INPLASY PROTOCOL

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Conflicts of interest:

None declared.

INTRODUCTION

Review question / Objective: To determine the utility of CEUS in the preoperative differentiation between pleomorphic

Contrast-enhanced ultrasound in the differentiation between the most common benign parotid gland tumors: a systematic review and meta-analysis

Rogalska, M1; Antkowiak, L2; Kasperczuk, A3; Scierski, W4.

Review question / Objective: To determine the utility of CEUS in the preoperative differentiation between pleomorphic adenomas (PAs) and Warthin's tumors (WTs) of the parotid gland.

Condition being studied: Recently, contrast-enhanced ultrasound (CEUS) has become a promising tool in distinguishing benign from malignant parotid gland tumors. However, its usefulness in differentiating various benign parotid tumors has not been determined so far. The preoperative differentiation of the most common parotid gland tumors (PAs and WTs) remains crucial to select an appropriate management strategy. Therefore, the purpose of the present study was to systematically review the literature to determine the utility of CEUS in the differentiation of PAs and WTs of the parotid gland.

INPLASY registration number: This protocol was registered with the International Platform of Registered Systematic Review and Meta-Analysis Protocols (INPLASY) on 10 December 2022 and was last updated on 10 December 2022 (registration number INPLASY2022120042).

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METHODS

Participant or population: Patients with PAs or WTs of the parotid gland, who underwent preoperative CEUS imaging.

Intervention: The evaluation of PAs and WTs by means of CEUS (number of assessed benign parotid gland tumors [PAs/WTs], reference standard (histopathology), contrast agent (Sonovue), time considered for analysis following contrast administration, the region of interest (ROI) selection, tumor characteristics on CEUS images, and the assessed CEUS parameters).

Comparator: Not applicable.

Study designs to be included: Nonrandomized prospective and retrospective studies.

Eligibility criteria: (1) Additional inclusion criteria: English-language full-text papers evaluating differential diagnosis of benign parotid tumors using CEUS. (2) Exclusion criteria: publications with an unrelated topic, conference papers, review articles, case reports, commentaries, technical notes, and letters to the editor.

Information sources: The PubMed, Embase, and Cochrane databases were searched by two authors independently for English-language full-text papers published from inception until July 21, 2022. Additionally, the reference lists in all preselected articles were screened for further relevant papers. Any discrepancies

between the researchers were discussed, until a consensus was reached.

Main outcome(s): To determine the presence of qualitative and quantitative CEUS characteristics, which significantly differ between PAs and WTs of the parotid gland.

Quality assessment / Risk of bias analysis: Two reviewers independently assessed the quality of the eligible studies (regarding their applicability and risk of bias) according to the Quality Assessment of Diagnostic Accuracy Studies (QUADAS-2)

according to the Quality Assessment of Diagnostic Accuracy Studies (QUADAS-2) tool. Any discrepancies between the reviewers were resolved through discussion until a consensus was reached.

Strategy of data synthesis: Since the studies included in the meta-analysis came from different centers and covered slightly different populations, the summary was performed by applying a random effect. As the end result, the mean value with a 95% confidence interval (CI) was chosen. Statistical heterogeneity in the studies was assessed using the I2 statistics. For values above 50%, further analysis was performed to identify the source of the heterogeneity, allowing the inclusion of homogeneous studies only. The mean values were then compared between the groups (PA and WT) with a series of t-tests for the two means in order to deter-mine the significance. Additionally, qualitative CEUS features of PAs and WTs available throughout the articles were compared using the Chi2 test. Differences were considered significant at p <0.05. The analysis was performed using Statistica 13.3 (StatSoft Polska, Krakow, Poland) and PQStat 1.8.4 (PQStat Software, Poznan, Poland) software.

Subgroup analysis: The evaluation of differences in various qualitative and quantitative CEUS-derived features between PAs and WTs of the parotid gland.

Sensitivity analysis: Not applicable.

Language restriction: English.

Country(ies) involved: Poland.

Keywords: ultrasonography; salivary gland; pleomorphic adenoma; Warthin's tumor; perfusion.

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