent with previous research showing that dog walking is associated with higher physical activity in
both older healthy adults and patients with type 2 diabetes.20–22 Whether this association may be extrapolated into an effective intervention to increase physical activity in patients with COPD is still a matter of debate. A recent review highlighted that dog walking constitutes a specific behaviour different from other physical activities or even other types of walking because it has its specific barriers and facilitators and takes place in a specific individual, mental, physical, social and environmental context. For example, concerns about dogs coming into contact with children or other park users, crowded places, and large or uncontrolled dogs have been identified as barriers to dog walking. On the other hand, accessible appropriate public open spaces for dogs or neighbourhood walkability act as facilitators for this activity.20 Therefore, before interventions are designed for improving physical activity by means of dogs, research is needed to understand the specific characteristics of patients with COPD who walk their dogs and the factors influencing dog walking behaviour in this population.23

The patients with COPD who reported grandparenting exhibited higher levels of physical activity amount and intensity. Indeed, previous studies had already shown that grandparenting had significant benefits for grandparents’ physical and mental health,24–26 but this had never been tested in chronically ill patients, such as those with COPD. Interestingly, patients with COPD have reported that caring for and playing with grandchildren is a motivator to undertake a pulmonary rehabilitation programme.27 Thus, grandparenting may represent an excellent opportunity for promoting physical activity in this aging population. Our finding that the benefits for physical activity may be restricted to grandfathers needs a cautious interpretation because of a lack of statistical power in the female group (15% of the sample). We hypothesise that grandmothers caring for grandchildren may carry out domestic tasks, like cooking or feeding the grandchildren, which do not require intense physical activity, while grandfathers may spend more time outdoors, playing with grandchildren, or playing with them. Previous studies have reported different effects of grandparenting depending on sex.24–26

We did not find any association between residential surrounding greenness and residential proximity to green or blue spaces and physical activity in our patients with COPD. In the general adult population, the proximity and number of green spaces around the home has been related to higher levels of physical activity,11 but this association has not been shown to be consistent across studies.26 In fact, a recent study carried out in Barcelona showed that perceived proximity to green spaces, but not objective proximity, was associated with higher physical activity.28 It has been reported that the relationship between the built environment and physical activity characteristics is not equal for the entire population but modulated by sex, age, SES or working status.30 31 We hypothesise that, in subjects with limited exercise capacity like patients with COPD, the ability of a favourable built environment to stimulate physical activity could be lower than in healthy subjects. This is supported by a recent study showing that the aesthetics of the built environment were related to physical activity levels only in healthy subjects but not in those with chronic disease.32 Therefore, despite the increasing evidence of the association between the built environment and physical activity in the general adult population, qualitative and quantitative research is required to determine how the built environment affects behaviour in patients with COPD or other chronic diseases before considering the use of green or blue spaces to promote their physical activity.

Neighbourhood deprivation was not associated with physical activity in our study, which could be partially explained by the small variability in the urban vulnerability score in the five included municipalities. However, low SES was associated with a higher amount and greater intensity of physical activity, at variance with previous studies in patients with COPD from Barcelona that reported higher physical activity in the higher SES categories.3 Such a difference could be attributed to the fact that the current study included patients from middle-sized municipalities with population densities and historical urban backgrounds different from that of Barcelona, which could result in different social behaviour.33

The results of this study provide novel and original information to help enhance physical activity for patients with COPD in the clinical setting, highlighting the potential of interpersonal relationships as determinants of behaviour. Questioning the patients about their family duties and/or leisure time preferences can help professionals customise physical activity recommendations. In patients with diabetes for example, the necessity of encouraging dog walking or identifying other activities that promote a regular commitment to undertake physical activity has been recognized.34 Our results support the advice concerning dog walking, which can be used in patients who already own a dog but otherwise needs a deep understanding of patient preferences and fears. Grandparenting and, in general, encouraging grandparents to have an active role in family

Table 2 Adjusted association* between socio-environmental characteristics and physical activity amount and intensity in COPD patients

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Time in physical activity of moderate-to-vigorous intensity (min/day)</th>
<th>VMU/min</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>β (95% CI)</td>
<td>p Value</td>
</tr>
<tr>
<td>Dog walking</td>
<td>19 (7 to 32)</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>Grandparenting</td>
<td>8 (−0.3 to 17)</td>
<td>0.06</td>
</tr>
<tr>
<td>Age (years)</td>
<td>−0.7 (−1.3 to −0.2)</td>
<td>0.01</td>
</tr>
<tr>
<td>Sec. male</td>
<td>3 (−9 to 15)</td>
<td>0.62</td>
</tr>
<tr>
<td>Low SES: IV, V</td>
<td>10 (1 to 20)</td>
<td>0.03</td>
</tr>
<tr>
<td>Active smoking</td>
<td>−20 (−30 to −10)</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>Dyspnoea (mMRC, 0–4)</td>
<td>−11 (−16 to −5)</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>6MWD (m)</td>
<td>0.1 (0.1 to 0.2)</td>
<td>&lt;0.01</td>
</tr>
<tr>
<td>Anxiety (HAD-A score, 0–21)</td>
<td>1.1 (0.01 to 2.2)</td>
<td>0.04</td>
</tr>
</tbody>
</table>

*Each column is a single linear regression multivariable model. Coefficients are expressed as minutes of time in activity or VMU/min (one at each time), per (i) each unit of the continuous covariates, or (ii) a change with respect to reference category in categorical covariates.

Chronic obstructive pulmonary disease

Table 3 Adjusted* association between socio-environmental characteristics and physical activity amount and intensity in COPD patients according to sex, COPD severity, municipality and neighbourhood deprivation

<table>
<thead>
<tr>
<th>Time in physical activity of moderate-to-vigorous intensity (min/day)</th>
<th>By sex</th>
<th>By COPD severity</th>
<th>By municipality</th>
<th>By neighbourhood deprivation</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Women</strong></td>
<td><strong>Men</strong></td>
<td><strong>By sex</strong></td>
<td><strong>By COPD severity</strong></td>
<td><strong>By sex</strong></td>
</tr>
<tr>
<td>Dog walking</td>
<td>n=61 (15%)</td>
<td>n=350 (85%)</td>
<td>FEV1&lt;50%</td>
<td>FEV1&lt;50%</td>
</tr>
<tr>
<td>18 (9 to 45)</td>
<td>24 (9 to 39)</td>
<td>15 (1 to 30)</td>
<td>31 (6 to 56)</td>
<td>28 (8 to 48)</td>
</tr>
<tr>
<td>Grandparenting</td>
<td>n=61 (15%)</td>
<td>n=350 (85%)</td>
<td>FEV1&lt;50%</td>
<td>FEV1&lt;50%</td>
</tr>
<tr>
<td>−16 (−38 to 6)</td>
<td>12 (3 to 22)</td>
<td>−12 (1 to 23)</td>
<td>1 (−14 to 15)</td>
<td>1 (−12 to 15)</td>
</tr>
</tbody>
</table>

**VMU/min**

<table>
<thead>
<tr>
<th>By sex</th>
<th>By COPD severity</th>
<th>By municipality</th>
<th>By neighbourhood deprivation</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Women</strong></td>
<td><strong>Men</strong></td>
<td><strong>By sex</strong></td>
<td><strong>By COPD severity</strong></td>
</tr>
<tr>
<td>Dog walking</td>
<td>n=61 (15%)</td>
<td>n=350 (85%)</td>
<td>FEV1&lt;50%</td>
</tr>
<tr>
<td>75 (88 to 273)</td>
<td>110 (22 to 196)</td>
<td>99 (12 to 187)</td>
<td>43 (−109 to 197)</td>
</tr>
<tr>
<td>Grandparenting</td>
<td>n=61 (15%)</td>
<td>n=350 (85%)</td>
<td>FEV1&lt;50%</td>
</tr>
<tr>
<td>7 (14 to 142)</td>
<td>5 (−84 to 94)</td>
<td>13 (−58 to 84)</td>
<td>173 (24 to 322)</td>
</tr>
</tbody>
</table>

*Models adjusted for age, sex, socio-economic status, dyspnoea, 6MWD and HAD-anxiety.
†6MWD, 6-min walking distance; HAD, Hospital Anxiety and Depression scale; UVI, Urban Vulnerability Index; NC, non-computable; VMU, vector magnitude units.

Chronic obstructive pulmonary disease
Chronic obstructive pulmonary disease

Gimeno-Santos, Jaume Torrent-Pallier; FCS Blanquerna, Universitat Ramon Llull, Barcelona: Jordi Vilard; Servei de Pneumologia, Hospital Clinic de Barcelona, Barcelona; Anel Barberan-Garcia; Universitat de Barcelona: Robert Rodriguez-Roisin; Hospital del Mar, Institut Hospital del Mar d'Investigacions Mèdiques (IMIM), Barcelona: Eva Balcells, Diego A Rodriguez, Hospital Universitari Germans Trias i Pujol, Badalona: Alicia Marin; Hospital de Matarranya, Consorci Sanitari del Maestrat, Pilar Ortega; Hospital de Viladecans, Viladecans: Nuria Celorio; Institut Universitari d'Investigació en Atenció Primària Jordi Gol (IDIAP Jordi Gol): Mònica Monteagudo, Nuria Montellà, Laura Muñoz, Pere Toran; Centre d’Atenció Primària Viladecans 2, Institut Català de la Salut, Viladecans: Pere Simonet; Centre d'Atenció Primària Passeig de Sant Joan, Institut Catalá de la Salut, Barcelona: Carmen Jané, Carlos Martín-Cartera; Centre d’Atenció Primària Sant Roc, Institut Català de la Salut, Badalona: Eulàlia Borrell; Universitat Internacional de Catalunya (UIC), Barcelona: Pere Vall-Casas.

Contributors AAE and JGA wrote the initial draft. AAE, MB and JGA had full access to the data and carried out statistical analysis. AAE, ABG, EB, AM and MM contributed to data collection and coordination. All authors (i) provided substantial contributions to the conception or design of the work, or the acquisition, analysis or interpretation of data for the work, (ii) revised the manuscript for important intellectual content, (iii) approved the final version, and (iv) agreed to be accountable for all aspects of the work.

Funding The study was funded by grants from Fondo de Investigación Sanitaria, Instituto de Salud Carlos III (ISCIII), Spain (PI11/00140 and P11/01409); Sociedad Española de Neumología y Cirugía Torácica (SEPAR) (14AP/2011 and 2011/2017); Societat Catalana de Pneumologia (Ajuts al millor projecte en Atenció Primària, 2015; AP2015/91); Fundació Jordi Gol (IDIAP Jordi Gol): Mònica Monteagudo, Nuria Montellà, Laura Muñoz, Pere Toran; Centre d’Atenció Primària Viladecans 2, Institut Català de la Salut, Viladecans: Pere Simonet; Centre d’Atenció Primària Passeig de Sant Joan, Institut Català de la Salut, Barcelona: Carmen Jané, Carlos Martín-Cartera; Centre d’Atenció Primària Sant Roc, Institut Català de la Salut, Badalona: Eulàlia Borrell; Universitat Internacional de Catalunya (UIC), Barcelona: Pere Vall-Casas.

Patient consent Obtained.

Ethics approval Comitè d'Ètica d’Investigació Clínica Hospital Universitari Germans Trias i Pujol; Comitè d’Investigació Hospital Clínic; Comitè d’Ètica d’Investigació Clínica IDIAP Jordi Gol; Comitè d’Ètica d’Investigació Clínica Hospital de Matarranya; Comitè d’Ètica d’Investigació Clínica Parc de Salut MAR; Comitè d’Ètica d’Investigació Clínica Hospital Universitari de Bellvitge.

Provenance and peer review Not commissioned; externally peer reviewed.

Open Access This is an Open Access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/}

REFERENCES