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Systematic review of the metabolic outcomes in adult-onset craniopharyngioma

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ADMINISTRATIVE INFORMATION

Support - None.

Review Stage at time of this submission - Completed but not published.

Conflicts of interest - None declared.

INPLASY registration number: INPLASY202450087

Amendments - This protocol was registered with the International Platform of Registered Systematic Review and Meta-Analysis Protocols (INPLASY) on 17 May 2024 and was last updated on 17 May 2024.

INTRODUCTION

Review question / Objective Systematically review the literature in order to compile evidence of metabolic outcomes in adultonset craniopharyngioma, performed metaanalysis to compare prevalence of metabolic disease with Childhood Onset Craniopharyngioma (COCP).

Rationale Craniopharyngioma is a rare brain tumour with a bi-modal age distribution that can be subcategorised into adult-onset (AOCP) and childhood-onset craniopharyngioma (COCP). Whilst histologically benign, these tumours are locally aggressive and damage to structures may occur due to compression by tumour or during treatment. Damage may lead to hypothalamic obesity and a range of metabolic diseases - Type 2 diabetes (T2DM), Hypertension (HTN), lipid disorders, and Metabolic Syndrome (MetS). **Condition being studied** Craniopharyngioma, metabolic diseases - Type 2 diabetes (T2DM), Hypertension (HTN), lipid disorders, and Metabolic Syndrome (MetS).

METHODS

Search strategy Three databases (PubMed, Embase, and Scopus) were searched for relevant published articles on 04/10/2023. The detailed search strategies for each database are presented in Appendix 1. Search terms were as follows: (metabolism) OR (obesity)) OR (hypertension)) OR (health) OR (quality of life)) OR (metabolic syndrome)) AND (craniopharyngioma). Adult-onset was not specified in the search to avoid missing studies which did not explicitly state age of onset.

Participant or population Inclusion criteria

 Studies that investigate metabolic sequalae* of craniopharyngioma

• Studies that investigate metabolic sequalae* of craniopharyngioma treatment such as surgery and radiotherapy

Adult -onset craniopharyngioma

• Cohort studies, case-control studies, and crosssectional studies.

*Metabolic sequalae - metabolic syndrome, hypertension, diabetes mellitus, biochemical markers.

Exclusion criteria

- Childhood-onset craniopharyngioma
- Studies not in the English language
- Case reports and secondary literature.

Intervention NA.

Comparator NA.

Study designs to be included Systematic review.

Eligibility criteria Inclusion criteria

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• Studies that investigate metabolic sequalae* of craniopharyngioma treatment such as surgery and radiotherapy

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Information sources Three databases (PubMed, Embase, and Scopus) were searched for relevant published articles on 04/10/2023.

Main outcome(s) Initial search retrieved 2295 articles with 11 included after screening, 4 studies were suitable for meta-analysis. In total 523 patients were included F=253 (48%). Reporting of metabolic outcomes was heterogeneous throughout the studies. Pooled prevalence was 117/292 (40%) MetS, 75/315 (24%) T2DM, 205/363 (56%) HTN, and 211/425 (50%) lipid disorders. Meta-analysis suggested significantly greater likelihood of T2DM (Odds Ratio (OR) 2.49, Confidence interval (Cl) 1.3 – 4.79, p=0.01) and HTN (OR 4.21, Cl 1.28 – 13.81, p=0.02) with no significant difference in likelihood of MetS in AOCP (OR 0.75, Cl 0.48 – 1.17, p=0.21) compared to COCP. **Data management** Zotero will be used to catalogue all the studies from the searches, and then used to process the studies utilising specified inclusion and exclusion criteria. Endnote 2.0 will be used to manage references for the study but not for the searches themselve.

Quality assessment / Risk of bias analysis Systematic review will be performed according to the preferred reporting items for systematic reviews and meta-analyses (PRISMA) framework. Studies will be included and excluded according to a pre-specified inclusion / exclusion criteria which will include criteria such as exclusion of paediatric craniopharyngioma and studies not written in the English language. Studies will be included / excluded by title and abstract, after this full texts will be reviewed to assess for eligibility. If applicable the references of included studies can also be reviewed in order to identify additional studies to include.

Strategy of data synthesis Basic descriptive statistics were performed to determine pooled sample prevalence, range and mean prevalence across the studies. Wilson's score interval was used to calculate 95% confidence intervals for the pooled prevalence. Coefficient of variation was calculated to describe variability across studies.

A meta-analysis was performed for 3 outcomes -T2DM, HTN, MetS using the 4 studies which compared AOCP and COCP. Binary meta-analysis was performed where effect size was derived from pooled odds ratio and 95% confidence intervals. Random effects model was utilised to take into account variation in studies. Results were displayed as a forest plot. Chi2 test of homogeneity and I2, Tau2, H2 tests of heterogeneity were performed. Publication bias could not be assessed by funnel plot due to the low number of studies included.

Meta-analysis was performed in IBM SPSS Statistics Version 29.0.1.1 (244) (Released 2022. IBM Corp., Armonk, New York).

Subgroup analysis Meta analysis.

Sensitivity analysis Meta analysis.

Language restriction English.

Country(ies) involved UK.

Other relevant information Supervisor - Dr Pratibha Natesh (previously known as Pratibha Machenahalli). **Keywords** Craniopharyngioma, Metabolic syndrome, Type 2 Diabetes Mellitus, Hypertension.

Dissemination plans 1. Publication ina journal 2. Poster presentation at a scientific conference.

Contributions of each author

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