

## Editorial

## COPD: Challenges in Primary Care

Ioanna Tsiligianni\*

School of Medicine, University of Crete, Greece

\*Corresponding author: Ioanna Tsiligianni, Asites rural practice, Agia Barbara Health Care Centre, Heraklion, Crete, Greece Cretan Practice-based Primary Care Research Network 'G. Lambrakis' & Clinic of Social and Family Medicine, School of Medicine, University of Crete, Heraklion, Crete, Greece

Received: July 11, 2015; Accepted: July 27, 2015;

Published: July 30, 2015

## Introduction

Chronic obstructive pulmonary disease is a progressive disease characterized by persistent airflow limitation [1] that it is estimated to be the third leading cause of death worldwide [2]. The challenges for Primary Care (PC) are known to be immense and among them include, primary prevention with reduction of risk factors, secondary prevention with early detection and monitoring of COPD and tertiary prevention to improve health status, decrease disease progression and reduce exacerbations rate [1,3].

## Prevention

In PC, general preventive strategies are influenced by time, cost, availability and practice capacity that makes it a challenge to manage behavioral risk factors [4]. Undoubtedly, cigarette smoking is by far the most important risk factor for COPD [1]. Fletcher and Peto demonstrated that smoking cessation abated the lung function decline that is characteristic of COPD [5]. Effective smoking cessation interventions in primary care are based on an awareness of which strategies have been shown to work and on making the most of available resources [6,7]. To that direction the International Primary Care Respiratory Group (IPCRG) has published a practical guidance for PC [6]. Despite that it is well known that smoking cessation in PC is feasible [6,7], physicians are not always prepared to deliver effective interventions for smoking cessation to their patients [8]. Furthermore, education and awareness to patients that pharmacological treatment as well as behavioural support for smoking cessation is available at a PC clinic is needed [9]. However, the preventive role of PC regarding COPD should not be limited to smoking. The Copenhagen City Heart Study highlighted that moderate to high levels of regular physical activity are associated with reduced lung function decline and COPD risk among smokers [10]. Additionally, a systematic review by the PRO active consortium showed that the physical activity level in COPD is consistently associated with mortality and exacerbations, yet there is poor evidence about determinants of physical activity [11]. Therefore, it is obvious that health-care providers should promote physical activity. Moreover data indicate that an increase in consumption of vegetables, fruit and (although with less evidence) fibre may contribute to the prevention of COPD [12].

When designing prevention strategies for COPD, a variety of

risk factors should be considered; host factors (i.e; atopy), perinatal factors (i.e; maternal smoking, maternal exposure to air pollution), childhood exposures (i.e; respiratory tract infections, indoor-outdoor air pollution, obesity) and adult exposures (i.e; occupational, indoor-outdoor air pollution) [13]. Therefore efforts to prevent COPD should also focus on optimization of lung health in the early years of life, before birth, and possibly even before conception [13]. Provider education, raising community awareness and the support and capacity building may improve the uptake of lifestyle modification interventions [4].

## Challenging Mis and Under Diagnosis- The Role of Spirometry

The PLATINO [14] and the IBEPROC [15] study showed that 88.7% and 78.2% respectively, of patients with COPD had not been diagnosed while in the USA, more than 50% of patients with COPD have been either undiagnosed or misdiagnosed [16]. Patients with COPD may experience respiratory symptoms that are often disregarded by patients who do not always report them to their PC physicians [17].

Misdiagnosis of COPD and asthma is common [18]. Tinkelman et al. have developed a symptom-based questionnaire to differentiate between asthma and COPD that could potentially be used extensively in PC in the future [19]. Spirometry is an important tool for the

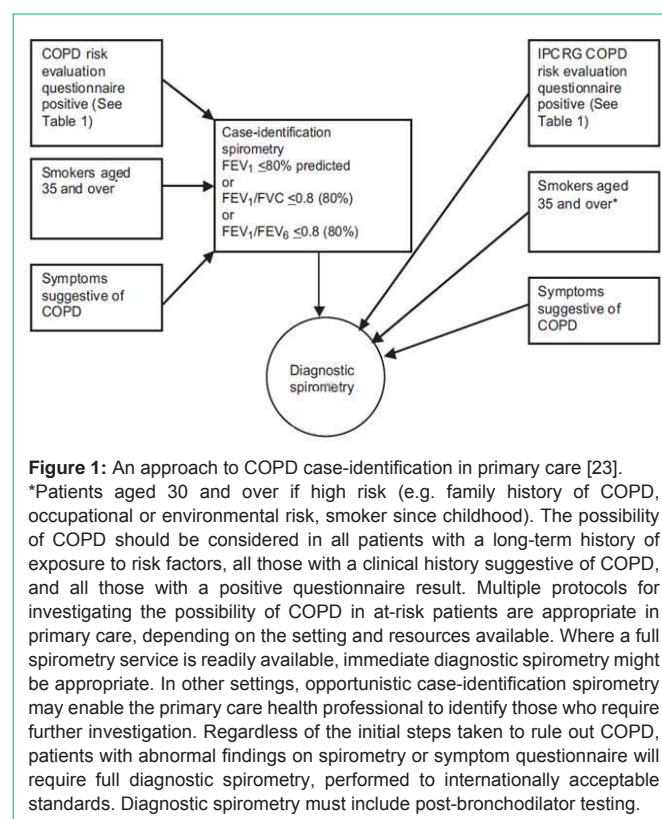
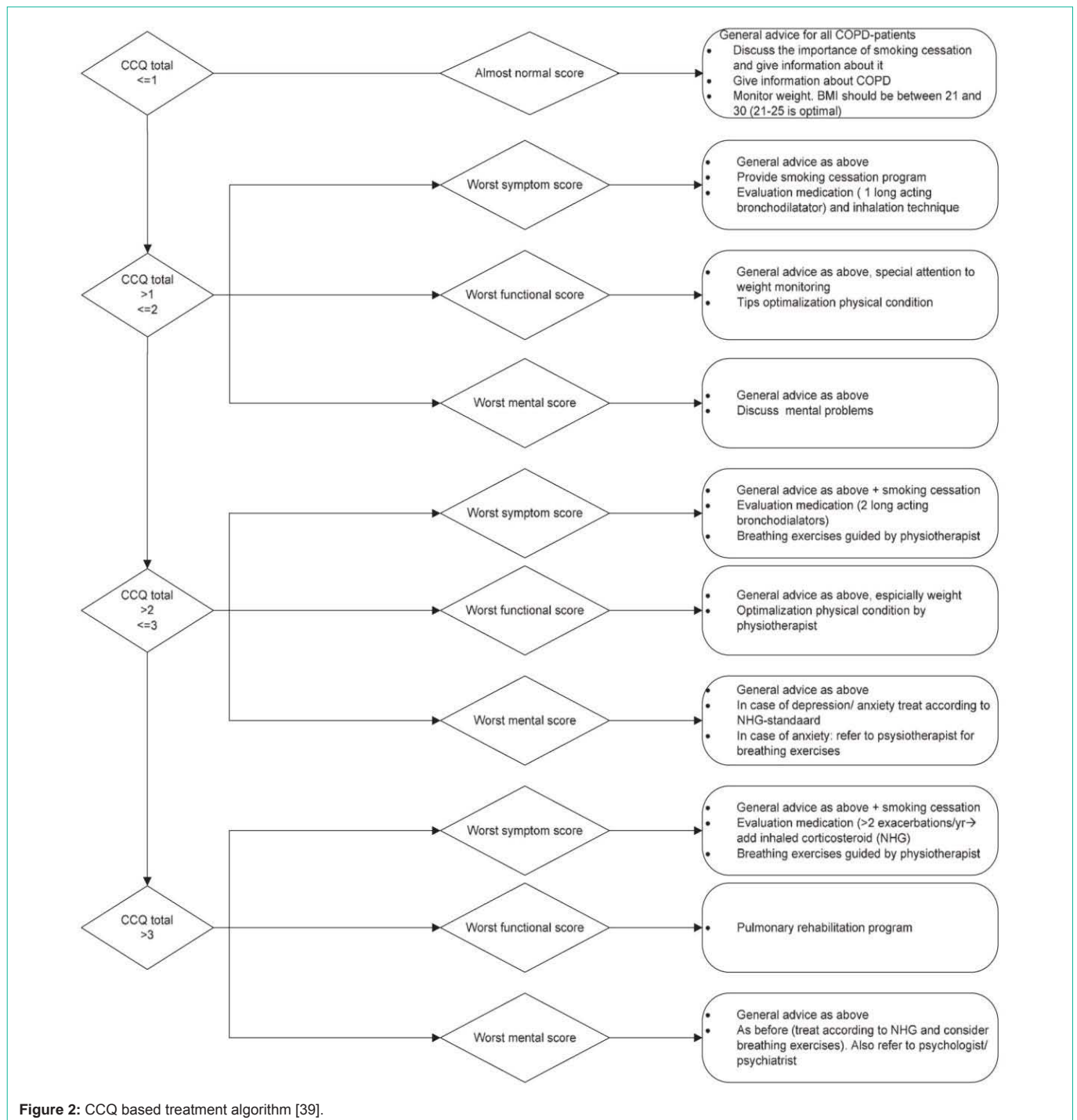


Figure 1: An approach to COPD case-identification in primary care [23].

\*Patients aged 30 and over if high risk (e.g. family history of COPD, occupational or environmental risk, smoker since childhood). The possibility of COPD should be considered in all patients with a long-term history of exposure to risk factors, all those with a clinical history suggestive of COPD, and all those with a positive questionnaire result. Multiple protocols for investigating the possibility of COPD in at-risk patients are appropriate in primary care, depending on the setting and resources available. Where a full spirometry service is readily available, immediate diagnostic spirometry might be appropriate. In other settings, opportunistic case-identification spirometry may enable the primary care health professional to identify those who require further investigation. Regardless of the initial steps taken to rule out COPD, patients with abnormal findings on spirometry or symptom questionnaire will require full diagnostic spirometry, performed to internationally acceptable standards. Diagnostic spirometry must include post-bronchodilator testing.



accurate diagnosis of COPD [1,3] and lack of routine spirometry [20] is a key cause of this misdiagnosis. The DIDASCO study illustrated a high accuracy with the use of a portable spirometer when PC physicians received training in performance and interpretation [21]. Despite that, spirometry is still underused in PC [22,23] due to barriers such as, lack of access to calibrated spirometers and inadequate training and interpretation skills [22-24]. However it is promising to assure the quality of spirometry in PC settings [25] and solutions to overlap technical interpretation barriers have been extensively

proposed [23]. Some examples are [23] basic training, operators that practice performing the man oeuvre to reinforce and improve skills, automatic feedback and technical error messages from the machine, use of IPCRG opinion sheets for spirometry [26], financial incentives for PC physicians to offset the cost of training, etc [23]. Increased use of spirometry in PC is a significant challenge as it is expected not only to increase rate of detection of COPD but also reporting to smokers their lung age (the age of the average healthy individual who would perform similar to them on spirometry), which is found to be

effective to significantly improve the likelihood of someone to quit smoking [27].

### Mass Population Screening or Case Finding?

The use of spirometry in mass population screening remains controversial and is not recommended however, its use in targeted case identification is more feasible [1]. Although when case-finding using full diagnostic spirometry is not possible in the PC setting in patients with symptoms, there are other approaches that could be useful. Such approaches are the use of questionnaires [28-31] designed to assess the risk for COPD and/or micro-spirometers such as copd-6 [32] PiKo-6 [33,34] as initial screening tools. All these studies [32-34] on microspirometry demonstrated a high negative predictive value meaning that they predict with a range of 91-98% accuracy level that the patient does not have COPD. In this direction, IPCRG has developed an algorithm driving GPs towards an effective case-finding (Figure 1) [23]. More studies are needed to confirm their role in PC.

### COPD Management in Primary Care Daily Clinical Practice

Even if spirometry is performed and a diagnosis is established, COPD patients present with a large variety of symptoms and signs, and patients with a similar degree of airway obstruction report different performance and Quality of Life (QoL) scores [35,36]. In general, patients with COPD tend to underreport their symptoms and impairments thus leaving physician with the impression that the situation is better than it actually is [36]. Indeed, research on health-related QoL shows that physician's consistently assess their patients' QoL better than when measured by questionnaires [37]. Therefore GOLD suggests the COPD management tree to be based on A,B,C,D categories; a classification based on symptomatology (as assessed by CAT, or CCQ, or mMRC), spirometry and exacerbations [1]. However this classification has been criticized that 'these categories are too complex to be used in PC' and suggested that COPD severity assessment could be accomplished by using multi-component indices such as Body mass index, airflow Obstruction, Dyspnoea, and Exercise capacity (BODE), Age, Dyspnoea and airflow Obstruction (ADO), or the Dyspnoea, Obstruction, Smoking and the Exacerbation index (DOSE) [38]. PC has suggested management should follow an evidence-based algorithm involving smoking cessation, pulmonary rehabilitation, and drug treatment [38]. An example could be seen in figure 2 [39].

### Therapeutic Challenges in Primary Care

There is strong evidence on which medications to suggest according to the A,B,C,D, categories in GOLD guidelines mainly based on large pharmaceutically sponsored COPD studies (LPCS) [1]. However PC COPD patients stand out from patients enrolled in LPCS in terms of gender, lung function, quality of life and exacerbations [40]. More research is needed to determine the effect of pharmacological treatment in PC patients that often present a lot of co morbidities that are usually excluded from LPCS at the moment [40]. Therefore when developing future guidelines for COPD, it is encouraged to involve PC populations in their recommendations [40].

### Phenotypes and Co Morbidities

COPD is a complex disease with many phenotypes and co morbidities. For example there is strong evidence that there is a specific phenotype characterized by an increased susceptibility to COPD exacerbations [41] and another characterized by eosinophil-related airway inflammation [42], and both may benefit from inhaled corticosteroids therapy. To what extent does PC use these phenotypes when managing patient's remains a challenge that should be established in the future as well as how PC could use these phenotypes towards the patients' benefit.

Multimorbidity in patients with COPD is common and a great challenge for PC [43]. Anecchino and colleagues conducted a large COPD cohort of 126,838 individuals in Italy and reported that the vast majority of them (98%) had at least one prescription for 'non-respiratory drugs' [44]. A Canadian, population study using data of more than >7 million adults showed that patients with COPD had more health service claims for comorbid disease than for COPD itself [45]. This could be used as a motivation to PC physicians to search for COPD when they present with another chronic condition, especially if they share common risk factors. Moreover GPs should search for some diseases that are quite prevalent in COPD patients and are outside the classical known co morbidities (cardiovascular diseases and diabetes) especially depression, anxiety and osteoporosis [43]. It is obvious that COPD medications can have a beneficial or harmful effect on other disease outcomes [43]. On the other hand, co administration of medications for co morbidities can have beneficial, neutral or harmful effects on COPD outcomes [43]. However, even if GPs are aware of single disease guidelines it may be that they are reluctant to use medications when a patient has more than two co morbidities. For example although the benefits of b-blockers in COPD are well known in reducing the risk of exacerbations and improving survival [46] they are still underused in COPD even if there is a clear need to use them ie; in concomitant cardiovascular diseases [47,48]. The above mentioned are challenges not only for COPD but also for other chronic diseases and this highlights the need for multimorbid guidelines and not single disease-centred clinical guidelines [49].

### References

1. Global Initiative for Chronic Obstructive Lung Disease (GOLD): Global Strategy for the Diagnosis Management and prevention of COPD. 2015.
2. WHO. The top 10 causes of death. May, 2014.
3. Soriano JB, Zielinski J, Price D. Screening for and early detection of chronic obstructive pulmonary disease. *Lancet*. 2009; 374: 721-732.
4. Ampt AJ, Amoroso C, Harris MF, McKenzie SH, Rose VK, Taggart JR. Attitudes, norms and controls influencing lifestyle risk factor management in general practice. *BMC Fam Pract*. 2009; 10: 59.
5. Fletcher C, Peto R. The natural history of chronic airflow obstruction. *Br Med J*. 1977; 1: 1645-1648.
6. Van Schayck OC, Pinnock H, Ostrem A, Litt J, Tomlins R, Williams S, et al. IPCRG Consensus statement: tackling the smoking epidemic - practical guidance for primary care. *Prim Care Respir J*. 2008; 17: 185-193.
7. Litt J. How to provide effective smoking cessation advice in less than a minute without offending the patient. *Aust Fam Physician*. 2002; 31: 1087-1094.
8. Nobile CG, Bianco A, Biafore AD, Manuti B, Pileggi C, Pavia M. Are primary care physicians prepared to assist patients for smoking cessation? Results of a national Italian cross-sectional web survey. *Prev Med*. 2014; 66: 107-112.

9. Van Rossem C, Spigt MG, Kleijnsen JR, Hendrick M, Van Schayck CP, Kotz D. Smoking cessation in primary care: Exploration of barriers and solutions in current daily practice from the perspective of smokers and healthcare professionals. *Eur J Gen Pract.* 2015; 21: 111-117.
10. Garcia-Aymerich J, Lange P, Benet M, Schnohr P, Antó JM. Regular physical activity modifies smoking-related lung function decline and reduces risk of chronic obstructive pulmonary disease: a population-based cohort study. *Am J Respir Crit Care Med.* 2007; 175: 458-463.
11. Gimeno-Santos E, Frei A, Steurer-Stey C, de Battle J, Rabinovich RA, Raste Y, et al. Determinants and outcomes of physical activity in patients with COPD: a systematic review. *Thorax.* 2014; 69: 731-739.
12. Annesi-Maesano I, Roche N. Healthy behaviours and COPD. *Eur Respir Rev.* 2014; 23: 410-415.
13. Postma DS, Bush A, Van den Berge M. Risk factors and early origins of chronic obstructive pulmonary disease. *Lancet.* 2015; 385: 899-909.
14. Tálamo C, de Oca MM, Halbert R, Perez-Padilla R, Jardim JR, Muñio A, et al. Diagnostic labeling of COPD in five Latin American cities. *Chest.* 2007; 131: 60-67.
15. Peña VS, Miravittles M, Gabriel R, Jiménez-Ruiz CA, Villasante C, Masa JF, et al. Geographic variations in prevalence and underdiagnosis of COPD: results of the IBERPOC multicentre epidemiological study. *Chest.* 2000; 118: 981-989.
16. Wise RA, Tashkin DP. Preventing chronic obstructive pulmonary disease: what is known and what needs to be done to make a difference to the patient? *Am J Med.* 2007; 120: S14-22.
17. Van Weel C. Underdiagnosis of asthma and COPD: is the general practitioner to blame? *Monaldi Arch Chest Dis.* 2002; 57: 65-68.
18. Pearson M, Ayres JG, Sarno M, Massey D, Price D. Diagnosis of airway obstruction in primary care in the UK: the CADRE (COPD and Asthma Diagnostic/management REassessment) programme 1997-2001. *Int J Chron Obstruct Pulmon Dis.* 2006; 1: 435-443.
19. Tinkelman DG, Price DB, Nordyke RJ, Halbert RJ, Isonaka S, Nonikov D, et al. Symptom-based questionnaire for differentiating COPD and asthma. *Respiration.* 2006; 73: 296-305.
20. Tinkelman DG, Price DB, Nordyke RJ, Halbert RJ. Misdiagnosis of COPD and asthma in primary care patients 40 years of age and over. *J Asthma.* 2006; 43: 75-80.
21. Buffels J, Degryse J, Heyrman J, Decramer M. DIDASCO Study. Office spirometry significantly improves early detection of COPD in general practice: the DIDASCO Study. *Chest.* 2004; 125: 1394-1399.
22. Yawn BP, Wollan PC. Knowledge and attitudes of family physicians coming to COPD continuing medical education. *Int J Chron Obstruct Pulmon Dis.* 2008; 3: 311-317.
23. Price D, Crockett A, Arne M, Garbe B, Jones RC, Kaplan A, et al. Spirometry in primary care case-identification, diagnosis and management of COPD. *Prim Care Respir J.* 2009; 18: 216-223.
24. Bolton CE, Ionescu AA, Edwards PH, Faulkner TA, Edwards SM, Shale DJ. Attaining a correct diagnosis of COPD in general practice. *Respir Med.* 2005; 99: 493-500.
25. Levy ML, Quanjer PH, Booker R, Cooper BG, Holmes S, Small IR. Diagnostic spirometry in primary care: proposed standards for general practice compliant with American Thoracic Society and European Respiratory Society recommendations. *Prim Care Resp J* 2009; 18: 130-147.
26. International Primary Care Respiratory Group. Opinion Sheet on Spirometry.
27. Parkes G, Greenhalgh T, Griffin M, Dent R. Effect on smoking quit rate of telling patients their lung age: the Step2quit randomised controlled trial. *BMJ.* 2008; 336: 598-600.
28. Price DB, Tinkelman DG, Halbert RJ, Nordyke RJ, Isonaka S, Nonikov D, et al. Symptom-based questionnaire for identifying COPD in smokers. *Respiration.* 2006; 73: 285-295.
29. Price DB, Tinkelman DG, Nordyke RJ, Isonaka S, Halbert RJ. COPD Questionnaire Study Group. Scoring system and clinical application of COPD diagnostic questionnaires. *Chest.* 2006; 129: 1531-1539.
30. Van Schayck CP, Loozen JM, Wagena E, Akkermans RP, Wesseling GJ. Detecting patients at a high risk of developing chronic obstructive pulmonary disease in general practice: cross sectional case finding study. *BMJ.* 2002; 324: 1370.
31. Martinez FJ, Raczek AE, Seifer FD, Conoscenti CS, Curtice TG, D'Eletto T, et al. Development and initial validation of a self-scored COPD Population Screener Questionnaire (COPD-PS). *COPD.* 2008; 5: 85-95.
32. Frith P, Crockett A, Beilby J, Marshall D, Attewell R, Ratnanesan A, et al. Simplified COPD screening: validation of the PiKo-6® in primary care. *Prim Care Respir J.* 2011; 20: 190-198, 2 p following 198.
33. Thorn J, Tilling B, Lisspers K, Jörgensen L, Stenling A, Stratelis G. Improved prediction of COPD in at-risk patients using lung function pre-screening in primary care: a real-life study and cost-effectiveness analysis. *Prim Care Respir J.* 2012; 21: 159-166.
34. Sichelidis L, Spyrtos D, Papaioannou M, Chloros D, Tsiotsios A, Tzagaraki V, et al. A combination of the IPAG questionnaire and PiKo-6® flow meter is a valuable screening tool for COPD in the primary care setting. *Prim Care Respir J.* 2011; 20: 184-189, 1 p following 189.
35. Westwood M, Bourbeau J, Jones PW, Cerulli A, Capkun-Niggli G, Worthy G. Relationship between FEV1 change and patient-reported outcomes in randomised trials of inhaled bronchodilators for stable COPD: a systematic review. *Respir Res.* 2011; 12: 40.
36. van der Molen T, Diamant Z, Kocks JW, Tsiligianni IG. The use of health status questionnaires in the management of chronic obstructive pulmonary disease patients in clinical practice. *Expert Rev Respir Med.* 2014; 8: 479-491.
37. Kocks JW, Kerstjens HA, Snijders SL, de Vos B, Biermann JJ, Van Hengel P, et al. Health status in routine clinical practice: validity of the clinical COPD questionnaire at the individual patient level. *Health Qual Life Outcomes.* 2010; 8: 135.
38. Jones R, Price D, Chavannes N, Van de Molen T, Thomas M, Tsiligianni I, et al. GOLD COPD categories are not fit for purpose in primary care. *Lancet Respir Med.* 2013; 1: e17.
39. Kocks J, de Jong C, Berger MY, Kerstjens HA, Van der Molen T. Putting health status guided COPD management to the test: protocol of the MARCH study. *BMC Pulm Med.* 2013; 13: 41.
40. Kruis AL, Stållberg B, Jones RC, Tsiligianni IG, Lisspers K, Van der Molen T, et al. Primary care COPD patients compared with large pharmaceutically-sponsored COPD studies: an UNLOCK validation study. *PLoS One.* 2014; 9: e90145.
41. Hurst JR, Vestbo J, Anzueto A, Locantore N, Müllerova H, Tal-Singer R, et al. Susceptibility to exacerbation in chronic obstructive pulmonary disease. *N Engl J Med.* 2010; 363: 1128-1138.
42. Eltboli O, Brightling CE. Eosinophils as diagnostic tools in chronic lung disease. *Expert Rev Respir Med.* 2013; 7: 33-42.
43. Tsiligianni IG, Kosmas E, Van der Molen T, Tzanakis N. Managing comorbidity in COPD: a difficult task. *Curr Drug Targets.* 2013; 14: 158-176.
44. Anecchino C, Rossi E, Fanizza C, De Rosa M, Tognoni G, Romero M. Working group ARNO project. Prevalence of chronic obstructive pulmonary disease and pattern of comorbidities in a general population. *Int J Chron Obstruct Pulmon Dis.* 2007; 2: 567-574.
45. Gershon AS, Mecredy GC, Guan J, Victor JC, Goldstein R, To T. Quantifying comorbidity in individuals with COPD: a population study. *Eur Respir J.* 2015; 45: 51-59.
46. Rutten FH, Zuithoff NP, Hak E, Grobbee DE, Hoes AW. Beta-blockers may reduce mortality and risk of exacerbations in patients with chronic obstructive pulmonary disease. *Arch Intern Med.* 2010; 170: 880-887.
47. Komajda M, Follath F, Swedberg K, Cleland J, Aguilar JC, Cohen-Solal A, et

- al. The EuroHeart Failure Survey programme--a survey on the quality of care among patients with heart failure in Europe. Part 2: treatment. *Eur Heart J*. 2003; 24: 464-474.
48. Egred M, Shaw S, Mohammad B, Waitt P, Rodrigues E. Under-use of beta-blockers in patients with ischaemic heart disease and concomitant chronic obstructive pulmonary disease. *QJM*. 2005; 98: 493-497.
49. Fabbri LM, Boyd C, Boschetto P, Rabe KF, Buist AS, Yawn B, et al. ATS/ERS Ad Hoc Committee on Integrating and Coordinating Efforts in COPD Guideline Development. How to integrate multiple comorbidities in guideline development: article 10 in integrating and coordinating efforts in COPD guideline development. An official ATS/ERS workshop report. *Proc Am Thorac Soc*. 2012; 9: 274-281.