

### RESEARCH ARTICLE

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# Long-term smoking increases the need for acute care among asthma patients: a case control study

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#### **Abstract**

**Background:** To examine risk factors for asthma patients' emergency room (ER) visits in a well organized asthma care setting.

**Methods:** A random sample of 344 asthma patients from a Pulmonary Clinic of a University Hospital were followed through medical records from 1995 to 2006. All the ER visits due to dyspnea, respiratory infections, chest pain, and discomfort were evaluated.

**Results:** The mean age of the study population was 56 years (SD 13 years), 72% being women. 117 (34%) of the patients had had at least one ER visit during the follow-up (mean 0.5 emergency visits per patient year, range 0–7). Asthma exacerbation, lower and upper respiratory infections accounted for the 71% of the ER visits and 77% of the hospitalizations. The patients with ER visits were older, had suffered longer from asthma and more frequently from chronic sinusitis, were more often ex- or current smokers, and had lower lung function parameters compared to the patients without emergency visits. Previous (HR 1.9, Cl 1.3-3.1) and current smoking (HR 3.6, Cl 1.6-8.2), poor self-reported health related quality of life (HRQoL) (HR 2.5, Cl 1.5-4), and poor lung function (FEV1 < 65%, HR 2.2, Cl 1.3-3.7) remained independent risk factors for ER visits after adjustment for age and gender.

**Conclusions:** Asthma patients who are or have been long-term smokers are more likely to require ER care compared to never smokers.

Keywords: Obstructive lung disease, Emergency visits, Hospitalizations, Smoking cessation

#### **Background**

Modern pharmaceuticals such as inhaled steroids, long-acting  $\beta 2$ -agonists, their combinations, and leukotriene antagonists have allowed high quality asthma management [1,2]. The present concept of asthma management is to intensify the treatment until full control has been achieved. In Finland, asthma management has developed not only because of efficient medication but also due to a nationwide educational programme during years 1994–2004 [3]. The program was focused to improve the patients' self-management capabilities by implementation of a self-management plan, and to update the referral and treatment networks in primary health care [3]. Improved care was proven by several indicators, such as decrease in mortality, disability pensions, and hospital days, and other savings in health care. Better control of asthma management was also

shown as a decreased proportion of young men exempted from military service [3,4].

Among younger adults exacerbations requiring hospitalizations frequently reflect a failure in asthma management. At older age, however, co-existing chronic diseases are common [5,6]. Co-morbidities, underdiagnosis and undertreatment of asthma, and also in turn potential adverse events due to long-term asthma medication complicate the treatment of asthma and thus, increase the risk of an exacerbation [5-7]. Obesity, gastro-oesophageal reflux, sleep apnoea, hyperlipidemia, diabetes, and osteoporosis have been associated both to aging and asthma [7,8]. These conditions are also closely related to adverse effects of corticosteroids. Asthma has also been reported to double the risk of coronary heart disease [8,9]. Depression and anxiety are found more often in asthma patients and further, depression and anxiety are associated with poor asthma control, long-term work disability, and increase in emergency treatments [8,10-12]. Many of elderly asthma

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patients have been long-term smokers and developed irreversible airway obstruction [13]. Overall, elderly asthma patients have poorer self-estimated health status compared to that in the age matched general population. Inadequate asthma treatment is also more common in older than in younger age groups [7].

In the present study, we examined the factors that are associated with emergency visits in an elderly asthma population. In these events, asthma was the primary cause or the primary contributing factor of the visit. All visits due to dyspnea, airway infection, chest discomfort, or palpitation were included. We concentrated in the events that require the emergency department and thus, urgent actions to prevent a serious outcome [14].

#### **Methods**

#### Study population

Adult asthma patients who had visited a Pulmonary Clinic of a University Hospital during the years 1995-2006 were invited to a 10 year follow-up study. Recruitment strategy and diagnostic criteria have been described previously in detail [13]. A total of 1855 patients participated in the study. Patients asthma diagnosis was confirmed and a random sample the patients (N = 344) were studied in detail regarding their emergency treatment history. We did a retrospective comparative study between asthma patients with emergency visits (N=117, those who had ≥1 emergency visit during the years 1995-2005) and asthma patients without emergency visits (N=227, patients without emergency visits). The event free time was accounted for each patient starting from the diagnosis or year 1995 in the case diagnosis was made earlier and ending to the first emergency visit or at the latest year 2005 when the patients were contacted by the study personnel.

The study population was a random sample (N = 344) of the original asthma population (N = 1855). In the original cohort the proportion of women was 69%, mean age at diagnosis 44 years, and at recruitment 55 years.

#### Assessment of emergency visits

Medical records from all health-care providers that had treated the patient were merged retrospectively. Referral diagnoses were used to identify all the ER visits at the emergency departments of the hospitals open 24/7. Visits in the primary health care were not included. Discharge diagnosis given by the physician of the emergency department was used to refine the diagnoses. All visits to the emergency clinics at hospitals open 24/7 due to asthma or respiratory infection were included into the analysis as the primary endpoint. All the visits due to upper or lower respiratory or cardiovascular systems (asthma exacerbation, respiratory infections, cardiac arrhythmias and palpitations, ischemic heart disease or heart failure, unspecified dyspnea, chest discomfort or

pain) were considered as the asthma-related and were recognized in the analysis as the ER visit. Information on age, gender, BMI, smoking history and pack years, onset of chronic diseases and lung function parameters [13] were collected with a structured manner. ATS/ERS guidelines and national reference values were used in interpretation of spirometer results [15,16]. In addition, the last scheduled contact to health care prior the emergency visit was evaluated.

#### Assessment of co-morbidities

Information on asthma related conditions (allergic rhinitis, asetylsalicylic intolerance, atopic eczema, chronic sinusitis, COPD, and nasal polyps), and other co-morbidities (cardioor cerebrovascular disease, diabetes, psychiatric diseases, and alcohol abuse) were gathered in the medical records. 'Asthma-COPD overlap' diagnosis was given those patients who had first asthma, but who in addition developed persistent irreversible airway obstruction (FEV<sub>1</sub>/FVC < 70%) due to long-term smoking [17]. 'Allergic rhinitis' was diagnosed by a pulmonologist or a rhinologist and was based on IgE mediated allergy and rhinitis symptoms. 'Asetylsalicylic intolerance' was diagnosed by a pulmonologist and was based on positive symptom history. 'Atopic eczema' was diagnosed by a pulmonologist or a dermatologist according to eczema manifestation on typical skin areas. 'Chronic sinusitis' and 'nasal polyposis' was diagnosed by a rhinologist according to clinical symptoms and findings. Coronary or cerebrovascular disease, or peripheral arterial occlusive disease were categorized as 'cardiovascular disease'. 'Cerebrovascular disease' (CVD) was diagnosed by a neurologist and was based on a stroke, brain haemorrhage or a transient ischemic attack. 'Coronary disease' included the patients who had been diagnosed with a myocardial infarct, an acute coronary syndrome, or angina pectoris by an internist. Both type I and type II diabetes were categorized as 'diabetes'. 'Psychiatric disorders' included patients with a medication for psychotic disorders, depression, or anxiety. 'Alcohol abuse' included patients with chronic alcoholism and treatment because of alcohol related disorder.

#### Assessment of self-reported information

Self-reported smoking status (never, ex-smoker, current smoker and pack years) was gathered by a questionnaire. In addition, health-related quality of life (HRQoL) was measured with an airway symptom specific questionnaire (AQ20 questionnaire) including 20 items [18-21]. "Yes" responses were scored as 1, and "no" and "not applicable" as 0. The summary score varied from 0 (=no symptoms or worries over the disease) to 20 (=full range of symptoms and worries about the disease). The summary scores were normally distributed among the participants. At recruitment all participants returned the questionnaire. Ambiguous or missing responses per question ranged

between 1% and 2%. HRQoL was measured at the study recruitment and on average 3.3 (SD 2.9) years after the latest emergency visit. The patients also reported their working status (at work, on disability pension, on old-age pension).

The study approach was approved by the Coordinating Ethics Committee of the Hospital District and the permission to conduct this research was granted by the University Central Hospital. The study population had given their written informed consents for the study.

#### Statistical analysis

The SPSS computer package (version 16.0) was used to compute differences in demographic and clinical variables. Continuous variables were compared using the non-parametric Mann–Whitney U –test and categorical variables using the Chi square test.

All associated clinical variables were included in the Cox proportional hazard model to determine the independent predictors for the first emergency visit. The respective estimated hazard ratios (HR) with confidence intervals (CI) of 95% are reported. Differences were considered statistically significant when the p value was less than 0.05.

Cumulative emergency visit free time (Figure 1) among asthma patients was stratified according to their smoking status and adjusted for age, gender, duration of disease,  $\text{FEV}_1$  of predicted, chronic sinusitis, and the number of co-morbidities.

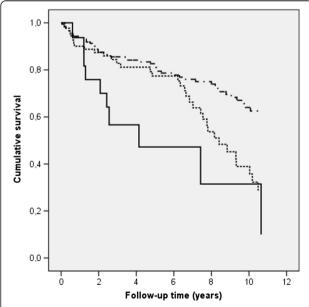


Figure 1 Cumulative emergency visits free time among asthma patients who were current smokers (solid line), ex-smokers (dotted line) and never smokers (dashed line) using the Cox regression model adjusted for age, gender, duration of disease, FEV1, chronic sinusitis, and number of co-morbidities.

#### Results

#### Frequency of emergency (ER) visits

A random sample (N = 344) of elderly (mean age of 56 years, 70% women) asthma patients of a Pulmonary Clinic of a University Central Hospital were included in the analysis. Their asthma related ER visits were followed through medical records since the asthma diagnosis or at the most, for 10 proceeding years (in total 1398 patient years). Altogether 117 patients (34%) had one or more (range 1-12) asthma related ER visits. These patients were considered as cases (a mean of 2.5 emergency visits per patient). Ninety-four (80%) of the cases had 1-2, and 23 (20%) had three or more asthma related ER visits. During the follow-up, the cases had in total 295 asthma-related emergency visits, a mean of 0.5 ER visits (range 0–7) per year per patient. The asthma patients with no asthma-related ER visits during the follow-up comprised the control group (N = 227).

#### Differences in clinical characteristics

The patients with ER visits were on average four years older and had also suffered asthma on average four years longer than the controls (Table 1). The cases were more frequently current (14.5% vs 4.0%) or ex-smokers (38.2% vs 27.1%,  $p \le 0.001$  for the trend), reported more pack years (13.2 vs 8.4, p = 0.02), displayed lower lung functions in spirometry (mean FEV1 75% vs 87% of predicted, p < 0.001), and were more frequently diagnosed with COPD second to asthma (23% vs. 8%, p < 0.001) than the controls. The mean BMI did not differ statistically significantly between the groups. A trend of an excess of co-morbidities (cardiovascular diseases, psychiatric disorders, or alcohol abuse) was found among cases compared to controls, even though not statistically significant. Except of chronic sinusitis, asthma associated allergic conditions such as allergic rhinitis, ASA intolerance, nasal polyposis, or atopic eczema did not associate with ER visits.

In the end of the retrospective follow-up, the cases reported poorer respiratory specific HRQoL than the controls (total score 8.9 vs. 6.3, p = 0.001). Further, those with  $\geq 3$  ER visits had lower HRQoL than those with 1-2 visits (total score mean 11.2 vs. 8.3, p = 0.003) and their PEF values were lower than that among the controls (71.9% vs. 82.9% of the predicted, p < 0.0.001).

There were no statistically significant differences in work status between the cases and controls. Among the cases 39% were in active workforce, 35% were on oldage, and 21% on disability pension. Among the controls, the corresponding proportions were 45%, 28%, and 14%, respectively.

When the participants' use of health care services were analysed prior to the ER visit, the last scheduled visit due to asthma was made an average 115 days

Table 1 Clinical characteristics of the asthma patients who had visited emergency clinic at least once during the follow-up compared to those who had not

Clinical characteristics	Emergency visit	Emergency visit			
	Yes	No	p-value <sup>1</sup>		
N of patients	117	227			
Proportion of women	65.8%	75.3%	0.06		
Age at diagnosis (mean $\pm$ SD)	42.6 ± 16.2	43.7 ± 17.1	not significant		
<b>Age at recruitment</b> (mean ± SD)	58.5 ± 11.8	$54.7 \pm 12.6$	0.006		
<b>Duration of asthma, years</b> (mean $\pm$ SD)	$15.7 \pm 14.3$	11.1 ± 12.0	0.001		
<b>BMI</b> (mean ± SD)	$26.8 \pm 5.5$	$26.5 \pm 5.6$	not significant		
Smoking status (%)					
never	47.3%	68.9%	0.001(trend)		
ex smoker	38.2%	27.1%			
current smoker	14.5%	4.0%			
Pack years (mean $\pm$ SD)	$13.2 \pm 19.7$	$8.4 \pm 17.4$	0.02		
COPD as second diagnosis (%)	23.1%	8.4%	< 0.001		
<b>Baseline FEV1 percentage of expected</b> (mean $\pm$ SD)	75.3 ± 21.1	$86.6 \pm 17.2$	< 0.001		
Baseline FVC percentage of expected (mean $\pm$ SD)	82.9 ± 17.2	$91.3 \pm 14.6$	< 0.001		
Baseline PEF percentage of expected (mean $\pm$ SD)	$80.7 \pm 20.3$	$94.0 \pm 19.6$	< 0.001		
Baseline FEV1/FVC ratio (mean ± SD)	$73.4 \pm 11.8$	$77.9 \pm 10.5$	0.001		
<b>HRQoL score</b> (mean ± SD)	$8.9 \pm 4.7$	$6.3 \pm 4.5$	< 0.001		
Proportion (%)of patients with					
no co-morbidities	59.8	71.8	0.06		
1 co-morbidity	33.3	23.3			
≥2 co-morbidities	6.8	4.8			
Proportion (%) of patients with co-existing					
Cardiovascular disease <sup>2</sup>	16.2	9.3	0.06		
Psychiatric disorder <sup>3</sup>	21.4	16.3	not significant		
Diabetes	3.4	5.7	not significant		
Alcohol abuse	7.0	4.0	not significant		
Chronic sinusitis	16.1	8.5	0.04		
Allergic rhinitis	39.3	35.1	not significant		
Allergic exzema	19.3	20.9	not significant		
ASA intolerance	10.4	6.6	not significant		
Nasal polyposis	10.3	8.0	not significant		

<sup>&</sup>lt;sup>1</sup>Continuous variables were compared using nonparametric Mann–Whitney U-test and categorical variables using the Chi-Square test.

(range 1–1588) prior. The last visit due to asthma was made in 64% in specialist care, and in 36% in primary health care, communal or occupational health care or at private doctor's office (Table 2). In 93% of the emergency visits, the last control had occurred within one year prior to the ER visit and no statistically significant difference was observed between primary health care and specialist care among those whose latest control exceeded one year (Table 2).

#### Reasons for ER visits among elderly asthma patients

The reason for the ER visits were analysed by using both the referral information and the more refined discharge diagnosis data. According to the both referral and the discharge diagnoses, the majority of emergency visits (47.4% and 44.6%, respectively) were due asthma exacerbation (Table 3). Asthma exacerbation was also the most common reason in hospitalizations (N = 168, 57% of all the visits). 22.4% of the visits were referred to emergency

<sup>&</sup>lt;sup>2</sup>Included coronary disease, cerebrovascular disease, and peripheral arterial occlusive disease.

<sup>&</sup>lt;sup>3</sup>Medication used.

Table 2 Last scheduled, asthma-related visit in primary health care or in specialist care before the ER visit

	L	Last scheduled visit before the ER visit			
	<1 year prior N = 250 (%)	≥1 year prior N = 19 (%)	TotalN = 269 (%)		
Primary health care	88 (35.2)	8 (42.1)	96 (35.7)		
Specialist care	162 (64.8)	11 (57.9)	173 (64.3)		
Total	250 (100)	19 (100)	269 (100)		

room care with a suspicion of respiratory infection, sinusitis or fever. The proportion of infections did not change in the ER evaluation. 14.1% of the ER visits was estimated to be of cardiovascular origin (pulmonary embolism, arrhythmia, ischemic coronary disease, or heart failure). 13% of the visits remained unspecified (dyspnoea, thoracic pain, hyperventilation, haemoptysis) when the patient was discharged from the emergency department.

The role of clinical determinants in the occurrence of ER visits was estimated by using the Cox-regression

analysis. Previous (HR 1.8) and current smoking (HR 3.9) compared to never smoking, impaired lung functions (FEV $_1$  < 65%, HR 2.9) compared to normal lung function, secondary diagnosis of COPD (HR 2.3), and poor HRQoL (HR 1.4) increased the risk significantly (Table 4). Current smoking (HR 3.6), poor HRQoL (HR 2.5), and FEV $_1$  < 65% (HR 2.2) remained independent risk factors for ER visit when the model was adjusted for gender, duration of asthma, cardiovascular disease, chronic sinusitis, and COPD. The ER visit free time was significantly shorter

Table 3 Referral and discharge diagnoses of the emergency visits made by the asthma patients

Diagnosis	Reference visit N = 117 (%)	All visits N = 295(%)	Visits leading to hospitalization N = 168 (%)				
Referral							
Asthma exacerbation	50 (42.7)	140 (47.4)	87 (51.8)				
Arrhythmias	5 (4.3)	22 (7.5)	4 (2.4)				
Chest pain	15 (12.8)	25 (8.5)	11 (6.5)				
Dyspnoea	19 (16.2)	40 (13.5)	26 (15.5)				
Fever	5 (4.3)	26 (8.8)	15 (8.9)				
Respiratory infection	17 (14.6)	33 (11.2)	22 (13.1)				
Sinusitis	5 (4.3)	7 (2.4)	2 (1.2)				
Other causes	1 (0.9)	2 (0.7)	1 (0.6)				
Total	117 (100)	295 (100)	168 (100)				
Emergency discharge							
Asthma exacerbation	50 (42.7 )	135 (44.6 )	86 (51.2)				
Asthma-COPD overlap	6 (5.1)	8 (2.9)	5 (3.0)				
Pneumonia	11 (9.4)	22 (8.7)	21 (12.5)				
Unspecified respiratory infection	9 (7.7)	26 (8.6)	12 (7.1)				
Otitis	1 (0.9)	6 (2.2)	2 (1.2)				
Sinusitis	4 (3.4)	12 (4.2)	3 (1.8)				
Dyspnoea	3 (2.6)	4 (1.9)	2 (1.2)				
Pulmonary embolism	3 (2.6)	4 (1.3)	4 (2.4)				
Arrhythmia	4 (3.4)	20 (6.4)	3 (1.8)				
Coronary disease or suspicion	9 (7.7)	15 (4.8)	10 (5.9)				
Heart failure	1 (0.9)	4 (1.6)	4 (2.4)				
Thoracic pain	8 (6.8)	11 (3.8)	2 (1.2)				
Allergic reaction	1 (0.9)	4 (1.3)	1 (0.6)				
Hyperventilation	4 (3.4)	6 (2.2)	2 (1.2)				
Haemoptysis	1 (0.9)	2 (0.6)	1 (0.6)				
Other causes	2 (1.8)	16 (5.4)	10 (6.0)				
Total	117 (100)	295 (100)	168 (100)				

Table 4 Hazard ratios and their 95% confidence intervals (CI) for patient characteristics that explained emergency visits among the asthma patients by the Cox regression analyses

	N of patiens <sup>1</sup>	% of patients with emergency visits	Crude HR	95% CI	P-value	Adjusted HR	95% CI	P-value
Male gender	96	34.2%						
Female gender	248	65.8%	0.68	0.46-1.00	0.051	1.11	0.67-1.84	NS
Age until first emergency care visit or at evaluation (increase of 1 year)			1.02	1.00-1.03	0.051	1.01	0.99-1.03	NS
Duration of asthma								
<15 years	231	60.0%	ref. 1.00			ref. 1.00		
>=15 years	100	40.0%	1.06	0.72-1.56	NS	1.29	0.82-2.03	NS
FEV1% of expected								
> 80%	164	42.7%	ref. 1.00			ref. 1.00		
-EV1 65-80%	65	24.5%	1.90	1.16-3.11	0.011	1.57	0.91-2.71	NS
FEV1 < 65%	54	32.7%	2.90	1.85-4.54	<0.001	2.20	1.29-3.74	0.004
Smoking								
Never	174	47.3%	ref. 1.00			ref. 1.00		
Ex-smoker	90	38.2%	1.75	1.14-2.66	0.10	1.86	1.30-3.06	0.015
Current	23	14.5%	3.94	2.20-7.07	<0.001	3.59	1.58-8.15	0.002
AQ_20								
score 12 or less	290	74.4%	ref. 1.00			ref 1.00		
score 13 or more	54	25.6%	2.17	1.41-3.33	<0.001	2.51	1.52-4.15	<0.001
CVD								
No	304	83.8%	ref. 1.00			ref. 1.00		
⁄es	40	16.2%	1.47	0.89-2.44	NS	1.61	0.91-2.91	NS
Sinusitis								
No	299	83.9%	ref. 1.00			ref 1.00		
⁄es	47	16.1%	1.47	0.87-2.51	NS	1.10	0.60-2.02	NS
COPD as a second diagnosis								
No	298	76.9%	ref. 1.00			ref 1.00		
r'es	48	23.1%	2.32	1.45-3.71	<0.001	0.60	0.29-1.23	NS

among current smokers compared to that among the examd never smokers (Figure 1).

#### Discussion

In this hospital-based adult asthma study population, one third of the patients had had an asthma related ER visit (mean 0.5 visits per patient per year). Asthma exacerbation and upper or lower respiratory tract infections covered 72% of the ER visits. Rest of the visits were due to other respiratory (15%) or cardiovascular events (13%). 57% of the ER visits led to hospitalization of the patient. The patients with emergency visits had experienced asthma longer (on average 16 years at the recruitment) than the control group (on average 11 years), and their asthma was more severe according to the lung function test results (FEV<sub>1</sub> 75% vs. 87% of the predicted, respectively). 59% of the patients with ER visits had been in special care prior to the ER visit also reflecting the severity of asthma. Asthma patients with ER visits were more frequently ex- or current smokers and displayed asthma/ COPD overlapping features. Poor airway specific HRQoL was associated with the number of ER visits. Current smoking, poor HRQoL, and moderate to severe airway obstruction (FEV1 less than 65%) remained independent risk factors for ER visits in the cohort.

Since the availability of health care services has an impact on ER visits and our study design was completely based on retrospective data, the analysis was conducted only in one hospital district. The number of study subjects was small which decreased variability between the patients. Also, the small recruitment rate was limited and it was not possible to analyse further the nonresponders. The follow-up time, however, was 1398 patient years, which increased the informativeness of the study design and the robustness of the major findings. When frequent visitors of ER have been studied in US, several of them had low income, inadequate medication, and no written asthma action plan [22]. By the implementation of the National Asthma Program special attention was paid to improve patients' capabilities to selftreatment and to the health care organisations to avoid simple caps in the asthma treatment. More than 90% of the ER visitors had had a schedule appointment due to asthma within a year and 57% of all visits led to hospital admissions suggesting rather well organised asthma care in the region.

Asthma and asthma related co-morbidities have shown to be important determinants of acute care. Co-morbidities have been associated in 9% of all ER room visits and 6% of all hospitalisations in asthma [23]. Our study supports the role of co-morbidities and expands findings on smoking and the level of bronchus obstruction as risk factors of emergency care. Current smoking already in mild asthma has been associated with impaired response to inhaled

corticosteroids [24-26]. With the same dose these patients report poorer asthma control including more frequent symptoms and lower peak flow values than their non-smoking counterparts. Furthermore, smoking cessation has been reported to gain better lung function in asthmatics compared to those asthma patients who continue to smoke [24-26]. Differential diagnostics with other conditions and overlap with COPD has been previously reported as a challenge for diagnosis of asthma, while co-morbidities being more of a challenge for asthma management [5].

The importance of obstruction was shown also in a prospective study of 554 asthma patients (18-55 years of age) identified from a large health maintenance organization. The patients were followed for 30 months to identify risks for acute care. The strongest predictors were airflow obstruction (RR 4.3 when FEV1<60%) and again, current smoking (RR 1.6) [27]. Typically spirometer measurements are performed when the patient is at his best, not during the exacerbations. Thus most likely declined spirometer values reflected permanent loss of lung function among this patient group. Whether the ER visit resulted due to severe disease or insufficient medication can not be determined without detailed information on used asthma medication. Other risk factors were regular exposure to solvents and the ownership of dog or cat when being skinprick test positive. In the present older and hospital based patient cohort, allergic rhinitis or atopic eczema did not did not associate with ER visits or hospitalizations.

Allergen sensitation and exposure with simultaneous viral infection have been reported to increase the risk (OR 8.4) of hospital admissions among adult asthma patients [28]. In the present study chronic sinusitis increased the risk and acute infections explained one fourth of all visits. Tobacco smoke has been shown to increase viral replication [29,30] while inhaled corticosteroids protect against rhinovirus induced airway inflammation but not against viral replication which might partly explain poor treatment response among smokers [31]. The cases had more common co-morbidities, but compared to infections for example the significance of cardiovascular disease was much less, 13% of the visits. Diabetes, psychiatric disorders, and alcohol abuse, even though common risk factors of poorer control of asthma and acute care, were not increased in the ER-group (Table 1). Many co-morbidities such as anxiety and depression also have an impact on HRQoL and poor HRQoL on the other hand is a risk factor for ER visits [22,32,33]. Poor HRQoL also associates with known asthma triggers such as respiratory infections, environmental irritants, emotions, allergens, weather, and exercise. The patients who need acute care have frequently multiple triggers [34].

Recently, increase in severity of asthma has been suggested to be associated with an increase of obesity [35]. Obesity is associated with gastroesophageal reflux

disease, hyperlipidemia, and diabetes with may in turn worsen the asthma symptoms [9]. In the study of Mosen et al. obese individuals with persistent asthma reported more frequently poorer asthma-related quality of life and more asthma-related hospitalisations than patients with normal weight [36]. Obesity remained an independent risk for hospitalizations after adjustment for age, sex, smoking status, use of corticosteroids, and gastroesophageal reflux disease. In present cohort, the BMI did not differ between the cases and controls which may be partly due to the relatively small study size.

#### **Conclusions**

The majority of the ER visits in adult asthma were directly due to asthma exacerbation or respiratory infections indicating the importance of the education of the patients. They should have the means to recognise the symptoms early on and adapt their self-management plan accordingly. Smoking cessation plays the key role in the maintenance of normal pulmonary function and prevention of emergency care.

#### Competing interests

The authors declare that they have no competing interests.

#### Authors' contributions

PK participated in the design and analysis of the study, and drafting and revising of the manuscript. HK participated in the analyses of the study and revising of the manuscript. AK participated in the acquisition of data and revising of the manuscript. TH participated in the design of the study and revising of the manuscript. TL participated in the acquisition of data, design of the study, and revising of the manuscript. All the authors have approved the final version to be published.

#### Acknowledgements

The authors thank clinical research nurses Kerstin Ahlskog, Kirsi Elorinne, Päivi Laakso, Tuula Lahtinen, and Piritta Rantanen for patient recruitment and monitoring the project. This study was supported by Helsinki University Hospital (HUS EVO), University of Helsinki, the Foundation of the Finnish Anti-Tuberculosis Association, the Finnish Allergy Programme by the Ministry of Social Affairs and Health, Allergiasäätiö ry, Tekes, and the Yrjö Jahnsson Foundation.

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Received: 1 December 2013 Accepted: 3 July 2014 Published: 16 July 2014

#### References

- Bateman ED, Hurd SS, Barnes PJ, Bousquet J, Drazen JM, FitzGerald M, Gibson P, Ohta K, O'Byrne P, Pedersen SE, Pizzichini E, Sullivan SD, Wenzel SE, Zar HJ: Global Strategy for asthma management and prevention: GINA executive summary. Eur Respir J 2008, 31:143–178.
- Bousquet J, Clark TJ, Hurd S, Khaltaev N, Lenfant C, O'byrne P, Sheffer A: GINA guidelines on asthma and beyond. Allergy 2007, 62:102–112.
- Haahtela T, Tuomisto LE, Pietinalho A, Klaukka T, Erhola M, Kaila M, Nieminen MM, Kontula E, Laitinen LA: A 10 years asthma programme in Finland: a major change for the better. Thorax 2006, 61:663–670.

- Latvala J, von Hertzen L, Lindholm H, Haahtela T: Trends in prevalence of asthma and allergy in Finnish young men: nationwide study, 1966–2003. BMJ 2005, 330:1186–1187.
- Hanania NA, King MJ, Braman SS, Saltoun C, Wise RA, Enright P, Falsey AR, Mathur SK, Ramsdell JW, Rogers L, Stempel DA, Lima JJ, Fish JE, Wilson SR, Boyd C, Patel KV, Irvin CG, Yawn BP, Halm EA, Wasserman SI, Sands MF, Ershler WB, Ledford DK, Asthma in Elderlly Workshop participants: Asthma in elderly: Current understanding and future research needs – a report of a National Institute on Aging (NIA) workshop. J Allergy Clin Immunol 2011. 128:54–524.
- Kauppi P, Linna M, Martikainen J, Mäkelä MJ, Haahtela T: Follow-up of the Finnish Asthma Programme 2000–2010: reduction of hospital burden needs risk group rethinking. Thorax 2013, 68:292–293.
- Enright PL, NcClelland RL, Newman AB, Gottlieb DJ, Lebowitz MD: Underdiagnosis and undertreatment of asthma in elderly. Chest 1999, 116:603–613.
- Gibson PG, McDonald VM, Marks GB: Asthma in older adults. Lancet 2010, 376:803–813.
- Cazzola M, Calzetta L, Bettoncelli G, Novelli L, Cricelli C, Rogliani P: Asthma and comorbid medical illness. Eur Respir J 2011, 38:42–49.
- 10. Lee HM, Truong ST, Wong ND: **Association of adult-onset asthma with specific cardiovascular conditions.** *Respir Med* 2012, **106**:948–952.
- Strine TW, Mokdad AH, Balluz LS, Berry JT, Gonzalez O: Impact of depression and anxiety on quality of life, health behaviors, and asthma control among adults in the United States. J Asthma 2008, 45:123–133.
- Hakola R, Kauppi P, Leino T, Ojajärvi A, Pentti T, Oksanen T, Haahtela T, Kivimäki M, Vahtera J: Persistent asthma, comorbid conditions and the risk of work disability: a prospective cohort study. Allergy 2011, 66:1598–1603.
- Kauppi P, Kupiainen H, Lindqvist A, Tammilehto L, Kilpeläinen M, Kinnula VL, Haahtela T, Laitinen T: Overlap syndrome of asthma and COPD predicts low quality of life. J Asthma 2011, 48:279–285.
- 14. Reddel HK, Taylor DR, Bateman ED, Boulet LP, Boushey HA, Busse WW, Casale TB, Chanez P, Enright PL, Gibson PG, de Jongste JC, Kerstjens HA, Lazarus SC, Levy ML, O'Byrne PM, Partridge MR, Pavord ID, Sears MR, Sterk PJ, Stoloff SW, Sullivan SD, Szefler SJ, Thomas MD, Wenzel SE, American Thoracic Society/European Respiratory Society Task Force on Asthma Control and Exacerbations: An official american thoracic society/european respiratory society statement: asthma control and exacerbations. Am J Respir Crit Care Med 2009, 180:59–99.
- Viljanen AA, Halttunen PK, Kreus KE, Viljanen BC: Spirometric studies in non-smoking, healthy adults. Scand J Clin Lab Invest Suppl 1982, 159:5–20.
- Miller MR, Hankinson J, Brusasco V, Burgos F, Casaburi R, Coates A, Crapo R, Enright P, van der Grinten CP, Gustafsson P, Jensen R, Johnson DC, MacIntyre N, McKay R, Navajas D, Pedersen OF, Pellegrino R, Viegi G, Wanger J, ATS/ERS Task Force: Standardisation of spirometry. Eur Respir J 2005, 26:319–338.
- Rabe KF, Hurd S, Anzueto A, Barnes PJ, Buist SA, Calverley P, Fukuchi Y, Jenkins C, Rodriguez-Roisin R, van Weel C, Zielinski J: Global strategy for the diagnosis, management, and prevention of chronic obstructive pulmonary disease: GOLD executive summary. Am J Respir Crit Care Med 2007. 176:532–555.
- Barley EA, Quirk FH, Jones PW: Asthma health status measurement in clinical practice: validity of a new short and simple instrument. Respir Med 1998, 92:1207–1214.
- Hajiro T, Nishimura K, Jones PW, Tsukino M, Ikeda A, Koyama H, Izumi T: A novel, short, and simple questionnaire to measure health-related quality of life in patients with chronic obstructive pulmonary disease. Am J Respir Crit Care Med 1999, 159:1874–1878.
- Oga T, Nishimura K, Tsukino M, Sato S, Hajiro T, Mishima M: Comparison of the responsiveness of different disease-specific health status measures in patients with asthma. Chest 2002, 122:1228–1233.
- Kauppinen R, Rissanen P, Sintonen H: Agreement between a generic and disease-specific quality-of-life instrument: the 15D and the SGRQ in asthmatic patients. Qual Life Res 2000, 9:997–1003.
- Urrutia I, Aguirre U, Pascual S, Esteban C, Ballaz A, Arrizubieta I, Larrea I: Impact of anxiety and depression on disease control and quality of life in asthma patients. J Asthma 2012, 49:201–208.
- Rowe BH, Voaklander DC, Wang D, Senthilselvan A, Klassen TP, Marrie TJ, Rosychuk RJ: Asthma presentations by adults to emergency departements in Alberta, Canada: a large populaton-based study. Chest 2009, 135:57–65.

- 24. Chalmers GW, Macleod KJ, Little SA, Thomson LJ, McSharry CP, Thomson NC: Influence of cigarette smoking on inhaled corticosteroid treatment in mild asthma. *Thorax* 2002, **573**:226–230.
- Tomlinson JEM, McMahon AD, Chaudhuri R, Thompson JM, Wood SF, Thomson NC: Efficacy of low and high dose inhaled corticosteroid in smokers versus non-smokers with mild asthma. *Thorax* 2005, 604:282–287.
- Lazarus SC, Chinchilli VM, Rollings NJ, Boushey HA, Cherniack R, Craig TJ,
  Deykin A, DiMango E, Fish JE, Ford JG, Israel E, Kiley J, Kraft M, Lemanske RF
  Jr, Leone FT, Martin RJ, Pesola GR, Peters SP, Sorkness CA, Szefler SJ,
  Wechsler ME, Fahy JV, National Heart Lung and Blood Institute's Asthma
  Clinical Research Network: Smoking affects response to inhaled
  corticosteroids or leukotriene receptor antagonists in asthma. Am J Respir
  Crit Care Med 2007, 1758:783–790.
- Osborne ML, Pedula KL, O'Hollaren M, Ettinger KM, Stibolt T, Buist AS, Vollmer WM: Assessing future need for acute care in adult asthmatics. Chest 2007, 132:1151–1161.
- Green RM, Custovic A, Sanderson G, Hunter J, Johnston SL, Woodcock A: Synergism between allergens and viruses and risk of hospital admission with asthma: a case–control study. BMJ 2002, 324:1131.
- 29. Noah TL, Zhou H, Jaspers I: Alterations of the nasal responses to influenza virus by tobacco smoke. Curr Opin Allergy Clin Immunol 2012, 12:24–31.
- Feldman C, Anderson R: Cigarette smoking and mechanisms of susceptibility to infections of the respiratory tract and other organ systems. J Infect 2013. 67:169–84.
- Bochkov YA, Busse WW, Brockman-Schneider RA, Evans MD, Jarjour NN, McCrae C, Miller-Larsson A, Gern JE: Budesonide and formoterol effects on rhinovirus replication and epithelial cell cytokine responses. Respir Res 2013. 14:98.
- Fernandes AK, Mallman F, Steinhorst AM, Noqueira FL, Avila EM, Saucedo DZ, Machado FJ, Raymundi MG, Barreto SS, Dalcin PT: Characteristics of acute asthma patients attended frequently with those attended only occasionally in an emergency department. J Asthma 2003, 40:683–690.
- Kullowatz A, Kanniess F, Dahme B, Magnussen H, Ritz T: Association of depression and anxiety with health care use and quality of life in asthma patients. Respir Med 2007, 101:638–644.
- Peterson MJ, Gaeta TJ, Birkhahn RH, Fernáandez JL, Mancuso CA: History of symptom triggers in patients presenting to the emergency department for asthma. J Asthma 2012, 49:629–636.
- Browatzki A, Ulrik CS, Lange P: Prevalence and severity of self-reported asthma in young adults. Eur Respir J 2009, 34:1046–1051.
- Mosen DM, Schatz M, Magid DJ, Camargo CA Jr: The relationship between obesity and asthma severity and control in adults. J Allergy Clin Immunol 2008, 122:507–511.

#### doi:10.1186/1471-2466-14-119

**Cite this article as:** Kauppi *et al.*: **Long-term smoking increases the need** for acute care among asthma patients: a case control study. *BMC Pulmonary Medicine* 2014 **14**:119.

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