



ORIGINAL ARTICLE

Inhaler technique knowledge and skills before and after an educational program in obstructive respiratory disease patients: A real-life pilot study



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Abstract

Introduction and objective: Patients present poor knowledge and skills about their respiratory disease and inhaler device. We aimed to: (1) evaluate COPD and asthmatic patients' ability to manage inhaled drugs (2) identify differences among devices and (3) correlate clinical data with patient ability.

Material and methods: Patients ($n = 134$) admitted for pulmonary rehabilitation (PR) were given an ad-hoc questionnaire covering 0% as the worst and 100% the best value of global ability (indicating the sum of knowledge and skills in managing inhaled drugs) at baseline (T0) and discharge (T1). Educational program was provided during PR. Setting of rehabilitation, age, sex, diagnosis, spirometry, CIRS score, level of autonomy to use medications, if naïve about PR, educational level, and number/type of prescribed inhaled drugs were recorded.

Results: Most patients used 1 drug while 37% used 2 drugs. DPIs were the main device prescribed. At baseline, patients' mean level of knowledge and skills were 73% and 58%, respectively. There was a significant difference in level of skills ($p = 0.046$) among device families, DPIs resulting worst and pMDIs best. Global ability, skills and knowledge improved after educational support ($p < 0.001$) but did not reach the optimal level, 88%, 87% and 89%, respectively. Baseline global ability was positively correlated to female gender, younger age, previous PR access, outpatient status, higher education level and GOLD D class.

Conclusions: At hospital admission, global ability was not optimal. Education may improve this, irrespective of the type of device used, in particular in male, elderly, naïve to PR, low educational level patients.

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Introduction

Asthma and chronic obstructive pulmonary disease (COPD) are chronic inflammatory pulmonary diseases affecting millions of people worldwide.¹ Inhaled therapy can be delivered via nebulizers, pressurized metered-dose inhalers (MDIs), dry powder inhalers (DPIs), and soft mist inhalers (SMIs). Studies consistently report that many patients with asthma and COPD do not use their inhaler devices correctly.^{2,3} Indeed, medications cannot be effective if they do not reach the sites they are intended to target.^{4,5} Poor inhaler technique stems from the fact that patients often poorly understand the purpose of and how to use their inhalation device.⁶ Poor adherence is common, with 50% or more of patients with asthma and COPD not taking their inhaled therapy as prescribed or instructed.^{7,8} Non adherence can further perpetuate poor technique and can lead to costly exacerbations and worsening disease.^{7,8} A comprehensive patient education, including device training, can improve outcomes.^{9,10} However, even with training, not all patients are able to use their inhalers correctly.¹¹

The aims of this study were: (1) to evaluate with a simple interview the global ability, knowledge and skills in managing inhaled drugs before and after an educational program during rehabilitation for obstructive respiratory disease patients; (2) to compare, if any, differences among the prescribed devices; and (3) to correlate clinical and anthropometric data with the overall ability to manage drugs.

Methods

This observational qualitative study considered a cohort of patients attending the Respiratory Unit of the Istituti Clinici Scientifici (ICS) Maugeri of Lumezzane (Bs), Italy. The study was conducted in a single center and approved by ICS Maugeri IRCCS Ethics Committee (EC 2322; 16 July 2019). All participants were informed and gave their written consent to participation.

Patients

Consecutive patients aged >18 years with a diagnosis of COPD or severe asthma were eligible for enrolment. COPD and asthma were defined according to Global Initiative for Chronic Obstructive Lung Disease (GOLD) criteria¹² and Global Initiative for Asthma (GINA) guidelines.¹³ Patients were admitted either to an inpatient program with an average stay of 25 (SD 3) days or an outpatient program attending no less than 22 rehabilitative sessions with 2 or 3 weekly accesses over a 2-month period. Indication and prescription for pulmonary rehabilitation (PR) was not based only on FEV₁ value, but according to a previously published internal triage,^{14,15} generating a Pulmonary Rehabilitation Decisional score (PRDS). PRDS considers several items such as lung function, clinical parameters, disability, frailty, and participation in ADLs and has been used for staging the clinical priority of PR prescription and choosing the PR setting (inpatient or outpatient). Anyway, all patients coming from acute hospitals have been inserted in the “inpatient group”. Patients were followed by a pulmonologist and

nurse case managers, nurses, respiratory therapists and others pulmonologists. The proposed PR programs were based on classical guidelines indications¹⁶ based on aerobic training, calisthenics, lower and/or upper limb selective muscle strengthening, balance training, inspiratory muscle training and secretion assistance when indicated. During the PR program drugs adjustments (in terms of quality and quantity) were carried out according to guidelines and clinical needs.

Exclusion criteria were: dyspnea at rest with need for acute hospitalization, oncological disease, terminal illness, neuromuscular degenerative diseases, severe orthopedic diseases, subject bedridden or confined to a wheelchair, and altered cognitive status measured by MMSE¹⁷ < 22.

Intervention

Development of the interview

To obtain a questionnaire with face validity based on expert opinion involving a structured process of consensus, we engaged key stakeholders (4 doctors and 4 nurses) from among health staff employed in the rehabilitation field of our Institute. We performed a systematic review of the COPD literature identifying items to use for the questionnaire and prepared a preliminary draft of questions. During the meeting, using a Delphi-like procedure, we asked the experts to rate the accordance of preselected items on a 5-point Likert scale (0 = totally disagree; 1 = disagree; 2 = sufficiently agree; 3 = moderately agree; 4 = totally agree). Consensus was considered when more than 75% of the respondents rated each item as totally agree. Finally, the focus group checked that the wording of each question was simple, clear, and comprehensible (details in supplementary material).

The final tool consisted of 8 questions enquiring about: the name of the drug/drugs, dosage prescribed, time of administration during the day, ability to distinguish the drug/s from others, the usefulness, and how to prepare, use and replace the drugs (Fig. 1). The operator scored each item dichotomously, according to the patient’s response, as knows/does not know or correct maneuver/incorrect maneuver (Fig. 1). The score for part A (representing the percentage of correct answers for knowledge) and that for part B (representing the percentage of correct maneuvers for skills) were added up to give a final total global ability score (A+B), 0% was the worst and 100% the best value of global ability (details in supplementary material).

Interview

Before conducting the interviews, all nurse staff participated in a briefing session on how to conduct the interview in a standardized way. According to the previous literature¹⁸ a list of possible mistakes/error for incorrect drug use was prepared. Nursing staff used the list during his/her interview (details in supplementary material).

The score was calculated (T0) during a face to face visit and assigned to a nurse case-manager not involved in the educational program. Patients admitted for a PR program, either as inpatients or outpatients, underwent the

defined as autonomous if they had an acceptable level of cognitive status and absence of dysphagia.

Statistical analysis

Statistical analysis was performed using STATA 11 (StataCorp LLC, Texas 77845-4512 USA). Continuous variables were expressed as mean \pm standard deviation (SD). Binary and categorical outcomes were described as frequency and percentage in each group. A two-sample t-test comparing differences at baseline between groups and changes after education training was performed. To identify correlations between baseline characteristics and the risk of presenting a low baseline global ability (defining by value below the median of global baseline data), we performed a post hoc analysis to estimate the Odds Ratio (OR). A p value <0.05 was considered as statistically significant.

Results

One hundred and thirty-four patients were consecutively enrolled in the study. The time spent on the patient interview was 3.06 ± 1.70 min (range 1.51–6.55).

Table 1 shows baseline data of the patients. The majority of patients routinely used only one inhaled drug while 37% used 2 drugs (the majority of these had COPD). Almost all patients were autonomous regarding general use of their medication. No differences were found in device categories between COPD and asthma. The majority of patients used a DPI: Diskus (9%), Ellipta (19.8%), Genuair (23%), Nexthaler (5.7%), Turbuhaler (5.1%), HandiHaler (7.4%), Breezhaler (30.6%). Younger patients used MDIs more frequently while patients with lower FEV₁% pred. more frequently used MDIs and SMIs. When compared to the whole group, SMI group (9.7%) included patients admitted in the 60% of the cases as inpatients, older (72 ± 9 years), in 38% of cases with CRF and with higher cultural level (70% of cases). At discharge after PR, drug prescription was changed in 30% of COPD and 51% asthmatic patients, respectively (in all cases number and dosage of drugs were increased).

Patients with low education level more frequently used MDIs. No differences in the devices used at admission were found concerning comorbidities and previous PR access.

All patients attended 2 educational sessions, 37.3% of COPD patients and 18.5% of asthmatic patients needed 3 educational sessions, because the dedicated nurses found these patients were still insecure and not yet ready to perform the final interview. Patients needing more than 2 sessions were older, with lower educational level, with lower MMSE and more naïve to PR.

On hospital admission, knowledge, skills and global ability regarding inhaled drugs were not optimal with 69%, 74% and 83% of patients with level $<100\%$, respectively. Mean and median global ability in percentage at baseline were 67.27 ± 26.26 and 68.75 (IQR 50.0–87.5), and at discharge 88.06 ± 18.82 and 100 (IQR 83.3–100) ($p < 0.001$), with mean delta improvement 20.79 ± 18.50 . Fig. 2 (panel a) shows baseline (T0) and delta improvement at discharge (T1) in percentage of the global ability score and according to the components of skills and knowledge. The pre-to-post changes were all statistically significant ($p < 0.001$)

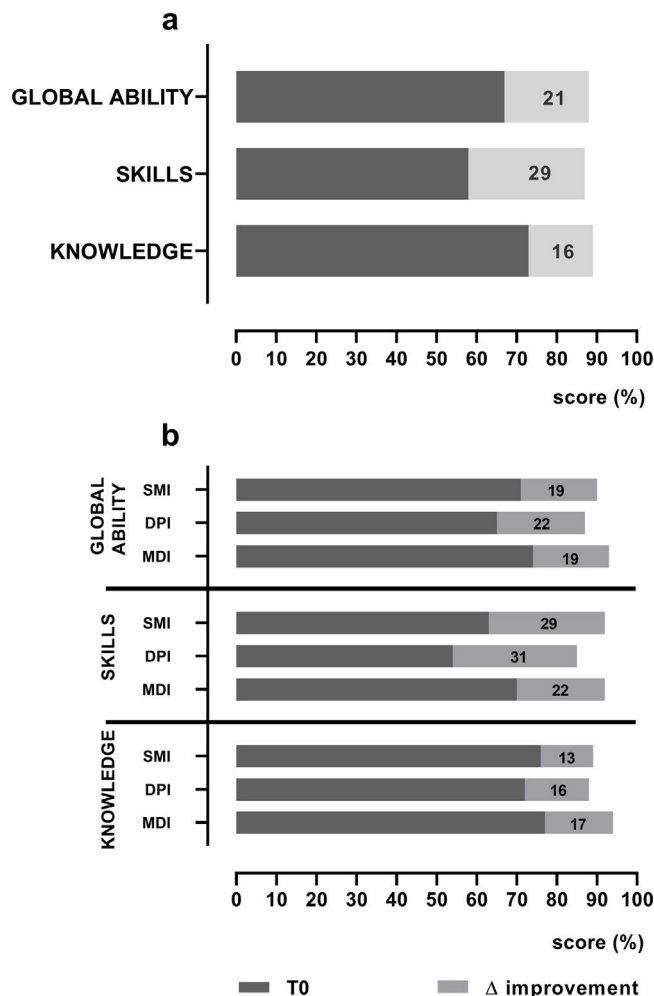


Figure 2 (panel a) shows baseline mean T0 score (dark bars) and mean delta score improvement at discharge (gray bars) in patient's global ability and the components of skills and knowledge. The pre-to-post changes (delta score improvement) were all statistically significant ($p < 0.001$); (panel b) shows baseline mean T0 scores (dark bars) and mean delta score improvement at discharge (gray bars) in global ability, in skills and knowledge according to different families of devices used. MDI = metered dose inhaler; DPI = dry powder inhaler; SMI = soft mist inhaler.

(panel a), the mean improvement in skills was 29.19 ± 29.61 and in knowledge 15.60 ± 20.17 . In detail, 100 patients (75%) improved global ability, while 34 (25%) worsened or remained unchanged. As to skills ability, at discharge, 30% of patients were still unable to use their devices correctly.

Fig. 2 (panel b) shows baseline values and delta improvement at discharge in global ability, skills and knowledge according to the 3 different families of devices tested (SMI, DPI and MDI). Differences in the patients' global ability at baseline were not significant among the devices tested ($p = 0.144$). At baseline only, there was a significant difference ($p = 0.046$) in the patients' level of skills in using different families of devices: the DPI family being the worst and the MDI the best; the delta skills improvement after education was otherwise not different ($p = 0.262$). Patients' knowledge of devices at baseline and delta knowledge improvement after education were not significantly dif-

Table 1 Baseline characteristics of patients.

	ALL	COPD	Asthma	<i>p</i>
<i>Patients, n</i>	134	107	27	
<i>Inpatient/outpatient, n</i>	49/85	42/65	7/20	0.199
<i>Age, years</i>	70.34 (8.26)	70.18 (8.53)	71 (7.20)	0.646
<i>Gender (M/F), n</i>	92/42	75/32	17/10	0.475
<i>FEV₁% pred.</i>	61.47 (25.27)	55.12 (23.01)	86.63 (16.90)	<0.001
<i>FVC % pred.</i>	84.51 (23.14)	82.05 (23.14)	94.30 (20.74)	0.013
<i>FEV₁/FVC</i>	59.22 (19.52)	52.63 (14.74)	85.37 (13.19)	<0.001
GOLD A, %		9.3		
GOLD B, %		38.3		
GOLD C, %		8.4		
GOLD D, %		43.9		
GINA I, %			7.4	
GINA II, %			14.8	
GINA III, %			48.1	
GINA IV, %			22.2	
GINA V, %			7.5	
<i>Educational level (elementary), n (%)</i>	71 (53)	52 (49)	19 (70)	0.103
<i>CIRS1, score</i>	1.70 (0.36)	1.70 (0.37)	1.69 (0.36)	0.897
<i>MMSE, score</i>	25.29 (1.24)	25.02 (1.37)	26.33 (1.01)	0.631
<i>Naive to pulmonary rehabilitation, n (%)</i>	48 (36)	35 (33)	13 (48)	0.135
<i>Respiratory drugs routinely used</i>	1.40 (0.52)	1.48 (0.54)	1.11 (0.32)	0.001
One, <i>n (%)</i>	82 (61)	58 (54)	24 (89)	0.004
Two, <i>n (%)</i>	50 (37)	47 (44)	3 (11)	0.004
Three, <i>n (%)</i>	2 (2)	2 (2)	0 (0)	0.004
<i>Autonomy in drug use, n (%)</i>	128 (96)	105 (98)	23 (85)	0.004
MDI, %	20.15	19.63	22.22	0.764
DPI, %	70.15	72.90	59.26	0.166
SMI, %	9.70	18.52	7.48	0.083

FEV₁ = forced expiratory volume at first second; FVC = forced volume capacity; GOLD = Global Initiative for Chronic Obstructive Lung Disease; GINA = Global Initiative for Asthma; CIRS = Cumulative Illness Rating Scale; MMSE = mini-mental state examination; MDI = metered dose inhaler; DPI = dry powder; SMI = soft mist inhaler. Where undefined, number in parentheses refers to (SD).

ferent ($p=0.516$ and $p=0.745$, respectively). At discharge global ability, skills and knowledge did not reach the optimal level being 88%, 87% and 89% respectively.

Table 2 shows the Odds Ratios risk to predict a low baseline global ability to use inhaler devices (less than the median value of 68.75). While female gender, younger age, previous PR access, outpatient status, higher education level and being into GOLD D class showed a protective effect on low baseline global ability to use inhaler devices, diagnosis, prior clinical classification of autonomy, FEV₁% < 30, CIRS > 1.6, type of device, and use of more than one inhaled drug did not influence baseline global ability.

Discussion

This study has demonstrated that at hospital admission knowledge and skills regarding inhaled drugs were not optimal (69% and 74% of patients with level <100%, respectively). A simple, clear and short interview at admission and discharge may be a good tool to monitor patients' knowledge and skills. A tailored educational program may strongly improve this gap, irrespective of the type of device used. Based on our findings, drugs education should be targeted

particularly at patients who are male, elderly, naïve to PR, with a low educational level.

Obstructive respiratory disease patients are poorly adherent to inhaler therapy. Low adherence was associated with age, current smoking status, number of respiratory drugs prescribed and poorer quality of life,²⁰ patients' perception of their health and medication effectiveness, and the presence of depressed mood and comorbidities.²¹

Eighty-two percent of COPD patients claimed to understand their disease and treatment, needed continuous education regarding inhalation devices.²² The Chronic Obstructive Pulmonary Disease knowledge Questionnaire (COPD-Q)²³ explores risk of infections, use of oxygen, utility of drugs, prevention, correct time to use long-acting drugs, symptoms, smoking cessation, SABA use, and disease time course. In our study, we proposed a short simple interview oriented exclusively on drugs knowledge and skills. The rationale of the questionnaire is to focus deeply on whether patients are able to remember the name, dosage, time of use, utility and can recognize their device amongst several other devices and perform correctly the preparation, use and post-use recommendations.

Table 2 Odds ratio (OR) to predict a low baseline global ability (less than the median value of 68.75).

	OR	Standard error	p	95% CI
Male	2.38	0.92	0.034	1.111–5.098
Age > 65 years	2.35	0.954	0.037	1.064–5.212
Not naïve to PR	0.11	0.04	<0.001	0.049–0.262
Outpatient	0.14	0.06	<0.001	0.064–0.335
Higher educational level	0.24	0.89	0.026	0.118–0.501
GOLD D class	0.33	0.13	0.004	0.156–0.708

PR = pulmonary rehabilitation, CI = confidence interval.

In outpatient settings mistakes using inhaler devices has been found in a range of 6–71%, with 40% of patients presenting at least one vital mistake.^{24–28} Muller et al.²⁹ found that 51.8% of patients present at least one mistake about using their devices. Molimard et al.,⁸ in a large cohort of patients outside of hospital, assessed inhaler device handling and its association with exacerbations. Handling errors were found in over 50% and exacerbations were less frequent in the absence of errors, while they doubled in the presence of critical error. Melani et al.³⁰ in a large cohort of both asthmatic and COPD patients found that critical mistakes were no fewer in DPIs than in MDIs: in 12% for MDIs vs. 35% for Diskus and HandiHaler, and 44% for Turuhaler. In patients referred for a chest visit, inhaler-specific errors were as follows: Aerolizer 9.1%, Diskus 26.7%, HandiHaler 53.1% and Turbuhaler 34.9%.³¹

Only one study was conducted in an inpatient setting—it found that misuse was common in COPD (86% with MDI and 71% with Diskus).³² The lower level of misuse and errors found in our sample (60% for drugs skills) may be due to the particular rehabilitative setting, i.e. 64% of patients were not naïve to PR, probably already confident with regard to drug education, with mixed diseases (asthma and COPD) and attending PR as both out- and inpatients. DPIs derive the energy for the emptying of the drug system from the user inspiratory flow: the failure to achieve a forceful inspiratory flow through a device was the most common critical mishandling error with DPIs in Melani et al.'s study.³⁰ In fact, in our study, COPD patients with a DPI were less able than patients with an MDI or SMI to use their devices, probably due to their severe functional limitation.

Critical errors and inability to improve after an education program found the following factors: older age,^{30,31,33} lower schooling^{30,33} and lack of instruction received,^{30,31,33–35} cognitive impairment or dyspraxia,³⁶ use of 2 or more inhalers,^{34,36} and severity of airway obstruction.³¹ Our study confirms that age, lack of a previous educational program, and low level of education are negative predictors for drugs use ability; as a novel finding, we also demonstrated that female gender and outpatient regime (as opposed to inpatient) are protective factors. It is noteworthy that female patients seem to do better: possible explanations are that this group presented a higher cultural level (in 57% of cases) and fewer naïve to PR (14%) when compared to males (43% and 40%, respectively). At the same time, it is not unexpected that outpatients group was better than inpatients due to a lower age (70 ± 8 years old), a better level of airway obstruction ($67 \pm 24\%$ pred), more frequently with asthma (23%) and with a very low number of patients unable to use

drugs (2.3%). Finally, it is interesting that only the more severe patients belonging GOLD D class presented a lower risk of having global inability: the possible explanation is that this group had attended previous PR program and education in the 76% of cases and, being more symptomatic, used more drugs with a higher adherence.

The lack of education by health caregivers on inhaler technique significantly increases the risk of misuse for all the studied devices.^{30,37} One study assessed errors with different devices in asthmatic and COPD patients after they had read the patient information leaflet:²⁴ fewer COPD patients made critical errors with Ellipta (5%) than with other devices, and most patients (57–70%) did not require health-operator instructions using Ellipta, but instructions were required for Diskus (65%), MDI (85%), Turbuhaler (71%), HandiHaler (62%) and Breezhaler (56%).

Different training programs of diverse intensity and content (individual interviews about beliefs and behavior related to adherence, information about the illness and training about inhalation techniques, video) have been proposed, with patient skills' improvement ranging from 20 to 79%.^{26,35,38–40} In our sample, the mean improvement in skills was 29.19% while that in knowledge was 15.60%: these delta differences are clearly related to lower baseline values allowing more space for improvement (Fig. 2).

Limitations and strengths

Limitations of the study are the small sample size, the subjective basis of our findings (based on researcher-s judgment), the lack of a comparison control group and the fact that knowledge and skills measurement were not related to clinical outcomes. Another limitation is the use of a questionnaire with a dichotomous yes/no answer where the different skill capacity steps depended on a single answer grouped together. Strengths of our study are: (1) the simplicity of a questionnaire exclusively dedicated to drugs knowledge and skills; (2) the relatively short time needed to administer it, favoring its potential for routine use; (3) the comparison among different devices available in the market and (4) the presented data on a selected population admitted to a rehabilitative center.

Conclusions

The findings were predominantly confirmatory of previous literature covering patients' lack of skills and knowledge about inhaled drugs, anthropometric, clinical and functional

features which influence patients' global ability. The study adds that an educational program may improve this gap, irrespective of the type of device used, in patients who are male, elderly, naïve to PR, with a low educational level.

Author contributions

MV, MF, EM, LC designed the study and draft the manuscript.

MF, EM, LC performed educational assessment and reviewed the manuscript.

MP and LB performed statistical analysis and reviewed the manuscript.

GB, CZ, AP, DF performed clinical assessment and reviewed the manuscript.

All the Authors approved the definitive version of the manuscript and declare that questions related to the accuracy or integrity of any part of it have been appropriately investigated and resolved.

Conflict of interest

MV declares conflict for consultancy for or receipt of speaker's fees from Boehringer; all the other Authors have no conflict of interest to disclose.

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Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at [doi:10.1016/j.pulmoe.2020.04.010](https://doi.org/10.1016/j.pulmoe.2020.04.010).

References

1. Global, regional, and national age-sex specific mortality for 264 causes of death, 1980–2016: a systematic analysis for the Global Burden of Disease Study 2016. *Lancet*. 2017;390:1151–210.
2. Chrystyn H, van der Palen J, Sharma R, Barnes N, Delafont B, Mahajan A, et al. Device errors in asthma and COPD: systematic literature review and meta-analysis. *NPJ Prim Care Respir Med*. 2017;27:22.
3. Price DB, Roman-Rodriguez M, McQueen RB, Bosnic-Anticevich S, Carter V, Gruffydd-Jones K, et al. Inhaler errors in the CRITIKAL study: type, frequency, and association with asthma outcomes. *J Allergy Clin Immunol Pract*. 2017;5:1071–81, e9.
4. Levy ML, Hardwell A, McKnight E, Holmes J. Asthma patients' inability to use a pressurised metered-dose inhaler (pMDI) correctly correlates with poor asthma control as defined by the global initiative for asthma (GINA) strategy: a retrospective analysis. *Prim Care Respir J*. 2013;22:406–11.
5. Lavorini F, Magnan A, Dubus JC, Voshaar T, Corbetta L, Broeders M, et al. Effect of incorrect use of dry powder inhalers on management of patients with asthma and COPD. *Respir Med*. 2008;102:593–604.
6. Darba J, Ramirez G, Sicras A, Garcia-Bujalance L, Torvinen S, Sanchez-de la Rosa R. Identification of factors involved in medication compliance: incorrect inhaler technique of asthma treatment leads to poor compliance. *Patient Prefer Adher*. 2016;10:135–45.
7. van Boven JF, Chavannes NH, van der Molen T, Rutten-van Molken MP, Postma MJ, Vegter S. Clinical and economic impact of non-adherence in COPD: a systematic review. *Respir Med*. 2014;108:103–13.
8. Molimard M, Raheison C, Lignot S, Balestra A, Lamarque S, Chartier A, et al. Chronic obstructive pulmonary disease exacerbation and inhaler device handling: real-life assessment of 2935 patients. *Eur Respir J*. 2017;49(2), pii: 1601794.
9. Wilson SR, Strub P, Buist AS, Knowles SB, Lavori PW, Lapidus J, et al. Shared treatment decision making improves adherence and outcomes in poorly controlled asthma. *Am J Respir Crit Care Med*. 2010;181:566–77.
10. Price D, Bosnic-Anticevich S, Briggs A, Chrystyn H, Rand C, Scheuch G, et al. Inhaler competence in asthma: common errors, barriers to use and recommended solutions. *Respir Med*. 2013;107:37–46.
11. Haughney J, Price D, Barnes NC, Virchow JC, Roche N, Chrystyn H. Choosing inhaler devices for people with asthma: current knowledge and outstanding research needs. *Respir Med*. 2010;104:1237–45.
12. Vogelmeier CF, Criner GJ, Martinez FJ, Anzueto A, Barnes PJ, Bourbeau J, et al. Global strategy for the diagnosis, management, and prevention of chronic obstructive lung disease 2017 report. GOLD Executive Summary. *Am J Respir Crit Care Med*. 2017;195:557–82.
13. Global Initiative for Asthma. Global strategy for asthma management and prevention. Available at: www.ginasthma.org; 2017 [updated 16.08.19].
14. Vitacca M, Comini L, Barbisoni M, Francolini G, Paneroni M, Ramponi JP. A pulmonary rehabilitation decisional score to define priority access for COPD patients. *Rehabil Res Pract*. 2017;2017:5710676, <http://dx.doi.org/10.1155/2017/5710676>.
15. Olivares A, Vitacca M, Comini L. Combining the pulmonary rehabilitation decisional score with the bode index and clinical opinion in assigning priority for pulmonary rehabilitation. *COPD*. 2018;15:238–44.
16. Spruit MA, Singh SJ, Garvey C, ZuWallack R, Nici L, Rochester C, et al. An official American Thoracic Society/European Respiratory Society statement: key concepts and advances in pulmonary rehabilitation. *Am J Respir Crit Care Med*. 2013;188:e13–64. Erratum in: *Am J Respir Crit Care Med*. 2014;189:1570.
17. Folstein MF, Folstein SE, McHugh PR. "Mini-mental state". A practical method for grading the cognitive state of patients for the clinician. *J Psychiatr Res*. 1975;12:189–98.
18. van der Palen J, Klein JJ, van Herwaarden CL, Zielhuis GA, Seydel ER. Multiple inhalers confuse asthma patients. *Eur Respir J*. 1999;14:1034–7.
19. Linn BS, Linn MW, Gurel L. Cumulative illness rating scale. *J Am Geriatr Soc*. 1968;16:622–6.
20. Agh T, Inotai A, Meszaros A. Factors associated with medication adherence in patients with chronic obstructive pulmonary disease. *Respiration*. 2011;82:328–34.
21. Khdour MR, Hawwa AF, Kidney JC, Smyth BM, McElnay JC. Potential risk factors for medication non-adherence in patients with chronic obstructive pulmonary disease (COPD). *Eur J Clin Pharmacol*. 2012;68:1365–73.
22. Dhand R, Mahler DA, Carlin BW, Hanania NA, Ohar JA, Pinto-Plata V, et al. Results of a patient survey regarding COPD knowledge. Treatment experiences, and practices with inhalation devices. *Respir Care*. 2018;63:833–9.
23. Maples P, Franks A, Ray S, Stevens AB, Wallace LS. Development and validation of a low-literacy Chronic Obstructive

- Pulmonary Disease knowledge Questionnaire (COPD-Q). *Patient Educ Couns*. 2010;81:19–22.
24. Rootmensen GN, van Keimpema AR, Jansen HM, de Haan RJ. Predictors of incorrect inhalation technique in patients with asthma or COPD: a study using a validated videotaped scoring method. *J Aerosol Med Pulm Drug Deliv*. 2010;23:323–8.
 25. Giraud V, Roche N. Misuse of corticosteroid metered-dose inhaler is associated with decreased asthma stability. *Eur Respir J*. 2002;19:246–51.
 26. Press VG, Arora VM, Shah LM, Lewis SL, Charbeneau J, Naureckas ET, et al. Teaching the use of respiratory inhalers to hospitalized patients with asthma or COPD: a randomized trial. *J Gen Intern Med*. 2012;27:1317–25.
 27. Quinet P, Young CA, Heritier F. The use of dry powder inhaler devices by elderly patients suffering from chronic obstructive pulmonary disease. *Ann Phys Rehabil Med*. 2010;53:69–76.
 28. Bryant L, Bang C, Chew C, Baik SH, Wiseman D. Adequacy of inhaler technique used by people with asthma or chronic obstructive pulmonary disease. *J Prim Health Care*. 2013;5:191–8.
 29. Muller T, Baas H, Kassubek J, Riederer P, Urban PP, Schrader C, et al. Optimizing inhalation technique using web-based videos in obstructive lung diseases. *Respir Med*. 2017;129:140–4.
 30. Melani AS, Bonavia M, Cilenti V, Cinti C, Lodi M, Martucci P, et al. Inhaler mishandling remains common in real life and is associated with reduced disease control. *Respir Med*. 2011;105:930–8.
 31. Press VG, Arora VM, Shah LM, Lewis SL, Ivy K, Charbeneau J, et al. Misuse of respiratory inhalers in hospitalized patients with asthma or COPD. *J Gen Intern Med*. 2011;26:635–42.
 32. Paneroni M, Clini E, Crisafulli E, Guffanti E, Fumagalli A, Bernasconi A, et al. Feasibility and effectiveness of an educational program in Italian COPD patients undergoing rehabilitation. *Respir Care*. 2013;58:327–33.
 33. Allen SC, Ragab S. Ability to learn inhaler technique in relation to cognitive scores and tests of praxis in old age. *Postgrad Med J*. 2002;78:37–9.
 34. Wieshammer S, Dreyhaupt J. Dry powder inhalers: which factors determine the frequency of handling errors? *Respiration*. 2008;75:18–25.
 35. Leiva-Fernandez J, Leiva-Fernandez F, Garcia-Ruiz A, Prados-Torres D, Barnestein-Fonseca P. Efficacy of a multifactorial intervention on therapeutic adherence in patients with chronic obstructive pulmonary disease (COPD): a randomized controlled trial. *BMC Pulm Med*. 2014;14:70.
 36. van der Palen J, Thomas M, Chrystyn H, Sharma RK, van der Valk PD, Goosens M, et al. A randomised open-label cross-over study of inhaler errors, preference and time to achieve correct inhaler use in patients with COPD or asthma: comparison of ELLIPTA with other inhaler devices. *NPJ Prim Care Respir Med*. 2016;26:16079.
 37. Chrystyn H, Small M, Milligan G, Higgins V, Gil EG, Estruch J. Impact of patients' satisfaction with their inhalers on treatment compliance and health status in COPD. *Respir Med*. 2014;108:358–65.
 38. Muller T, Baas H, Kassubek J, Riederer P, Urban PP, Schrader C, et al. Laboratory assessments in the course of Parkinson's disease: a clinician's perspective. *J Neural Transm (Vienna)*. 2016;123:65–71.
 39. Giraud V, Allaert FA, Roche N. Inhaler technique and asthma: feasibility and acceptability of training by pharmacists. *Respir Med*. 2011;105:1815–22.
 40. Sestini P, Cappiello V, Aliani M, Martucci P, Sena A, Vaghi A, et al. Prescription bias and factors associated with improper use of inhalers. *J Aerosol Med*. 2006;19:127–36.