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### INTRODUCTION

- GERD questionnaire [GerdQ] (2010) combines 3 questionnaires: • Reflux Disease Questionnaire [RDQ],
  - Gastrointestinal Symptom Rating Scale [GSRS], and
  - Gastro-oesophageal reflux disease Impact Scale [GIS])
- GerdQ is used as a **diagnostic tool** for GERD in primary care of many countries.
- The Asian guidelines for GERD diagnosis and treatment recommended using symptom-based diagnostic questionnaires such as **GerdQ** for diagnosing symptomatic GERD, despite the low level of evidence.
- The 2022 American College of Gastroenterology (ACG) guideline for GERD diagnosis recommended using the PPI test for establishing GERD diagnosis, which already has a moderate level of evidence.
- Aim: To summarize available evidence of the **diagnostic accuracy of** GerdQ compared to upper endoscopy and/or pH-metry for diagnosing GERD.

## METHODS

### • Inclusion criteria:

- Diagnostic Test Accuracy (DTA) studies comparing GerdQ to ambulatory pH-metry or upper endoscopy
- Adult patients presenting with symptoms suggestive of GERD who filled in the GerdQ and underwent gold standard test,
- Any GERD phenotypes (erosive or non-erosive reflux disease)

### • Exclusion criteria:

- Studies with patients presenting with GERD alarm symptoms, pregnant patients, patients who are breastfeeding or with malignancy, and patients with extraesophageal manifestations
- The Quality Assessment of Diagnostic Accuracy Studies-2 tool was used to assess the study quality

• Meta-analysis using the random-effects model (DerSimonian-Laird) was done to summarize the overall sensitivity, specificity, likelihood ratios (LRs), and diagnostic odds ratio (DOR).

• The hierarchical summary receiver operating characteristics (HSROC) curve was calculated using the bivariate model alongside the area under the HSROC (AUC).

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# Evaluating the Diagnostic Accuracy of GerdQ for Diagnosis of Gastroesophageal Reflux Disease: A Meta-Analysis

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Table 1. Characteristics of the Included Studies

| First Author                 | Country  | Study Design            | Period of Study       | Reference Test  | Sample<br>Size    | Patients<br>excluded from<br>analysis | Age (years)                  | Male (%)                           |
|------------------------------|--|-------------------------|-----------------------|---|-------------------|---------------------------------------|------------------------------|------------------------------------|
| Jones R (2009)               | Germany,<br>Sweden,<br>Canada,<br>Denmark,<br>Norway, UK | Non-randomized<br>trial | Sep 2005 - Nov 2006   | Upper endoscopy,<br>48-h wireless esophageal<br>pH monitoring | 308               | 5                                     | 47±14                        | 143 (46)                           |
| Lacy BE (2011)               | USA  | Cross-Sectional         | Oct 2008 - Oct 2009   | Wireless 48-h pH-metry  | 358 <sup>\$</sup> | 6#                                    | 51±14                        | 101 (28)                           |
| Bai Y (2013)                 | China  | Cross-Sectional         | Jan 2010 - Dec 2010   | Upper endoscopy   | 8065              | 0                                     | 46±14                        | 4043 (50)                          |
| Jonasson C<br>(2013)         | Norway   | Cross-Sectional         | Mar 2009 - Dec 2009   | Upper endoscopy, 24- h<br>pH-metry                            | 169               | 0                                     | 47±15                        | 90 (53)                            |
| Wang W (2014)                | China  | Cross-Sectional         | Feb 2011 - Dec 2011   | 24-h pH-metry   | 95                | 13                                    | 49±12                        | 44 (54)                            |
| Zavala-Gonzales<br>MA (2014) | Mexico   | Cross-Sectional         | Jan 2011 - Oct 2012   | Upper endoscopy, pH-<br>metry                                 | 252               | 0                                     | 49.49                        | 93 (37)                            |
| Zhou LY (2014)               | China  | Non-randomized<br>trial | Sep 2011 - Dec 2012   | Upper endoscopy,<br>24-h impedance pH<br>monitoring           | 636               | 0                                     | 49±14                        | 265 (42)                           |
| Siregar GA<br>(2015)         | Indonesia  | Cross-Sectional         | Oct 2015 - Dec 2015   | Upper endoscopy   | 85                | 0                                     | 45±13                        | 50 (59)                            |
| Wang M (2017)                | China  | Cross-sectional         | Aug 2014-<br>Dec 2015 | Upper endoscopy   | 1233              | 0                                     | 54±12                        | 532 (43)                           |
| Norder Grusell E<br>(2018)   | Sweden   | Cross-Sectional         | Oct 2009 - Apr 2014   | 24-h pH-metry   | 646               | 0                                     | 52 (15-84)                   | 296 (46)                           |
| Vadivelu S<br>(2019)         | Malaysia   | Cross-Sectional         | NR                    | Upper endoscopy, pH-<br>metry                                 | 104               | 0                                     | 47±1                         | 50 (48)                            |
| Zaika S (2020)               | Ukraine  | Cross-Sectional         | NR                    | 24-h impedance<br>pH monitoring                               | 28                | 0                                     | 47±2                         | 11 (39)                            |
| Chen G (2021)                | China  | Cross-Sectional         | Jul 2018 - Aug 2018   | Upper endoscopy   | 565               | 54                                    | NR                           | 290 (51)                           |
| Wang B (2021)                | China  | Cross-sectional         | Jun 2016 - Jun 2019   | Upper endoscopy   | 100               | 0                                     | GERD: 45±4<br>Non-GERD: 45±4 | GERD: 32 (52)<br>Non-GERD: 19 (50) |

GERD: Gastroesophageal Reflux Disease; NR: Not reported

Age presented in mean±SD / median (range) <sup>\$</sup>178 participants were off acid suppression, and 180 participants were on acid suppression

\*1 participant were off acid suppression, and 5 participants were on acid suppression

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| DISCUSSION  |
|---|
| <ul> <li>14 studies with a total of 12566 subjects were included, of which<br/>3564 subjects had confirmed GERD diagnosis.</li> </ul>   |
| <ul> <li>The overall pooled sensitivity, specificity, and AUC of GerdQ was 67.8%, 66.6%, and 0.705, respectively</li> <li>GerdQ with a cut-off value of ≥8 = slightly modest diagnostic value for GERD</li> <li>GerdQ had better specificity (70.3%) in ERD diagnosis</li> <li>GerdQ was suboptimal in establishing NERD diagnosis (sensitivity of 38% and specificity of 57%)</li> </ul> |
| <ul> <li>Despite better sensitivity, specificity, and AUC of GerdQ in GERD<br/>compared to previous meta-analysis, GerdQ was still insufficient to<br/>be used as the only screening or diagnostic tool for GERD</li> </ul>   |
| <ul> <li>While PPI test was shown to have moderate-to-high sensitivity (79%)<br/>for GERD, a lower specificity (45%) was expected due to the high<br/>placebo effect</li> </ul>   |
| <ul> <li>Upper endoscopy and esophageal pH/pH-impedance monitoring<br/>are known to be more sensitive and specific</li> <li>However, such reference tests are limited by the level of<br/>invasiveness and need for referral for primary care patients</li> </ul>   |
| <ul> <li>Some studies recommended increasing the GerdQ threshold to improve its diagnostic credibility in predicting GERD</li> <li>Only 1 study by Wang et al. used a GerdQ cut-off value &gt;9 with a resulting in sensitivity and specificity of 87.7% and 65.8%</li> </ul>   |
| <ul> <li>Strengths of this meta-analysis:</li> <li>Extensive and up-to-date search of studies that identified an additional 8 articles not included in a previous meta-analysis</li> </ul>  |
| <ul> <li>Sensitivity analysis confirmed the robustness of our study findings</li> </ul>   |
| <ul> <li>Limitations of this meta-analysis:</li> <li>Only studies published in English were included</li> </ul>   |
| • <b>Residual confounders (i.e PPI use)</b> cannot be entirely excluded due to different exclusion criteria definitions used in the included studies  |
| CONCLUSION  |
| <ul> <li>This meta-analysis demonstrated that:</li> <li>GerdQ had slightly moderate sensitivity and specificity for confirming GERD diagnosis, especially for the ERD phenotype</li> </ul>  |
| <ul> <li>GerdQ may still be considered a GERD diagnostic tool in resource-<br/>limited settings which lacks accepted reference tests and when<br/>PPI test is unavailable or contraindicated</li> </ul>   |