

Towards a collaborative approach in chronic obstructive pulmonary disease: an expert opinion paper on cardiopulmonary risk reduction in clinical practice in Romania

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Abstract

English:

Chronic obstructive pulmonary disease (COPD) is frequently complicated by cardiovascular diseases (CVDs), worsening patient outcomes. Recognising COPD as a multimorbid and syndemic condition, we developed this position paper to provide evidence-based recommendations for reducing cardiopulmonary (CP) risk through integrated, guideline-directed management in Romania. In June–September 2025, the multidisciplinary task force (pulmonologists/cardiologists/internal medicine specialists/general practitioners GPs), with expertise in diagnosis and treatment of patients with COPD and CVDs, engaged in working groups and developed algorithms for COPD management, focusing on CP risk reduction. Expert recommendations were formulated using current guidelines, scientific literature evidence and clinical expertise as follows: (i) prioritise early COPD diagnosis in at-risk populations; (ii) address CP risk assessment at every visit, with urgent pulmonology referral for severe COPD symptoms and cardiology referral for severe dyspnoea with abnormal electrocardiogram; (iii) ensure adequate assessment and management of mild–moderate COPD exacerbations, with urgent emergency referral/hospitalisation when severity cannot be reliably determined; (iv) refer all patients with severe COPD exacerbations to emergency care and/or hospitalisation; (v) implement standardised COPD hospital discharge protocol; (vi) optimise pharmacotherapy by initiating long-acting β_2 -agonist (LABA) + long-acting muscarinic antagonist (LAMA) and escalating to triple therapy (LABA/LAMA/inhaled corticosteroid) when indicated and ensure structured follow-up as per GOLD guidelines. This expert opinion paper calls for urgent action to bridge the gap between guideline-directed recommendations and routine practice, with a strict policy of zero tolerance for treatment delays.

Keywords


chronic obstructive pulmonary disease • cardiopulmonary risk • expert opinion • routine clinical practice • Romania

Către o abordare colaborativă în boala pulmonară obstructivă cronică: un articol de opinie al experților privind reducerea riscului cardiopulmonar în practica clinică din România

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Rezumat

Romanian:

Complicațiile bronhopneumopatiei obstructive cronice (BPOC) apar frecvent în contextul prezenței bolilor cardiovasculare (BCV), cu rezultate nefavorabile pentru pacienți. Recunoscând BPOC ca o afecțiune multimorbidă și sindemică, am elaborat acest document de poziție pentru a oferi recomandări bazate pe dovezi pentru reducerea riscului cardiopulmonar prin management integrat în România, conform recomandărilor ghidurilor. În perioada iunie-septembrie 2025, grupul de lucru multidisciplinar (pneumologie/cardiologie/medicină internă/medicină de familie) cu experiență în diagnosticarea și tratarea pacienților cu BPOC și BCV s-a implicat în sesiuni de lucru și a elaborat algoritmi pentru managementul pacienților cu BPOC, cu accent pe reducerea riscului cardiopulmonar. Recomandările experților au fost formulate pe baza ghidurilor actuale, a dovezilor din literatura științifică și a expertizei clinice, astfel: (i) prioritizarea diagnosticului BPOC precoce în populațiile cu risc; (ii) evaluarea riscului cardiopulmonar la fiecare consultație medicală, cu trimitere urgentă la pneumologie pentru simptome severe de BPOC și la cardiologie pentru dispnee severă cu modificări pe electrocardiogramă; (iii) asigurarea evaluării și gestionării adecvate a exacerbărilor ușoare-moderate BPOC, cu trimitere urgentă la camera de gardă/spitalizare atunci când severitatea nu poate fi stabilită clar; (iv) recomandare pentru tratament de urgență/spitalizare pentru toți pacienții cu exacerbări severe BPOC; (v) implementarea unui protocol standardizat la externarea pacienților cu BPOC; (vi) optimizarea tratamentului prin inițierea tratamentului cu β_2 -agonist cu durată lungă de acțiune (BADLA) + antimuscarinic cu durată lungă de acțiune (ACDLA) și trecerea la terapia triplă (ACDLA/BADLA/corticosteroid inhalator) atunci dacă este indicat, cu monitorizare structurată conform recomandărilor GOLD. Acest document de opinie a experților solicită măsuri urgente pentru a reduce decalajul de implementare a recomandărilor din ghiduri în practica clinică, cu o politică strictă de toleranță zero pentru inerția terapeutică.

Cuvinte-cheie

boala pulmonară obstructivă cronică • risc cardiopulmonar • opinie de expert • practica clinică curentă, România

Introduction

Chronic obstructive pulmonary disease (COPD) is a progressive disease with consequences that extend far beyond the respiratory system (1). The currently estimated prevalence of COPD is approximately 10% globally (2), and projections for Europe predict a rise at almost 40% by 2050 (3). In Romania, the prevalence of COPD has been estimated at 9.3% among individuals over 40 years old in a national survey performed by the Romanian Society of Pneumology (4).

Almost all patients with COPD have at least one additional significant chronic condition (1, 5). Due to the intricate reciprocal interactions between COPD and its comorbidities, the disease should not be viewed as a single entity, but rather as a multimorbid state, with significant implications for diagnosis, treatment and patient care (6). Consequently, the syndemic approach has been recently introduced for better characterisation of COPD, emphasising the interconnectivity of biological determinants and shared risk factors, while promoting a move towards an integrated management strategy (1, 7).

Among the many comorbid conditions associated with COPD, cardiovascular diseases (CVDs) are one of the most significant, affecting 20%–70% of patients (8), and are recognised as a major risk factor for patients by the Global Initiative for Chronic Obstructive Lung Disease (GOLD) guidelines (9, 10). Taken together, they have a greater impact on the health status and a more severe prognosis than each condition alone (7, 8, 11). The presence of

cardiovascular (CV) comorbidities increases the risk and severity of COPD exacerbations, which, in turn, increases the risk of acute CV events, hospitalisation and death (9, 10, 12–14). The vicious downward spiral linking these two conditions is representative of the cardiopulmonary (CP) risk, defined as ‘the risk of serious respiratory and/or CV events in patients with COPD’, including, but not limited to, COPD exacerbation, myocardial infarction, stroke, congestive heart failure, arrhythmias and death from any of these conditions (8).

A significant step in changing perspectives for the management of COPD patients has been taken with the inclusion of mortality reduction as a treatment goal in the 2023 update of the GOLD recommendations, based on results from the efficacy and safety of triple therapy in obstructive lung disease (ETHOS) and informing the pathway of COPD treatment (IMPACT) trials (15, 16). Moreover, the ETHOS trial showed that reducing mortality risk with triple therapy (long-acting β_2 -agonist [LABA] + long-acting muscarinic antagonist [LAMA] + inhaled corticosteroid) compared to dual bronchodilation (LAMA + LABA) has been driven by a decrease in CV death (16). According to the GOLD 2025, recently updated GOLD 2026 guidelines and considering the symptoms and/or exacerbations, the current guidelines-directed medical treatment (GDMT) for maintenance in COPD includes dual bronchodilation or triple therapy, preferably administered in a single inhaler (9, 10).

Considering all the above, prevention of CP risk and implementation of GDMT are now top priorities in COPD management (14, 17, 18). Because COPD patients require a multidisciplinary approach throughout their disease journey, all healthcare specialists commonly involved in their care should take appropriate actions to address CP risk as part of routine practice (1, 7, 17, 18).

Objectives

The algorithms and recommendations for COPD management included in this paper were developed specifically for healthcare professionals (HCPs) to provide clear, practical guidance on decision-making strategies that help reduce CP risk in these patients.

The objective was to outline key recommendations to improve continuity of care, facilitating timely identification and management of risk factors for acute COPD exacerbations and CV events. The target audience includes pulmonologists, cardiologists, internists and family physicians, as well as other clinicians involved in COPD–CVD care.

Materials and methods

During June–September 2025, a multidisciplinary task force including 11 HCPs with expertise in the diagnosis and treatment of patients with COPD and CVDs in both inpatient and outpatient settings gathered to discuss how to better address CP risk reduction in patients with COPD in Romania, based on the updated 2025 GOLD.

At the initial meeting, this multidisciplinary group of pulmonologists, cardiologists and specialists in internal and family medicine engaged in working groups to review evidence from the scientific literature related to the prevention of acute cardio-pulmonary events and mortality in COPD. No systematic literature search was conducted.

The algorithms, further developed and presented here, are based on international guidelines and clinical experience and include landmarks for referral of patients with COPD to other specialists for additional investigations and adequate treatment. The recommendations included in this document reflect the group discussions and have been approved by all members of the task force.

Results

A consensus was reached on the critical need to raise awareness of CP risk and improve the current standard of care in COPD in Romania. The following recommendations for CP risk reduction strategies in COPD through an interdisciplinary approach were developed and validated.

Modifiable risk factors

Recommendation I

We strongly recommend the early identification of COPD in at-risk populations (e.g. current/former smokers, individuals with persistent respiratory symptoms), recognising that timely diagnosis contributes significantly to the prevention of CP complications and overall reduction in morbidity and mortality.

Rationale

COPD and CVDs share common risk factors and pathogenic mechanisms, a concept now recognised as a syndemic occurrence. Although anatomical and pathophysiological links between COPD and CVDs are not fully understood, they appear to be multifactorial, interactive and mutually aggravating: lung hyperinflation, hypoxaemia, pulmonary hypertension, systemic inflammation and oxidative stress, exacerbations, shared risk factors and shared genetics, as well as COPD phenotype (11, 18–20).

Diagnosis of COPD and assessment of CP risk

Recommendation II

Irrespective of the medical speciality, every medical visit should be an opportunity to estimate CP risk and take appropriate actions in patients with COPD: identify CV risk factors from the COPD diagnosis (10), refer to an appropriate specialist if necessary and optimise treatment (7–12, 18). Figure 1 includes potential touchpoints and assessments.

Rationale

Diagnosis of COPD is based on medical history (smoking, recurrent respiratory infections), clinical manifestations (dyspnoea, wheezing, cough and sputum not explained by other conditions) and spirometry (forced expiratory volume in 1 s [FEV₁]/forced vital capacity [FVC] <0.7) (9). In Romania, the diagnosis of COPD is mainly made by pulmonologists, due to current limitations in the competencies assigned to family physicians. Improved access to spirometry (micro spirometers or portable spirometers) could help diagnose COPD at early stages (21).

Recommendation III

If already implemented in routine practice, we recommend the use of QRISK3 (22) or SCORE2 (23) in COPD populations in addition to lung function parameters of the spirometry, where available.

Rationale

Related to CV risk assessment in clinical practice, the current risk calculators neither include COPD, nor were they validated in populations with COPD (7). Therefore, they

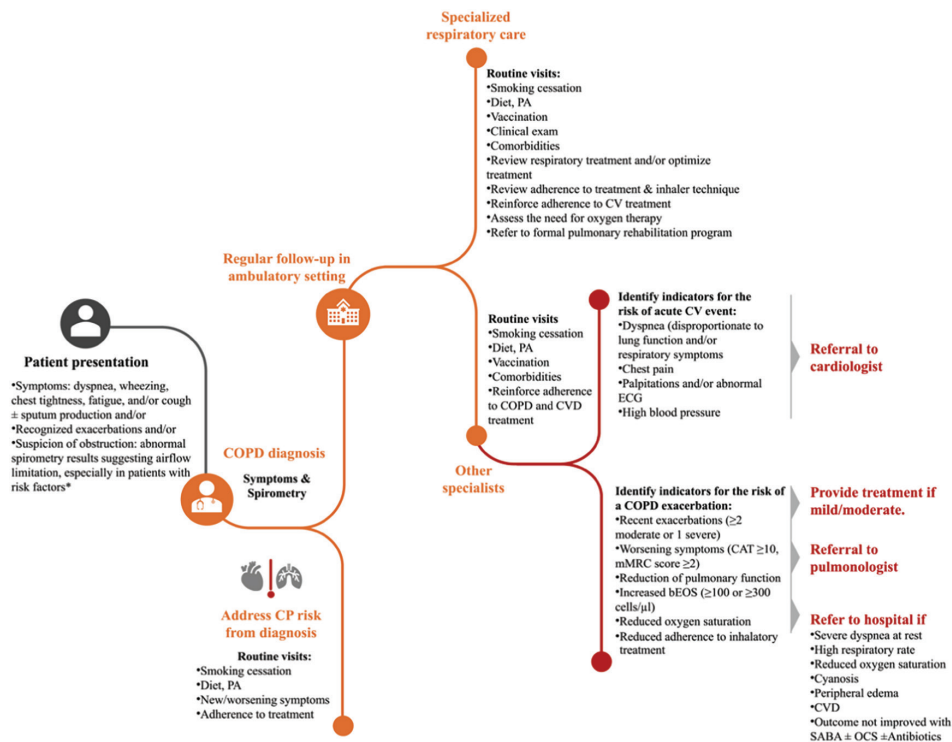


Figure 1. Routine assessment of cardiopulmonary CP risk in COPD and triggers for referral (9, 11, 18). *Smoking history, environmental exposure, family history of respiratory diseases, and so forth. bEOS, blood eosinophils; CAT, COPD assessment test; COPD, chronic obstructive pulmonary disease; CP, cardiopulmonary; CV, cardiovascular; CVD, cardiovascular disease; ECG, electrocardiogram; mMRC, (modified) Medical Research Council Dyspnea/Dyspnoea; OCS, oral corticosteroid; PA, physical activity; SABA, short-acting bronchodilator.

may underestimate CVD in patients with COPD (24, 25). Incorporating spirometry results, such as FEV₁% predicted value and FEV₁/FVC ratio, into CV assessments may improve predictive accuracy of standard CV risk scores in COPD populations, as reduced lung function has been shown to independently predict incident CV events beyond traditional risk factors (25, 26).

COPD exacerbations

Recommendation IV

Irrespective of the medical speciality, the mild and moderate exacerbations of COPD (ECOPDs) should be adequately assessed, investigated and managed. When it is not possible to evaluate the severity, patients should be referred to emergency care and/or hospitalised.

Rationale

Mild and moderate ECOPDs are frequently underreported by patients (27), as they may perceive symptom fluctuations as 'normal'. Therefore, clinicians interacting with patients with COPD should be aware of trigger words that may reflect the experience of an exacerbation. This is essential because the CV risk, including CV death, increases

significantly during and after acute episodes, likely in relation to worsening of the factors that contribute to CVDs during periods of clinical stability (2, 9–11). Moreover, the risk remains high for several weeks after discharge for ECOPD and may persist for up to 1 year later (9, 28–30). Routine assessment of CVD markers in case of suspicion of exacerbation is preferred for an accurate differential diagnosis and management strategy (18).

Recommendation V

All patients with severe ECOPD should be referred to emergency care and/or hospitalisation (9).

Rationale

Hospital treatment is indicated if at least one of the following is present (9):

- Severe symptoms such as sudden worsening of resting dyspnoea, high respiratory rate, decreased oxygen saturation, confusion and drowsiness;
- Acute respiratory failure;
- Onset of new physical signs (e.g. cyanosis, peripheral oedema);
- Failure of an exacerbation to respond to initial medical management;

- Presence of severe comorbidities (e.g. heart failure, newly occurring arrhythmias)
- Moderate to severe exacerbation, with insufficient home support.

Hospitalisation should include comprehensive education for the patient and caregivers, providing information on the disease, impact of exacerbations, action plans and reasons for adherence to treatment (31, 32).

Recommendation VI

Provide a standardised discharge protocol for all hospitalised patients with COPD. The main aspects to be included in the hospital discharge protocol are presented in Figure 2 (9, 25, 31).

Recommendation VII

Ensure timely initiation of dual long-acting bronchodilation with LABA + LAMA and escalation to triple therapy with LABA + LAMA + inhaled corticosteroid when indicated, in line with GOLD 2025 recommendations (9), with structured follow-up to monitor adherence, inhaler technique, symptoms/exacerbations and lung function and to adjust therapy accordingly.

Rationale

The discharge letters, usually provided to patients, include a summary of investigations, procedures and treatments

performed during hospitalisation, as well as recommendations for pharmacological and non-pharmacological treatment and follow-up time intervals. Improved discharge protocols can better support communication between HCPs managing COPD patients, further promoting CP risk reduction (9, 31).

Discussions

A unique feature of this expert opinion paper is that it was specifically created for CP risk reduction in COPD, in accordance with GOLD recommendations and recent literature addressing this topic (1, 9–11, 18). This may serve as a formal starting point for the development of specific protocols to be implemented in the Romanian clinical practice. CVDs significantly impact the health status of patients with COPD, being a major cause of mortality, both in acute exacerbations and in clinically stable patients with mild–moderate airflow limitations. The frequent coexistence of COPD and CVD reflects bidirectional pathophysiology. COPD-related abnormalities, such as persistent systemic inflammation, abnormal pulmonary gas exchange, lung hyperinflation and exertional dyspnoea, promote CV injury and events. In turn, CVD worsens the COPD status by causing abnormal myocardial contractility, increased left-ventricular filling pressures and subsequent lung stasis, and reduced skeletal muscle oxygen delivery, thereby amplifying dyspnoea and exercise intolerance (2, 7, 8) (Figure 3). This reciprocal

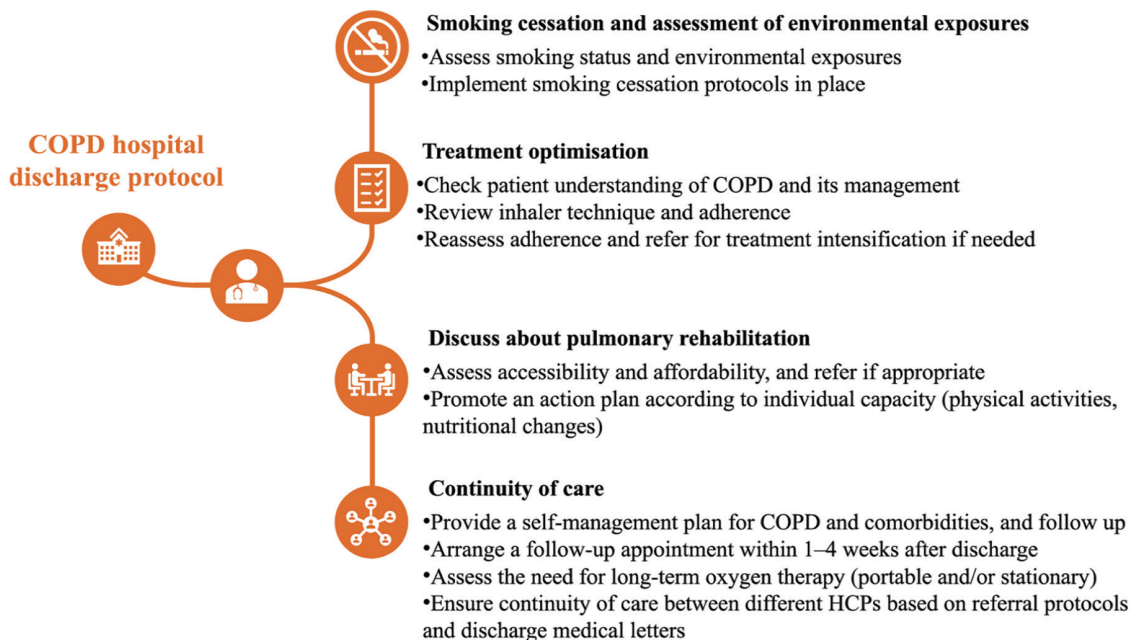


Figure 2. Main aspects to be included and addressed in the COPD hospital discharge protocol (9, 25, 31). COPD, chronic obstructive pulmonary disease; HCPs, healthcare professionals.

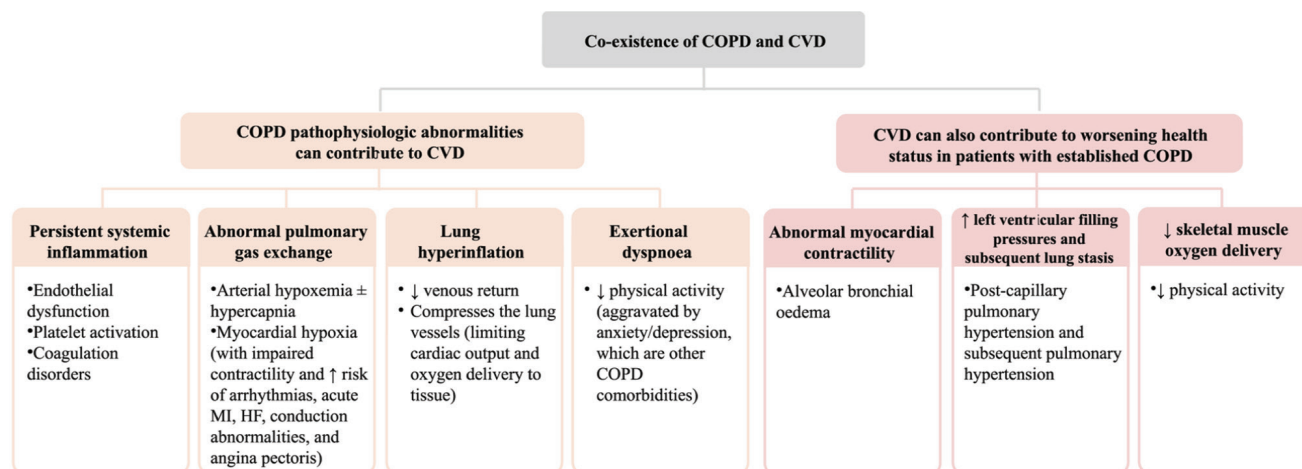


Figure 3. The mechanism underlying the frequent coexistence of COPD and CVD (2, 7, 8). COPD, chronic obstructive pulmonary disease; CVD, cardiovascular disease; HF, heart failure; MI, myocardial infarction.

loop explains their high co-occurrence and supports the integration of CP risk assessment and management.

COPD remains largely underdiagnosed, with estimates suggesting that more than 70% of individuals with airflow limitation are unaware of their condition (33), pointing to the need to improve access to spirometry with early diagnosis and appropriate management (34). In Romania, COPD is diagnosed predominantly by pulmonologists because current scope-of-practice regulations restrict spirometry in family physicians' offices. Considering the high rate of underdiagnosis and the challenges in accessing specialised care, extending the competencies of family physicians to perform spirometry in primary healthcare settings is a topic of interest to be further explored. Incorporation of spirometry results (such as FEV₁ or airflow-obstruction indices) into CV risk stratification could improve the accuracy of standard CV risk scores (26), aligning pulmonary diagnostics with CV prevention and helping to reduce CP complications, as well as overall morbidity and mortality. Moreover, early diagnosis of COPD allows for timely modification of risk factors (such as smoking cessation, vaccination and pulmonary rehabilitation), optimisation of inhaled therapy, and systematic screening and management of CV comorbidities. Nevertheless, the updated 2026 GOLD recommendations include testing of natriuretic peptide at COPD diagnosis and annually thereafter (10), emphasising the efforts needed for timely identification of prevalent heart disease.

In patients with COPD with worsening respiratory symptoms and/or reduced lung function, ECOPD should be suspected. As per the current definition (9, 10), a change in symptom severity should require a change in the current treatment. Considering the high impact of an exacerbation on the CV risk and our aim to minimise the CP events, we recommend an

initial treatment trial with a bronchodilator [short acting] and, where indicated, an antibiotic and/or systemic corticosteroid for 5 days. To avoid antimicrobial overuse and resistance, antibiotic therapy should be reserved for patients with increased sputum purulence, with or without increased sputum volume or worsening dyspnoea, according to the GOLD 2025 recommendations (9).

Regarding COPD hospitalisations, a standardised discharge protocol with a focus on CP risk can strengthen care transitions and reduce early post-discharge events. Key elements include (i) patient and caregiver education on COPD, identification of exacerbations and CP risk; (ii) personalised written action plans for early recognition and management of exacerbation symptoms; (iii) reinforcement of adherence to COPD maintenance treatment, including correct inhaler technique; (iv) clear post-discharge management plan that integrates pharmacological and non-pharmacological measures, with triple therapy (LABA + LAMA + inhaled corticosteroid) from discharge in patients with high blood eosinophil levels and (v) scheduled follow-up within 1–4 weeks with appropriate specialists (pulmonology, cardiology, family medicine), ensuring cross-level communication (discharge summary sent to family physicians and relevant specialists) (9, 10).

Conclusions

Considering the burden of COPD and the risks that COPD patients face daily, we strongly advocate for the adoption of a policy of zero tolerance for clinical inertia in COPD care in Romania, especially for reducing exacerbations and associated CV risk (9, 10, 23, 35). In conclusion, it is imperative to take decisive actions. Bridging the gap between GDMT

recommendations and clinical practice remains a priority. Efforts to translate GDMT recommendations into real-world clinical settings need to be strengthened. We should start by identifying patients at risk in a timely manner and making immediate treatment adjustments. The implementation of GDMT in clinical practice, namely dual bronchodilation or triple therapy, and intensive treatment of CV risk factors according to the current clinical guidelines, should be closely followed up for optimal outcomes in patients with COPD.

The interdisciplinary care of patients with COPD will be based on the following guiding principles:

- Modifiable risk factors for CP events, such as smoking cessation, vaccination, comorbidity screening and lifestyle modification, should be part of every visit.
- Provide an early and accurate diagnosis. All patients with severe symptoms of COPD should be immediately referred to a pulmonologist and, in case of severe dyspnoea and abnormal electrocardiogram, the immediate referral should be to a cardiologist.
- Expand access to spirometry in primary care, including training and certification for family physicians.
- Implement clear, standardised protocols at the institutional level for exacerbation management, hospital discharge and inter-speciality communication to streamline referrals and management of patients with COPD and CP risk.
- Optimise treatment by ensuring the timely initiation of dual bronchodilation or triple therapy, in line with the GOLD guidelines (9, 10), with close follow-up for adherence and outcomes.
- Call for action – Zero Tolerance for Clinical Inertia. Act early, optimise therapy and coordinate care.

Disclosures

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Conflict of interest

The authors declare no conflict of interest.

Informed consent statement

No informed consent was necessary.

Author Contributions

All authors made significant contributions to the development of the manuscript submitted (review of medical literature and interpretation, participation to group discussion, paper writing and reviewing the manuscript).

References

1. Fabbri LM, Celli BR, Agustí A, Criner GJ, Dransfield MT, Divo M, et al. COPD and multimorbidity: recognising and addressing a syndemic occurrence. *The Lancet Respiratory Medicine*. 2023;11: 1020–1034. doi:10.1016/S2213-2600(23)00261-8
2. Boers E, Barrett M, Su JG, Benjafield AV, Sinha S, Kaye L, et al. Global burden of chronic obstructive pulmonary disease through 2050. *JAMA Network Open*. 2023;6: e2346598. doi:10.1001/jamanetworkopen.2023.46598
3. Benjafield A, Tellez D, Barrett M, Gondalia R, Nunez C, Wedzicha J, et al. An estimate of the European prevalence of COPD in 2050. *European Respiratory Journal*. 2021;58: OA2866.

4. Mihălțan F, Nemes R, Daramus I, Paunescu B, Farcasanu D. Prevalence of chronic obstructive pulmonary disease (COPD) in Romania. *Chest*. 2012;142: 658A. doi:10.1378/chest.1389539
5. Duszyk K, McLoughlin RF, Gibson PG, McDonald VM. The use of treatable traits to address COPD complexity and heterogeneity and to inform the care. *Breathe (Sheffield, England)*. 2021;17: 210118. doi:10.1183/20734735.0118-2021
6. Morgan AD, Zakeri R, Quint JK. Defining the relationship between COPD and CVD: what are the implications for clinical practice? *Therapeutic Advances in Respiratory Disease*. 2018;12: 1753465817750524. doi:10.1177/1753465817750524
7. Hawkins NM, Kaplan A, Ko DT, Penz E, Mohit Bhutani M. Is 'Cardiopulmonary' the new 'Cardiometabolic'? making a case for systems change in COPD. *Pulmonary therapy*. 2024;10: 363–376. doi:10.1007/s41030-024-00270-2
8. Singh D, Han MK, Hawkins NM, Hurst JR, Kocks JWH, Skolnik N. Implications of cardiopulmonary risk for the management of COPD: a narrative review. *Advances in Therapy*. 2024;41: 2151–2167. doi:10.1007/s12325-024-02855-4
9. *Global initiative for chronic obstructive lung disease (GOLD) 2025 report*. Available from: <https://goldcopd.org/2025-gold-report/> [Accessed June 2025].
10. *Global initiative for chronic obstructive lung disease (GOLD) 2026 report*. Available from: <https://goldcopd.org/2026-gold-report-and-pocket-guide/> [Accessed November 2025].
11. de Miguel-Díez J, Núñez Villota J, Santos Pérez S, Manito Lorite N, Alcázar Navarrete B, Delgado Jiménez JF, et al. Multidisciplinary management of patients with chronic obstructive pulmonary disease and cardiovascular disease. *Archivos de Bronconeumología*. 2024;60: 226–237. doi:10.1016/j.arbres.2024.01.013
12. Hurst JR, Gale CP. MACE in COPD: addressing cardiopulmonary risk. *The Lancet Respiratory Medicine*. 2024;12: 345–348. doi:10.1016/S2213-2600(24)00038-9
13. Bourbeau J, Bhutani M, Hernandez P, Aaron SD, Beauchesne M-F, Kermelley SB, et al. 2023 Canadian Thoracic Society Guideline on pharmacotherapy in patients with stable COPD. *Chest*. 2023;164: 1159–1183. doi:10.1016/j.chest.2023.08.014
14. Shrikrishna D, Taylor CJ, Stonham C, Gale CP. Exacerbating the burden of cardiovascular disease: how can we address cardiopulmonary risk in individuals with chronic obstructive pulmonary disease? *European Heart Journal*. 2024;45: 247–249. doi:10.1093/eurheartj/ehad669
15. Martinez FJ, Rabe KF, Ferguson GT, Wedzicha JA, Singh D, Wang C. Reduced all-cause mortality in the ETHOS trial of budesonide/glycopyrrolate/formoterol for Chronic obstructive pulmonary disease. A randomized, double-blind, multicenter, parallel-group study. *American Journal of Respiratory and Critical Care Medicine*. 2021;203: 553–564. doi:10.1164/rccm.202006-2618OC
16. Wells JM, Criner GJ, Halpin DMG, Han MK, Jain R, Lange P, et al. Mortality Risk and Serious Cardiopulmonary Events in Moderate-to-Severe COPD: Post Hoc Analysis of the IMPACT Trial. *Chronic Obstr Pulm Dis*. 2023; 10(1): 33–45. doi: 10.15326/jcopdf.2022.0332
17. Stolz D, Mkorombindo T, Schumann DM, Agusti A, Ash SY, Bafadhel M. Towards the elimination of chronic obstructive pulmonary disease: a Lancet Commission. *Lancet*. 2022;400: 921–972. doi:10.1016/S0140-6736(22)01273-9
18. Gale CP, Hurst JR, Hawkins NM, Bourbeau J, Han MLK, Lam CSP, et al. Identification and management of cardiopulmonary risk in patients with COPD: a multidisciplinary consensus and modified Delphi study. *European Journal of Preventive Cardiology*. 2025;32(15): 1445–1460. doi:10.1093/eurjpc/zwaf119
19. Rabe KF, Hurst JR, Suissa S. Cardiovascular disease and COPD: dangerous liaisons? *European Respiratory Review*. 2018;27(149): 180057. doi:10.1183/16000617.0057-2018
20. Rogliani P, Ritondo BL, Laitano R, Chetta A, Calzetta L. Advances in understanding of mechanisms related to increased cardiovascular risk in COPD. *Expert Review of Respiratory Medicine*. 2021;15: 59–70. doi:10.1080/17476348.2021.1840982
21. Jankowski P, Mycroft K, Górska K, Korczyński P, Krenke R. How to enhance the diagnosis of early stages of chronic obstructive pulmonary disease (COPD)? The role of mobile spirometry in COPD screening and diagnosis—a systematic review. *Advances in Respiratory Medicine*. 2024;92: 158–174. doi:10.3390/arm92020018
22. *QRISK 3 score*. Available from: [QRISK.org](https://www.qrisk.org/) [Accessed October 2025].
23. SCORE2 Working Group and ESC Cardiovascular Risk Collaboration. SCORE2 risk prediction algorithms: new models to estimate 10-year risk of cardiovascular disease in Europe. *European Heart Journal*. 2021;42: 2439–2454. doi:10.1093/eurheartj/ehab309
24. Amegadzie JE, Gao Z, Quint JK, Russell R, Hurst JR, Lee TY, et al. QRISK3 underestimates the risk of cardiovascular events in patients with COPD. *Thorax*. 2024;79: 718–724. doi:10.1136/thorax-2023-220615
25. Fawzy A, Vogelmeier CF. Standard risk scores inadequately estimate subclinical Coronary artery disease in Chronic obstructive pulmonary disease. *American Journal of Respiratory and Critical Care Medicine*. 2025;211: 889–890. doi:10.1164/rccm.202411-2240ED
26. Zhou L, Yang H, Zhang Y, Wang Y, Zhou X, Liu T. Predictive value of lung function measures for cardiovascular risk: a large prospective cohort study. *Thorax*. 2024;79: 250–258. doi:10.1136/thorax-2023-220703
27. Jones P, Alzaabi A, Casas Herrera A, Polatli M, Rabahi MF, Telles AC. Understanding the gaps in the reporting of COPD exacerbations by patients: a review. *COPD*. 2024;21: 2316594. doi:10.1080/15412555.2024.2316594
28. Calabria S, Ronconi G, Dondi L, Dondi L, Dell'Anno I, Nordon C, et al. Cardiovascular events after exacerbations of chronic obstructive pulmonary disease: results from the EXAcerbations

- of COPD and their OutcomeS in CardioVascular diseases study in Italy. *European Journal of Internal Medicine*. 2024;127: 97–104. doi:10.1016/j.ejim.2024.04.021
29. Hawkins NM, Nordon C, Rhodes K, Talukdar M, McMullen S, Ekwaru P, et al. Heightened long-term cardiovascular risks after exacerbation of chronic obstructive pulmonary disease. *Heart*. 2024;110: 702–709. doi:10.1136/heartjnl-2023-323487
30. Nordon C, Simons SO, Marshall J, Müllerová H, Pollack M, Bengtsson C, et al. The sustained increase of cardiovascular risk following COPD exacerbations: meta-analyses of the EXACOS-CV studies. *ERJ Open Research*. 2025;11(3): 01091–2024. doi:10.1183/23120541.01091-2024
31. Miravittles M, Bhutani M, Hurst JR, Franssen FME, van Boven JFM, Khoo EM. Implementing an evidence-based COPD hospital discharge protocol: a narrative review and expert recommendations. *Advances in Therapy*. 2023;40: 4236–4263. doi:10.1007/s12325-023-02609-8
32. Lindh A, Giezeman M, Theander K, Zakrisson A-B, Westerdahl E, Stridsman C. Factors associated with patient education in patients with Chronic obstructive pulmonary disease (COPD) - A primary health care register-based Study. *International Journal of Chronic Obstructive Pulmonary Disease*. 2024;19: 1069–1077. doi:10.2147/COPD.S455080
33. Diab N, Gershon AS, Sin DD, Tan WC, Bourbeau J, Boulet LP, et al. Underdiagnosis and overdiagnosis of Chronic obstructive pulmonary disease. *American Journal of Respiratory and Critical Care Medicine*. 2018;198: 1130–1139. doi:10.1164/rccm.201804-0621CI
34. Guecamburu M, Klein E, Verdy G, Nocent-Ejnaini C, Macey J, Portel L, et al. Factors associated with undiagnosed COPD in patients hospitalised for severe COPD exacerbation. *BMJ Open Respiratory Research*. 2024;11(1): e002620. doi:10.1136/bmjresp-2024-002620
35. Singh D, Holmes S, Adams C, Bafadhel M, Hurst JR. Overcoming therapeutic inertia to reduce the risk of COPD exacerbations: four action points for healthcare professionals. *International Journal of Chronic Obstructive Pulmonary Disease*. 2021;16: 3009–3016. doi:10.2147/COPD.S329316