# INPLASY PROTOCOL

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Conflicts of interest: None declared.

### **INTRODUCTION**

**Review question / Objective:** To systematically evaluate the efficacy of apitherapy in the treatment of allergic rhinitis.

Condition being studied: Allergic rhinitis (AR) is an inflammatory, IgE mediated disease characterized by recurrent nasal congestion, runny nose, sneezing, and/or nasal itching. It is estimated to affect up to

## Systematic evaluation and Mate analysis of apitherapy for allergic rhinitis

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Review question / Objective: To systematically evaluate the efficacy of apitherapy in the treatment of allergic rhinitis. Condition being studied: Allergic rhinitis (AR) is an

inflammatory, IgE mediated disease characterized by recurrent nasal congestion, runny nose, sneezing, and/or nasal itching. It is estimated to affect up to 25% of children and 40% of adults. This has significant implications for the quality of life and economic burden on those affected, the health care system and society as a whole.

**INPLASY registration number:** This protocol was registered with the International Platform of Registered Systematic Review and Meta-Analysis Protocols (INPLASY) on 09 August 2021 and was last updated on 09 August 2021 (registration number INPLASY202180032).

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### **METHODS**

Search strategy: We will identify published and ongoing research by searching the following databases from the outset: CENTRAL Registry of Cochrane Controlled Trials, PubMed, EMBASE. Chinese databases: China National Knowledge Infrastructure Database (CNKI), Chinese Scientific Journals Database (VIP), Wanfang Database, Chinese Biomedical Literature Database.

Participant or population: Patients who have intermittent or persistent AR will be included, regardless of age, gender, race, or other demographical factors. The diagnosis of AR is not universally standardised, but is predominantly made clinically. We define AR based on history and physical exam consistent with an allergic cause and one or more of the following symptoms: nasal congestion, rhinorrhoea, sneezing or nasal itch. This may be supported with a positive skin prick test or the detection of high circulating blood levels of allergen-specific IgE antibody via a radioallergosorbent (RAST) test.

Intervention: Apitherapy involves a series of procedures that involve simulating body points

**Comparator:** Placebo, sham acupuncture, pharmacotherapy, other types of Chinese medicine therapies.

Study designs to be included: All relevant RCTs published in English and Chinese on needle-embedding for AR can be included. Non-RCTs, reviews, case report, experimental studies, expert experience, and duplicated publications will be excluded.

Eligibility criteria: Inclusion criteria: Study designs included case-control, crosssectional, or cohort studies.Exclusion criteria:1) For duplicate literatures, only literatures with good quality and complete information should be retained, and other literatures should be deleted.2) Unable to extract data for statistical analysis;3) Poor literature quality or statistical errors;4) review; Comments. Letters.

Information sources: We will search PubMed(MEDLINE), EMBASE, Cochrane Library, Web of Science, and National Library of China Cnki, VIP, The Chinese Academy of Biomedical Research and Wanfang will identify all published studies, regardless of language. (Deadline: August 9, 2021).

Main outcome(s): Symptom score will be assessed as the primary outcome. The main evaluation indicators of symptom score included nasal symptoms (sneezing, runny nose, nasal itching and nasal congestion) and ocular symptoms (eye itching, foreign body sensation, red eyes, tearing). If asthma is combined, additional asthma symptoms such as wheezing, cough, shortness of breath and chest tightness should be recorded.

Additional outcome(s): I. Rhinoconjunctivitis quality of life questionnaire (RQLQ), participants with asthma can use asthma control test (ACT); II. Medicine usage and scoring; III. Nasal function examination & nasal provocation test; IV. Allergen detection (e.g., skin prick test (SPT), total serum IgE detection, serum specific IgE detection will be included); V. Supplementary examination (e.g., nasal secretion smear, determination of specific IgE in nasal lavage fluid); VI. Side effects & adverse events.

Quality assessment / Risk of bias analysis: Each working group consists of three members. Two members will independently screen studies and extract relevant data according to predefined selection criteria. A third member will decide whether to include or exclude indeterminate studies and examine the data extraction table. For the selected study, the first author, publication date, title, source, region, sample size and the correlation coefficient between units and AR were extracted and entered into Microsoft Excel.

Strategy of data synthesis: Data will be analyzed using Excel and STATA13.1. When heterogeneity occurs across studies, if the p-value was <0.05, a random effect model would be selected; if not, a fixed-effects model.Summary effect sizes will be calculated as RR(or AR), with 95% credible intervals (CrIs).The Q and I<sup>2</sup> statistics will be used to test for heterogeneity between included studies.Sensitivity analysis was performed to investigate the influence of a single study on the overall meta-analysis estimate.

Subgroup analysis: Subgroup analyses were based on the different continent, country or district (depends on the actually extracted data).

Sensitivity analysis: To assess the influence of each individual study, leave-one-out sensitivity analysis was performed iteratively by removing one study at a time to confirm that the findings were not influenced by any single study.

Language: English.

Country(ies) involved: China.

Keywords: Allergic rhinitis, Rhinitis, Allergic; apitherapy.

#### Contributions of each author:

Author 1 - Yao Xiao - The author drafted and improved the manuscript. Email: 1399073323@qq.com Author 2 - Jun Xiong - Revise this protocol; search strategy. Email: xiongjun196071@163.com Author 3 - Yuntao Zeng - Data collection; analysis of results. Email: 1451911996@qq.com