

Online Supplement

Determinants for changing the treatment of COPD: a regression analysis from a clinical audit

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Table 1S. Bivariate analysis of factors associated with a step-up of treatment.

| | No step-up (n = 466) | Step-up (n = 99) | P-Value* |
|---|-------------------------|---------------------|----------|
| Monographic COPD clinic | 187 (40.1) | 42 (42.4) | NS |
| Outpatient clinic located within the hospital | 455 (97.6) | 96 (97.0) | NS |
| Male gender | 69 (14.8) | 16 (16.2) | NS |
| Age (years) | 68.4 (9.5) | 69.4 (10.4) | NS |
| Rural area | 135 (20.9) | 28 (28.3) | NS |
| Current smoker | 104 (23.4) | 27 (28.7) | NS |
| Tobacco history (pack-year) | 56.0 (32.1) | 51.1 (25.3) | NS |
| Acute myocardial infraction | 43 (9.2) | 15 (15.2) | 0.099 |
| Charlson | 2.1 (1.6) | 2.2 (1.3) | NS |
| GesEPOC phenotype A | 222 (47.6) | 37 (37.4) | NS |
| GesEPOC phenotype B | 47 (10.1) | 12 (12.1) | NS |
| GesEPOC phenotype C | 39 (8.4) | 7 (7.1) | NS |
| GesEPOC phenotype D | 68 (14.6) | 17 (17.2) | NS |
| GOLD type A | 89 (19.1) | 20 (20.2) | NS |
| GOLD type B | 45 (9.7) | 14 (14.1) | NS |
| GOLD type C | 81 (17.4) | 16 (16.2) | NS |
| GOLD type D | 88 (18.9) | 20 (20.2) | NS |
| Dyspnea (mMRC) | 1.6 (1.0) | 1.8 (0.8) | 0.021 |
| Chronic bronchitis | 215 (46.1) | 61 (61.6) | 0.006 |
| Purulent sputum | 33 (7.1) | 9 (9.1) | NS |
| Asthma-like symptoms | 77 (16.5) | 21 (21.2) | NS |
| Exacerbations previous year | 0.8 (1.3) | 1.3 (1.6) | 0.004 |
| Hospitalizations previous year | 0.2 (0.5) | 0.2 (0.6) | NS |

| | No step-up (n = 466) | Step-up (n = 99) | P-Value* |
|--|-------------------------|---------------------|----------|
| Performs exercise | 116 (24.9) | 23 (23.2) | NS |
| Influenza vaccine | 159 (34.1) | 39 (39.4) | NS |
| Pneumococcal vaccine | 45 (9.7) | 12 (12.1) | NS |
| Receiving non-combined LAMA | 408 (87.6) | 60 (61.2) | < 0.001 |
| Receiving non-combined tiotropium | 330 (70.8) | 45 (45.9) | < 0.001 |
| Receiving non-combined aclidinium | 60 (12.9) | 8 (8.2) | NS |
| Receiving non-combined glycopyrronium | 24 (5.2) | 9 (9.3) | NS |
| Receiving non-combined LABA | 99 (21.2) | 22 (22.4) | NS |
| Receiving non-combined salmeterol | 1 (0.2) | 1 (1.0) | NS |
| Receiving non-combined formoterol | 3 (0.6) | 0 (0) | NS |
| Receiving non-combined indacaterol | 95 (20.4) | 22 (22.4) | NS |
| Receiving non-combined LABA + LAMA | 91 (19.5) | 16 (16.2) | NS |
| Receiving non-combined ICS | 28 (6.0) | 13 (13.1) | 0.019 |
| Receiving combined ICS-LABA | 315 (67.6) | 42 (42.4) | < 0.001 |
| Receiving combined salmeterol-fluticasone | 191 (41.0) | 27 (27.3) | 0.012 |
| Receiving combined formoterol-budesonide | 100 (21.5) | 11 (11.1) | 0.018 |
| Receiving combined formoterol-beclomethasone | 25 (5.4) | 4 (4.0) | NS |
| Receiving any form of ICS | 341 (73.2) | 53 (53.5) | < 0.001 |
| Receiving low dose of ICS | 36 (7.7) | 8 (8.1) | NS |
| Receiving medium dose of ICS | 164 (35.2) | 19 (19.2) | 0.094 |
| Receiving high dose of ICS | 119 (25.5) | 23 (23.2) | NS |
| Receiving triple therapy with LAMA+FDC | 273 (58.6) | 26 (26.3) | < 0.001 |
| Receiving roflumilast | 55 (11.8) | 8 (8.2) | NS |
| Receiving mucolytics | 42 (9.0) | 3 (3.1) | 0.062 |
| Receiving antibiotics | 13 (2.8) | 0 (0) | NS |

| | No step-up (n = 466) | Step-up (n = 99) | P-Value* |
|---|-------------------------|---------------------|----------|
| Receiving oral methylxanthines | 21 (4.5) | 3 (3.0) | NS |
| Receiving long-term oxygen therapy | 88 (18.9) | 18 (18.2) | NS |
| Receiving home mechanical ventilation | 23 (4.9) | 3 (3.0) | NS |
| Receiving nebulized home therapy | 17 (3.6) | 4 (4.0) | NS |
| Treatment adherence evaluation | 255 (54.7) | 65 (65.7) | 0.057 |
| Inhaler device satisfaction evaluation | 81 (17.4) | 20 (20.2) | NS |
| Adverse effects recorded | 110 (23.6) | 30 (30.3) | NS |
| FVC (%) | 74.9 (21.1) | 76.5 (19.8) | NS |
| FEV ₁ (%) | 52.3 (20.1) | 53.2 (18.0) | NS |
| FEV ₁ increase after the bronchodilator test (%) | 7.2 (9.8) | 9.2 (13.7) | NS |

Average value expressed as mean (standard deviation) or absolute (relative) frequencies depending on the nature of the variable.

* Calculated between groups using an unpaired Student t-test or a Chi-square test, depending on the nature of the variable.

LAMA: long-acting muscarinic antagonists. LABA: long-acting β_2 agonist. ICS: inhaled corticosteroids. FDC: fixed-dose combinations of an ICS and a LABA. FVC: forced vital capacity. FEV₁: forced expiratory volume in one second.

Table 2S. Factors associated with a step-down of treatment.

| | No step-down (n = 510) | Step-down (n = 55) | P-Value* |
|---|---------------------------|-----------------------|----------|
| Monographic COPD clinic | 205 (40.2) | 24 (43.6) | NS |
| Outpatient clinic located within the hospital | 497 (97.5) | 54 (98.2) | NS |
| Male gender | 81 (15.9) | 4 (7.3) | NS |
| Age (years) | 68.7 (9.8) | 68.0 (8.3) | NS |
| Rural area | 363 (71.2) | 39 (70.9) | NS |
| Current smoker | 133 (26.7) | 9 (16.4) | NS |
| Tobacco history (pack-year) | 54.0 (30.0) | 65.7 (38.3) | 0.011 |
| Depression | 47 (9.2) | 9 (16.4) | 0.098 |
| Charlson | 2.1 (1.5) | 1.9 (1.5) | NS |
| GesEPOC phenotype A | 231 (45.3) | 28 (50.9) | 0.020 |
| GesEPOC phenotype B | 57 (11.2) | 2 (3.6) | 0.049 |
| GesEPOC phenotype C | 44 (8.6) | 2 (3.6) | 0.082 |
| GesEPOC phenotype D | 79 (15.5) | 6 (10.9) | NS |
| GOLD type A | 93 (18.2) | 16 (29.1) | 0.031 |
| GOLD type B | 55 (10.8) | 4 (7.3) | NS |
| GOLD type C | 88 (17.3) | 9 (16.4) | NS |
| GOLD type D | 104 (20.4) | 4 (7.3) | 0.062 |
| Dyspnea (mMRC) | 1.7 (1.0) | 1.4 (0.9) | NS |
| Chronic bronchitis | 253 (49.6) | 23 (41.8) | NS |
| Purulent sputum | 38 (7.5) | 4 (7.3) | NS |
| Asthma-like symptoms | 95 (18.6) | 3 (5.5) | 0.036 |
| Exacerbations previous year | 0.9 (1.3) | 0.8 (1.5) | NS |
| Hospitalizations previous year | 0.2 (0.6) | 0.2 (0.6) | NS |

| | No step-down (n = 510) | Step-down (n = 55) | P-Value* |
|--|---------------------------|-----------------------|----------|
| Performs exercise | 125 (38.0) | 14 (40.0) | NS |
| Influenza vaccine | 180 (35.3) | 18 (32.7) | NS |
| Pneumococcal vaccine | 52 (10.2) | 5 (9.1) | NS |
| Receiving non-combined LAMA | 420 (82.5) | 48 (87.3) | NS |
| Receiving non-combined tiotropium | 335 (65.8) | 40 (72.7) | NS |
| Receiving non-combined acclidinium | 62 (12.2) | 6 (10.9) | NS |
| Receiving non-combined glycopyrronium | 31 (6.1) | 2 (3.6) | NS |
| Receiving non-combined LABA | 107 (21.0) | 14 (25.5) | NS |
| Receiving non-combined salmeterol | 1 (0.2) | 1 (1.8) | NS |
| Receiving non-combined formoterol | 2 (0.4) | 1 (1.8) | NS |
| Receiving non-combined indacaterol | 105 (20.7) | 12 (21.8) | NS |
| Receiving non-combined LABA + LAMA | 94 (18.4) | 13 (23.6) | NS |
| Receiving non-combined ICS | 35 (6.9) | 6 (10.9) | NS |
| Receiving combined ICS-LABA | 314 (61.6) | 43 (78.2) | 0.018 |
| Receiving combined salmeterol-fluticasone | 188 (36.9) | 30 (54.5) | 0.013 |
| Receiving combined formoterol-budesonide | 101 (19.8) | 10 (18.2) | NS |
| Receiving combined formoterol-beclomethasone | 26 (5.1) | 3 (5.5) | NS |
| Receiving any form of ICS | 345 (67.6) | 49 (89.1) | 0.001 |
| Receiving low dose of ICS | 39 (7.6) | 5 (9.1) | NS |
| Receiving medium dose of ICS | 153 (30.0) | 30 (54.5) | 0.027 |
| Receiving high dose of ICS | 131 (25.7) | 11 (20.0) | 0.035 |
| Receiving triple therapy with LAMA+FDC | 263 (51.6) | 36 (65.5) | 0.064 |
| Receiving roflumilast | 57 (11.2) | 6 (10.9) | NS |
| Receiving mucolytics | 37 (7.3) | 8 (14.5) | 0.068 |
| Receiving antibiotics | 9 (1.8) | 4 (7.3) | 0.030 |

| | No step-down (n = 510) | Step-down (n = 55) | P-Value* |
|---|---------------------------|-----------------------|----------|
| Receiving oral methylxanthines | 16 (3.1) | 8 (14.5) | 0.001 |
| Receiving long-term oxygen therapy | 103 (20.2) | 3 (5.5) | 0.006 |
| Receiving home mechanical ventilation | 26 (5.1) | 0 (0) | 0.097 |
| Receiving nebulized home therapy | 19 (3.7) | 2 (3.6) | NS |
| Treatment adherence evaluation | 284 (55.7) | 36 (65.5) | NS |
| Inhaler device satisfaction evaluation | 88 (17.3) | 13 (23.6) | NS |
| Adverse effects recorded | 123 (24.1) | 17 (30.9) | NS |
| FVC (%) | 74.9 (20.9) | 78.2 (20.7) | NS |
| FEV ₁ (%) | 51.7 (19.2) | 60.2 (22.3) | 0.057 |
| FEV ₁ increase after the bronchodilator test (%) | 7.7 (11.1) | 7.4 (7.5) | NS |

Average value expressed as mean (standard deviation) or absolute (relative) frequencies depending on the nature of the variable.

*Calculated between groups using an unpaired Student t-test or a Chi-square test, depending on the nature of the variable.

LAMA: long-acting muscarinic antagonists. LABA: long-acting β_2 agonist. ICS: inhaled corticosteroids. FDC: fixed-dose combinations of an ICS and a LABA. FVC: forced vital capacity. FEV₁: forced expiratory volume in one second.

Table 3S. Bivariate analysis indicating factors associated with discontinuing inhaled steroids.

| | No ICS withdrawal (n = 355) | ICS withdrawal (n = 39) | |
|---|-----------------------------------|-------------------------------|---------|
| Monographic COPD clinic | 146 (41.1) | 15 (38.9) | NS |
| Outpatient clinic located within the hospital | 348 (98.0) | 38 (97.4) | NS |
| Male gender | 301 (84.8) | 32 (82.1) | NS |
| Age (years) | 69.2 (9.7) | 67.3 (8.6) | NS |
| Rural area | 117 (33.0) | 11 (28.2) | NS |
| Current smoker | 82 (23.1) | 9 (23.1) | NS |
| Tobacco history (pack-year) | 55.1 (31.8) | 61.6 (41.3) | NS |
| Diabetes | 78 (22.0) | 3 (7.7) | 0.037 |
| Depression | 33 (9.3) | 8 (20.5) | 0.047 |
| Anxiety | 33 (9.3) | 7 (17.9) | 0.096 |
| Charlson | 2.2 (1.6) | 2.0 (1.7) | NS |
| GesEPOC phenotype A | 132 (37.2) | 21 (53.8) | 0.013 |
| GesEPOC phenotype B | 49 (13.8) | 2 (5.1) | NS |
| GesEPOC phenotype C | 31 (8.7) | 1 (2.6) | NS |
| GesEPOC phenotype D | 70 (19.7) | 4 (10.3) | NS |
| GOLD type A | 36 (54.4) | 11 (28.2) | < 0.001 |
| GOLD type B | 37 (10.4) | 2 (5.1) | NS |
| GOLD type C | 67 (18.9) | 6 (15.4) | NS |
| GOLD type D | 89 (25.1) | 3 (7.7) | 0.020 |
| Dyspnea (mMRC) | 1.8 (0.9) | 1.4 (0.9) | 0.032 |
| Chronic bronchitis | 186 (52.4) | 16 (41.0) | NS |

| | No ICS withdrawal (n = 355) | ICS withdrawal (n = 39) | |
|--|-----------------------------------|-------------------------------|-------|
| Purulent sputum | 36 (10.1) | 1 (2.6) | NS |
| Asthma-like symptoms | 75 (21.1) | 3 (7.7) | 0.011 |
| Exacerbations previous year | 1.1 (1.4) | 0.6 (0.8) | 0.035 |
| Hospitalizations previous year | 0.3 (0.6) | 0.3 (0.7) | NS |
| Performs exercise | 83 (23.4) | 10 (25.6) | NS |
| Influenza vaccine | 137 (38.6) | 15 (38.5) | NS |
| Pneumococcal vaccine | 44 (12.4) | 2 (5.1) | 0.060 |
| Receiving non-combined LAMA | 301 (84.8) | 28 (71.8) | 0.065 |
| Receiving non-combined Tiotropium | 250 (70.4) | 23 (59.0) | NS |
| Receiving non-combined aclidinium | 40 (11.3) | 5 (12.8) | NS |
| Receiving non-combined glycopyrronium | 14 (3.9) | 0 (0) | NS |
| Receiving non-combined LABA | 25 (7.0) | 7 (17.9) | 0.028 |
| Receiving non-combined salmeterol | 2 (0.6) | 0 (0) | NS |
| Receiving non-combined formoterol | 0 (0) | 1 (2.6) | 0.099 |
| Receiving non-combined indacaterol | 24 (6.8) | 6 (15.4) | NS |
| Receiving non-combined LABA + LAMA | 20 (5.6) | 6 (15.4) | 0.033 |
| Receiving non-combined ICS | 35 (9.9) | 6 (15.4) | NS |
| Receiving combined ICS-LABA | 324 (91.3) | 33 (84.6) | NS |
| Receiving combined salmeterol-fluticasone | 193 (54.4) | 25 (64.1) | NS |
| Receiving combined formoterol-budesonide | 105 (29.6) | 6 (15.4) | 0.063 |
| Receiving combined formoterol-beclomethasone | 27 (7.6) | 2 (5.1) | NS |
| Receiving low dose of ICS | 38 (10.7) | 6 (15.4) | NS |
| Receiving medium dose of ICS | 161 (45.4) | 22 (56.4) | NS |
| Receiving high dose of ICS | 133 (37.5) | 9 (23.1) | NS |

| | No ICS withdrawal (n = 355) | ICS withdrawal (n = 39) | |
|---|-----------------------------------|-------------------------------|-------|
| Receiving triple therapy with LAMA+FDC | 277 (78.0) | 22 (56.4) | 0.005 |
| Receiving roflumilast | 51 (14.4) | 2 (5.1) | NS |
| Receiving mucolytics | 30 (8.5) | 6 (15.4) | NS |
| Receiving antibiotics | 10 (2.8) | 1 (2.6) | NS |
| Receiving oral methylxanthines | 18 (5.1) | 1 (2.6) | NS |
| Receiving long-term oxygen therapy | 77 (21.7) | 1 (2.6) | 0.002 |
| Receiving home mechanical ventilation | 21 (5.9) | 0 (0) | NS |
| Receiving nebulized home therapy | 17 (4.8) | 1 (2.6) | NS |
| Treatment adherence evaluation | 200 (56.3) | 24 (61.5) | NS |
| Inhaler device satisfaction evaluation | 62 (17.5) | 6 (15.4) | NS |
| Adverse effects recorded | 97 (27.3) | 9 (23.1) | NS |
| FVC (%) | 71.8 (19.0) | 78.0 (21.2) | NS |
| FEV1 (%) | 47.6 (16.9) | 59.8 (22.2) | 0.016 |
| FEV1 increase after the bronchodilator test (%) | 7.8 (10.2) | 6.0 (6.7) | NS |

Average values are expressed as means (standard deviation) or absolute (relative) frequencies depending on the nature of the variable.

*Calculated between groups using an unpaired Student t-test or a Chi-square test, depending on the nature of the variable.

LAMA: long-acting muscarinic antagonists. LABA: long-acting β_2 agonist. ICS: inhaled corticosteroids. FDC: fixed-dose combinations of an ICS and a LABA. FVC: forced vital capacity. FEV₁: forced expiratory volume in one second.