Correlations of three polymorphisms in  $\beta_2$ -adrenergic receptor gene with chronic obstructive pulmonary disease risk and related phenotypes: a meta-analysis

| pplementary Table SI. The PRISMA checklist for the present meta-analysis |
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|--|

| Section/topic                         | #                                     | Reported<br>on page #   |     |  |  |  |
|---------------------------------------|---------------------------------------|---|-----|--|--|--|
| Fitle                                 |                                       |   |     |  |  |  |
| Title                                 | 1                                     | Identify the report as a systematic review, meta-analysis,<br>or both   | 1   |  |  |  |
| Abstract                              |                                       |   |     |  |  |  |
| Structured summary                    | 2                                     | Provide a structured summary including, as applicable:<br>background; objectives; data sources; study eligibility<br>criteria, participants, and interventions; study appraisal and<br>synthesis methods; results; limitations; conclusions and<br>implications of key findings; systematic review registration<br>number | 2   |  |  |  |
| ntroduction                           |                                       |   |     |  |  |  |
| Rationale                             | 3                                     | Describe the rationale for the review in the context of what is already known   | 3   |  |  |  |
| Objectives                            | 4                                     | Provide an explicit statement of questions being addressed<br>with reference to participants, interventions, comparisons,<br>outcomes, and study design (PICOS)   | 3–4 |  |  |  |
| Nethods                               |                                       |   |     |  |  |  |
| Protocol and registration             | 5                                     | Indicate whether a review protocol exists, if and where<br>it can be accessed (e.g., Web address), and, if available,<br>provide registration information including registration<br>number  | 4   |  |  |  |
| Eligibility criteria                  | 6                                     | Specify study characteristics (e.g., PICOS, length of follow-<br>up) and report characteristics (e.g., years considered,<br>language, publication status) used as criteria for eligibility,<br>giving a rationale   | 4–5 |  |  |  |
| Information sources                   | 7                                     | Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched   | 4–5 |  |  |  |
| Search                                | 8                                     | Present full electronic search strategy for at least one<br>database, including any limits used, such that it could be<br>repeated  | 4   |  |  |  |
| Study selection                       | 9                                     | State the process for selecting studies (i.e., screening,<br>eligibility, included in systematic review, and, if applicable,<br>included in the meta-analysis)  | 4–6 |  |  |  |
| Data collection process               | · · · · · · · · · · · · · · · · · · · |   | 6   |  |  |  |
| Data items                            | 11                                    | List and define all variables for which data were sought<br>(e.g., PICOS, funding sources) and any assumptions and<br>simplifications made  | 6–7 |  |  |  |
| Risk of bias in individual<br>studies | 12                                    | Describe methods used for assessing risk of bias of<br>individual studies (including specification of whether this<br>was done at the study or outcome level), and how this<br>information is to be used in any data synthesis  |     |  |  |  |
| Summary measures                      | 13                                    | State the principal summary measures (e.g., risk ratio,<br>difference in means)   | 6–7 |  |  |  |
| Synthesis of results                  | 14                                    | Describe the methods of handling data and combining<br>results of studies, if done, including measures of<br>consistency (e.g., I <sup>2</sup> ) for each meta-analysis   | 6–7 |  |  |  |

| Supplementary | Table | SI. | Cont. |
|---------------|-------|-----|-------|
|---------------|-------|-----|-------|

| Section/topic                    | #  | Checklist item   | Reported on page #  |  |  |
|----------------------------------|--|--|---|--|--|
| Risk of bias across studies      | of bias across studies 15 Specify any assessment of risk of bias that may affect<br>the cumulative evidence (e.g., publication bias, selective<br>reporting within studies)                              |  | 6–7   |  |  |
| Additional analyses              | 16 Describe methods of additional analyses (e.g., sensitivity<br>or subgroup analyses, meta-regression), if done, indicating<br>which were pre-specified   |  | 6–7   |  |  |
| Results                          |  |  |   |  |  |
| Study selection                  | 17   | Give numbers of studies screened, assessed for eligibility,<br>and included in the review, with reasons for exclusions at<br>each stage, ideally with a flow diagram   | Figure 1  |  |  |
| Study characteristics            | 18   | For each study, present characteristics for which data were<br>extracted (e.g., study size, PICOS, follow-up period) and<br>provide the citations  | 7; 14–15<br>Table I   |  |  |
| Risk of bias within<br>studies   | 19   | Present data on risk of bias of each study and, if available,<br>any outcome level assessment (see item 12)  | Figure 3  |  |  |
| Results of individual<br>studies | 20   | For all outcomes considered (benefits or harms), present,<br>for each study: (a) simple summary data for each<br>intervention group (b) effect estimates and confidence<br>intervals, ideally with a forest plot | Figure 2  |  |  |
| Synthesis of results             | 21   | Present results of each meta-analysis done, including confidence intervals and measures of consistency   | 7–9   |  |  |
| Risk of bias across<br>studies   | 22   | Present results of any assessment of risk of bias across<br>studies (see Item 15)  | Figure 4  |  |  |
| Additional analysis              |  |  | 7–9;16–20<br>Supplementary<br>Figure S1,<br>Table II,<br>Table III,<br>Figure 4 |  |  |
| Discussion                       |  |  |   |  |  |
| Summary of evidence              | evidence 24 Summarize the main findings including the strength of<br>evidence for each main outcome; consider their relevance<br>to key groups (e.g., healthcare providers, users, and policy<br>makers) |  | 9–11  |  |  |
| Limitations                      | 25   | Discuss limitations at study and outcome level (e.g., risk<br>of bias), and at review level (e.g., incomplete retrieval of<br>identified research, reporting bias)   | 11  |  |  |
| Conclusions                      | 26   | Provide a general interpretation of the results in the context of other evidence, and implications for future research   | 11  |  |  |
| Funding                          |  |  |   |  |  |
| Funding                          | 27   | Describe sources of funding for the systematic review and<br>other support (e.g., supply of data); role of funders for the<br>systematic review  | 12  |  |  |

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| First author<br>(year)                        | Represen-<br>tative-<br>ness of<br>cases | Represen-<br>tative-<br>ness of<br>controls |   |   | Genotyp-<br>ing exam-<br>ination |   | Asso-<br>ciation<br>assess-<br>ment | Response<br>rate | Total |
|---|--|---|---|---|----------------------------------|---|-------------------------------------|------------------|-------|
| Ho LI (2001)                                  | 2  | 2   | 2 | 1 | 0                                | 0 | 1                                   | 2                | 10    |
| Hegab AE (2004)<br>(Japan)                    | 2  | 2   | 2 | 0 | 0                                | 2 | 1                                   | 2                | 11    |
| Hegab AE (2004)<br>(Egypt)                    | 2  | 2   | 2 | 0 | 0                                | 2 | 1                                   | 2                | 11    |
| Brogger J (2006)                              | 2  | 2   | 2 | 2 | 0                                | 2 | 2                                   | 0                | 12    |
| Matheson MC<br>(2006)                         | 2  | 2   | 2 | 2 | 1                                | 2 | 2                                   | 2                | 15    |
| Ferdinands JM<br>(2007)<br>(American-African) | 2  | 2   | 2 | 2 | 1                                | 2 | 2                                   | 2                | 15    |
| Ferdinands JM<br>(2007)<br>(Caucasian)        | 2  | 2   | 2 | 2 | 1                                | 2 | 2                                   | 2                | 15    |
| Vacca G (2009)                                | 2  | 2   | 2 | 2 | 0                                | 2 | 1                                   | 2                | 13    |
| Papatheodorou A<br>(2010)                     | 2  | 2   | 2 | 1 | 0                                | 2 | 1                                   | 2                | 12    |
| Thomsen M<br>(2012)                           | 2  | 2   | 2 | 2 | 1                                | 2 | 2                                   | 2                | 15    |
| Ganbold C (2016)                              | 2  | 2   | 2 | 1 | 0                                | 2 | 2                                   | 2                | 13    |
| Hussein MH<br>(2017)                          | 2  | 2   | 2 | 1 | 0                                | 2 | 2                                   | 2                | 13    |
| Zhao H (2017)                                 | 2  | 2   | 2 | 2 | 0                                | 2 | 1                                   | 2                | 13    |
| Li JX (2018)                                  | 2  | 2   | 2 | 2 | 0                                | 2 | 2                                   | 2                | 14    |

Supplementary Table SII. Quality scores of all studies included in the present meta-analysis

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**Supplementary Figure S1.** Cumulative and influential plots of three polymorphisms in *ADRB2* gene associated with the risk of chronic obstructive pulmonary disease

 $\label{eq:correlations} \mbox{ of three polymorphisms in $$$$$$$$$$$$$$$$$_2$-adrenergic receptor gene with chronic obstructive pulmonary disease risk and related phenotypes: a meta-analysis a meta-analys$ 









Supplementary Figure S1. Cont.

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Supplementary Figure S1. Cont.