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1 **SEARCH STRATEGY**

2 The full content of each string is reported below. PubMed database was searched, by inserting one

3 string at the time on the research bar. No limitations or filters were set during our PubMed search.

4 No backward time limitation was set, we included studies published up to September 10th, 2023.

5

6 PubMed Strings:

7 • *Riociguat*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND

8 "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All

9 Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields]))

10 AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND

11 "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All

12 Fields] AND "hypertension"[All Fields])) AND ("riociguat"[Supplementary Concept] OR

13 "riociguat"[All Fields]).

14 • *Sildenafil*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND

15 "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All

16 Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields]))

17 AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND

18 "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All

19 Fields] AND "hypertension"[All Fields])) AND ("sildenafil citrate"[MeSH Terms] OR

20 ("sildenafil"[All Fields] AND "citrate"[All Fields]) OR "sildenafil citrate"[All Fields] OR

21 "sildenafil"[All Fields] OR "sildenafil s"[All Fields])

22 • *Tadalafil*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND

23 "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All

24 Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields]))

25 AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND

26 "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All

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Fields] AND "hypertension"[All Fields])) AND ("tadalafil"[MeSH Terms] OR "tadalafil"[All Fields])

- *Bosentan*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields])) AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All Fields] AND "hypertension"[All Fields])) AND ("bosentan"[MeSH Terms] OR "bosentan"[All Fields])

- *Macitentan*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields])) AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All Fields] AND "hypertension"[All Fields])) AND ("macitentan"[Supplementary Concept] OR "macitentan"[All Fields])

- *Ambrisentan*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields])) AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All Fields] AND "hypertension"[All Fields])) AND ("ambrisentan"[Supplementary Concept] OR "ambrisentan"[All Fields])

- *Treprostinil*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields]))

1. AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND
2. "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All
3. Fields] AND "hypertension"[All Fields])) AND ("treprostinil"[Supplementary Concept] OR
4. "treprostinil"[All Fields])

5. • *Iloprost*: ("lung diseases, interstitial"[MeSH Terms] OR ("lung"[All Fields] AND
6. "diseases"[All Fields] AND "interstitial"[All Fields]) OR "interstitial lung diseases"[All
7. Fields] OR ("interstitial"[All Fields] AND "lung"[All Fields] AND "diseases"[All Fields]))
8. AND ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND
9. "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All
10. Fields] AND "hypertension"[All Fields])) AND ("iloprost"[MeSH Terms] OR "iloprost"[All
11. Fields])

12. • *Selexipag*: ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND
13. "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All
14. Fields] AND "hypertension"[All Fields])) AND ("selexipag"[Supplementary Concept] OR
15. "selexipag"[All Fields])

16. • *Isosorbide*: ("hypertension, pulmonary"[MeSH Terms] OR ("hypertension"[All Fields] AND
17. "pulmonary"[All Fields]) OR "pulmonary hypertension"[All Fields] OR ("pulmonary"[All
18. Fields] AND "hypertension"[All Fields])) AND ("isosorbide"[All Fields] OR
19. "isosorbide"[MeSH Terms] OR "isosorbide"[All Fields])

20.

21. ClinicalTrials.gov was also employed to search for clinical trials. The words of [*vasoactive drug*] *(*in
22. eg., riociguat) and "*pulmonary hypertension*" were inserted in the research string panel for each
23. vasoactive drug. The resulting clinical trials were then considered for screening if their status was
24. addressed as "completed with results". No backward time limitation was set, we included studies
25. published up to September 10th, 2023.

26.

1   ClinicalTrial.gov string panel:

2   - *Riociguat* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

3   - *Sildenafil* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

4   - *Tadalafil* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

5   - *Bosentan* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

6   - *Macitentan* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

7   - *Ambrisentan* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

8   - *Treprostinil* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

9   - *Iloprost* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

10   - *Selexipag* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

11   - *Isosorbide* ("intervention/treatment") AND *pulmonary hypertension* ("condition/disease")

12

13

14