

## **e-Appendix 1.**

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**e-Table 1. Univariable Associations with Health Outcomes**

| Outcome          | $\beta$ (95% CI)           | p-value |
|------------------|----------------------------|---------|
| Obesity exposure |                            |         |
| SGRQ Total Score |                            |         |
| Obese class I    | 2.42 (0.63, 4.21)          | 0.008   |
| Obese Class II   | 5.52 (2.97, 8.08)          | <0.001  |
| Obese Class III  | 9.54 (6.27, 12.80)         | <0.001  |
| SF-36 T score    |                            |         |
| Obese class I    | -1.00 (-2.34, 0.35)        | 0.147   |
| Obese class II   | -2.66 (-4.61, -0.71)       | 0.008   |
| Obese class III  | -4.07 (-6.49, -1.66)       | 0.001   |
| 6MWD             |                            |         |
| Obese class I    | -52.24 (-84.70, -19.78)    | 0.002   |
| Obese class II   | -125.64 (-172.05, -79.23)  | <0.001  |
| Obese class III  | -305.06 (-364.74, -245.38) | <0.001  |
|                  | OR (95% CI)                | p-value |
| mMRC $\geq$ 2    |                            |         |
| Obese class I    | 1.19 (1.00, 1.41)          | 0.053   |
| Obese class II   | 1.80 (1.37, 2.35)          | <0.001  |
| Obese class III  | 3.42 (2.26, 5.19)          | <0.001  |
| Moderate AECOPD  |                            |         |
| Obese class I    | 1.27 (1.00, 1.60)          | 0.047   |
| Obese class II   | 1.20 (0.86, 1.68)          | 0.284   |
| Obese class III  | 1.06 (0.68, 1.65)          | 0.806   |
| Severe AECOPD    |                            |         |
| Obese class I    | 1.04 (0.85, 1.27)          | 0.685   |
| Obese class II   | 1.17 (0.88, 1.54)          | 0.276   |
| Obese class III  | 1.57 (1.13, 2.18)          | 0.007   |

Abbreviations:  $\beta$ , beta coefficient; 6MWD, 6-Minute Walk Distance; AECOPD, Acute Exacerbation of COPD; CI, confidence interval; mMRC, modified Medical Research Council; OR, odds ratio; SF-36, Short Form 35; SGRQ, Saint George's Respiratory Questionnaire

**e-Table 2. Health Outcome Multivariable Models with Individual Comorbidities**

| Exposure of Interest        | SGRQ Total Score    |         | SF-36 T score        |         | 6MWD                       |         |
|-----------------------------|---------------------|---------|----------------------|---------|----------------------------|---------|
|                             | $\beta$ (95% CI)    | p-value | $\beta$ (95% CI)     | p-value | $\beta$ (95% CI)           | p-value |
| Obesity                     |                     |         |                      |         |                            |         |
| Obese class I               | 2.04 (0.54, 3.54)   | 0.008   | -0.76 (-1.96, 0.43)  | 0.212   | -54.57 (-82.57, -26.56)    | <0.001  |
| Obese Class II              | 3.03 (0.84, 5.22)   | 0.007   | -0.86 (-2.66, 0.94)  | 0.349   | -123.17 (-164.08, -82.27)  | <0.001  |
| Obese Class III             | 5.40 (2.57, 8.22)   | <0.001  | -0.93 (-3.16, 1.31)  | 0.417   | -286.55 (-339.77, -233.34) | <0.001  |
| Allergies                   | 0.74 (-0.65, 2.13)  | 0.297   | -0.94 (-2.07, 0.18)  | 0.101   | 5.36 (-20.66, 31.39)       | 0.686   |
| Asthma                      | 6.60 (5.16, 8.04)   | <0.001  | -3.27 (-4.43, -2.12) | <0.001  | 15.31 (-11.60, 42.22)      | 0.265   |
| Cancer                      | 1.79 (-0.56, 4.15)  | 0.135   | -1.71 (-3.64, 0.23)  | 0.084   | -11.41 (-55.68, 32.85)     | 0.613   |
| Coronary Heart Disease      | 1.88 (0.18, 3.58)   | 0.030   | -1.86 (-3.21, -0.52) | 0.007   | -64.99 (-96.83, -33.14)    | <0.001  |
| Congestive Heart Failure    | 1.44 (-1.23, 4.11)  | 0.291   | -0.11 (-2.20, 1.97)  | 0.914   | -89.18 (-140.19, -38.16)   | 0.001   |
| CVA / TIA                   | 4.79 (2.29, 7.28)   | <0.001  | -2.01 (-3.94, -0.09) | 0.040   | -28.87 (-75.47, 17.73)     | 0.225   |
| Diabetes mellitus           | 1.18 (-0.62, 2.99)  | 0.198   | -1.64 (-3.06, -0.21) | 0.024   | -37.68 (-71.57, -3.79)     | 0.029   |
| GERD                        | 3.61 (2.27, 4.96)   | <0.001  | -1.71 (-2.78, -0.63) | 0.002   | -23.44 (-48.67, 1.79)      | 0.069   |
| Hypertension                | 0.82 (-0.44, 2.09)  | 0.201   | 0.52 (-0.51, 1.56)   | 0.319   | -17.00 (-40.60, 6.61)      | 0.158   |
| Hyperlipidemia              | -0.15 (-1.44, 1.14) | 0.820   | -0.56 (-1.60, 0.47)  | 0.285   | 32.58 (8.39, 56.77)        | 0.008   |
| Obstructive Sleep Apnea     | 5.84 (4.21, 7.47)   | <0.001  | -2.94 (-4.28, -1.61) | <0.001  | -46.39 (-77.00, -15.78)    | 0.003   |
| Osteoarthritis              | 1.45 (-0.06, 2.96)  | 0.060   | -0.03 (-1.23, 1.17)  | 0.965   | -29.00 (-57.22, -0.77)     | 0.044   |
| Osteoporosis                | 2.86 (1.24, 4.48)   | 0.001   | -2.66 (-3.99, -1.32) | <0.001  | -44.34 (-74.82, -13.87)    | 0.004   |
| Peptic ulcer disease        | 1.38 (-0.59, 3.34)  | 0.171   | -1.61 (-3.19, -0.04) | 0.045   | 0.52 (-36.53, 37.57)       | 0.978   |
| Peripheral arterial disease | 0.14 (-3.13, 3.40)  | 0.935   | -1.04 (-3.63, 1.56)  | 0.435   | -82.78 (-143.52, -22.02)   | 0.008   |
| Rheumatoid arthritis        | 5.28 (3.19, 7.36)   | <0.001  | -1.45 (-3.13, 0.23)  | 0.090   | -40.20 (-79.17, -1.24)     | 0.043   |
| Exposure of Interest        | mMRC                |         | Moderate AECOPD      |         | Severe AECOPD              |         |
|                             | OR (95% CI)         | p-value | OR (95% CI)          | p-value | OR (95% CI)                | p-value |

|                             |      |              |        |      |              |        |      |              |        |
|-----------------------------|------|--------------|--------|------|--------------|--------|------|--------------|--------|
| Obesity                     |      |              |        |      |              |        |      |              |        |
| Obese class I               | 1.22 | (0.99, 1.50) | 0.058  | 1.22 | (0.95, 1.57) | 0.124  | 1.04 | (0.84, 1.29) | 0.725  |
| Obese class II              | 1.67 | (1.21, 2.30) | 0.002  | 1.06 | (0.73, 1.52) | 0.771  | 1.10 | (0.81, 1.50) | 0.550  |
| Obese class III             | 2.92 | (1.82, 4.69) | <0.001 | 0.79 | (0.48, 1.28) | 0.337  | 1.42 | (0.97, 2.08) | 0.069  |
| Allergies                   | 0.91 | (0.75, 1.11) | 0.349  | 1.28 | (1.2, 1.60)  | 0.034  | 1.10 | (0.91, 1.34) | 0.317  |
| Asthma                      | 1.55 | (1.25, 1.92) | <0.001 | 1.56 | (1.24, 1.95) | <0.001 | 1.72 | (1.42, 2.08) | <0.001 |
| Cancer                      | 0.95 | (0.69, 1.31) | 0.758  | 0.93 | (0.61, 1.41) | 0.723  | 1.25 | (0.89, 1.76) | 0.193  |
| Coronary Heart Disease      | 1.42 | (1.11, 1.82) | 0.005  | 1.09 | (0.82, 1.46) | 0.537  | 1.03 | (0.81, 1.32) | 0.798  |
| Congestive Heart Failure    | 1.66 | (1.04, 2.66) | 0.035  | 0.92 | (0.60, 1.40) | 0.699  | 1.45 | (1.02, 2.04) | 0.036  |
| CVA / TIA                   | 1.40 | (0.98, 2.02) | 0.067  | 1.57 | (1.06, 2.32) | 0.025  | 1.52 | (1.08, 2.14) | 0.017  |
| Diabetes mellitus           | 1.07 | (0.82, 1.40) | 0.609  | 1.42 | (1.07, 1.89) | 0.016  | 1.15 | (0.89, 1.48) | 0.276  |
| GERD                        | 1.39 | (1.15, 1.69) | 0.001  | 1.17 | (0.93, 1.46) | 0.179  | 1.17 | (0.97, 1.42) | 0.103  |
| Hypertension                | 1.12 | (0.94, 1.34) | 0.199  | 1.01 | (0.81, 1.26) | 0.907  | 0.93 | (0.77, 1.11) | 0.410  |
| Hyperlipidemia              | 1.08 | (0.90, 1.29) | 0.430  | 0.92 | (0.74, 1.14) | 0.447  | 1.03 | (0.85, 1.24) | 0.750  |
| Obstructive Sleep Apnea     | 1.56 | (1.22, 1.98) | <0.001 | 1.47 | (1.14, 1.91) | 0.003  | 1.11 | (0.88, 1.40) | 0.370  |
| Osteoarthritis              | 1.17 | (0.94, 1.45) | 0.164  | 0.98 | (0.76, 1.26) | 0.861  | 0.82 | (0.66, 1.03) | 0.090  |
| Osteoporosis                | 1.40 | (1.10, 1.77) | 0.006  | 1.45 | (1.12, 1.87) | 0.005  | 1.25 | (0.99, 1.57) | 0.056  |
| Peptic ulcer disease        | 1.23 | (0.92, 1.64) | 0.164  | 1.12 | (0.82, 1.54) | 0.471  | 1.06 | (0.80, 1.39) | 0.690  |
| Peripheral arterial disease | 1.10 | (0.68, 1.80) | 0.699  | 0.94 | (0.54, 1.20) | 0.829  | 0.90 | (0.55, 1.45) | 0.652  |
| Rheumatoid arthritis        | 1.40 | (1.03, 1.90) | 0.033  | 0.83 | (0.57, 1.20) | 0.315  | 1.34 | (1.01, 1.77) | 0.042  |

Adjusted for: age, gender, race, education, FEV<sub>1</sub>/Ht<sup>2</sup>, smoking status, and smoking pack-years.

Referent for obesity class: Normal/overweight individuals (BMI 18.5 – 29.9kg/m<sup>2</sup>)

Abbreviations: β, beta coefficient; 6MWD, 6-Minute Walk Distance; AECOPD, Acute Exacerbation of COPD; CI, confidence interval; CVA, cerebrovascular attack; GERD, gastroesophageal reflux disease; mMRC, modified Medical Research Council; OR, odds ratio; SF-36, Short Form 35; SGRQ, Saint George's Respiratory Questionnaire; TIA, transient ischemic attack