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# Neurosurgical Complications Following Tooth Extraction: A Systematic Review and Individual Patient Meta-Analysis

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# Abstract

Background: We aimed to review the characteristics of patients with neurosurgical complications after tooth extraction. Materials and Methods: This systematic review followed PRISMA guidelines and searched PubMed/MEDLINE, Embase, Web of Science, and Scopus databases for studies investigating neurosurgical complications post-tooth extraction. Relevant keywords for dental extraction, adverse events or complications, and neurosurgery were searched using Boolean operators. Extracted data was synthesized using proper statistical tests. Results: Among 42 studies, 47 cases (34 males, 13 females) were included. The complications were distributed as follows: 25 brain abscesses, 11 meningitis cases, 8 cerebrovascular accidents, 2 cases with both meningitis and stroke, and 1 pituitary macroadenoma. Four deaths occurred in cerebrovascular accident cases. A significant association was found between preexisting diseases and death (odds ratio = 2.15, 95% CI: 1.08-4.29, P-value = 0.03). Three mucormycosis and two mycobacterium tuberculosis cases were reported. The most common symptoms were headache (55.32%), fever (38.3%), and laterality symptoms (25.53%). Neck pain/neck rigidity was more prevalent in females (30.77% vs. 8.82%, P = 0.042), as were nausea and vomiting (30.77% vs. 8.82%, P = 0.028). Overall, 31.91% of cases had no underlying diseases. The mean time from tooth extraction to emergency room visit was 19.73 days (SD = 31.01 days), ranging from 2 to 180 days. Fourteen cases (29.79%) involved the upper jaw, 6 (12.77%) the lower jaw, and 2 (4.26%) both jaws. Conclusion: The study introduces a novel approach by systematically reviewing and analyzing individual patient data to identify specific risk factors and symptoms associated with neurosurgical complications following tooth extraction. Healthcare providers can use the identified symptoms, such as headache and fever, as key indicators for prompt evaluation and management of patients presenting after tooth extraction, especially in male patients with pre-existing conditions who are undergoing upper jaw teeth extraction.

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**Keywords:** Neurosurgical Complications; Tooth Extraction; Neurosurgical Procedures; Brain Abscess; Meningitis; Cerebrovascular Accidents; Systematic Review and Meta-Analysis

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# Introduction

Interaction between dental sciences and neurosurgery is multifaceted, as is the shared anatomy of the head and neck region [1], the emergence of the concept of the "brainoral axis" in neurosciences [2], and the effect of dental health on neurological diseases [3]. Collaborations between dentists and neurosurgeons in combined surgeries have become increasingly common, like combined craniomaxillofacial and neurosurgical procedures [4].

Dental caries and its complications are significant reasons for tooth extractions, with a high prevalence observed in the population [5]. Socio-demographic factors like gender may also influence extraction rates, with higher prevalence observed among female individuals [6]. Dental extractions, while common procedures, can lead to adverse events, such as pain and discomfort, swelling and bruising, bleeding, infection, and nerve damage [7, 8]. Inferior Alveolar Nerve (IAN) is susceptible to injury during mandibular tooth extractions [9, 10]. Lingual nerve and trigeminal nerve injuries might also happen [9, 10]. Pneumomediastinum, pneumorrhachis, pneumothorax, and pneumopericardium, are infrequently encountered but documented [11]. Another uncommon complication is surgical emphysema [12], typically associated with the use of high-speed air rotors during extractions. Additionally, osteoradionecrosis, characterized by bone tissue death due to prior radiation therapy, and complications like bite collapse and improper tooth alignment can occur post-extraction, albeit rarely [13]. Severe trismus, although not exceedingly common, may also manifest after extractions, leading to difficulties in mouth opening [14]. Neurosurgery following tooth extraction is a rare occurrence but may be necessary in cases where dental procedures inadvertently lead to complications involving nearby neurological structures. Traumatic neuralgia and posttraumatic pain syndrome been reported as complications have necessitating neurosurgical evaluation after dental procedures [15]. In some cases, neurosurgery may be necessary to decompress the inferior alveolar nerve after endodontic

treatment complications [16]. Additionally, microsurgical repair of lingual nerve injuries may be required in cases of nerve damage during third molar removal [17]. A review focused on the neurological complications associated with local anesthesia in dentistry, including adverse effects such as diplopia, ptosis, ocular paralysis, blindness, paresthesia, trismus, neuralgia, and facial palsy [18]. Another review investigated the neurological sequelae following surgical interventions on the lower molars, including adverse effects such as transient and permanent sensory deficits, often resulting from the compression or irritation of the mandibular nerve [19]. As the existing literature on neurosurgical complications following tooth extraction is fragmented and lacks a comprehensive analysis of individual patient data, we aimed to systematically review and metaanalyze the characteristics of patients with neurosurgical complications after tooth extraction. Furthermore, previous studies have primarily focused on specific aspects of dental procedures or local anesthesia, without providing a thorough understanding of the risk factors and symptoms associated with neurosurgical complications. Our study introduces a novel approach by synthesizing individual patient data to identify specific risk factors and symptoms that can serve as key indicators for prompt evaluation and management of patients presenting after tooth extraction.

## **Materials and Methods**

A systematic review was conducted to investigate neurosurgical complications following tooth extraction. The Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines were followed for reporting the findings of this systematic review [20].

#### Information sources and Search strategy:

A search strategy was developed using relevant keywords and Medical Subject Headings (MeSH) terms. The following databases were searched from inception to January 2024: PubMed/MEDLINE, Embase, Web of Science, and Scopus. The search strategy utilized the following combination of terms:

("dental extraction" or "tooth extraction" OR "milk tooth" OR "dental Manipulation" OR "dental extraction" ) AND ((brain) OR (neurosurgery) OR (stroke) OR (spine) OR (cerebrovascular event)

# Eligibility criteria

Inclusion criteria: Studies that investigated neurosurgical complications associated with dental extraction procedures. Published in peer-reviewed journals and available in English language were included. Studies with individual patient data were only included. То diagnose neurosurgery originating from complications dental sources, three conditions should be met: the absence of alternative bacteremia sources, a microbiological profile in line with oral flora, and clinical or radiographic indications of dental infection [21].

Exclusion criteria: Orbital abscess cases were excluded. Sinusitis-related infections were not included. Iatrogenic traumatic brain injury cases were not included. Cluster Headache cases were not counted as neurosurgical cases.

## Selection and Data Collection Process

Two independent reviewers screened titles and abstracts of retrieved articles based on the predefined inclusion and exclusion criteria. Full texts of potentially relevant articles were then assessed for eligibility. Any disagreements between the reviewers were resolved through discussion or consultation with a third reviewer. A standardized data extraction form was used to extract relevant information from included studies. Data extracted included study characteristics (author, year of publication, study design), participant demographics, details of dental extraction procedures, neurosurgical complications reported, and relevant outcomes.

## Study risk of bias assessment

The methodological quality of the included studies was evaluated using The CARE guideline of case reporting [22].

## Synthesis methods

Data synthesis was performed summarizing

findings from included studies, including the prevalence and types of neurosurgical complications following dental extraction. Meta-analysis was performed using STATA software, with descriptive statistics of n (%) for categorical data and mean±SD for continuous ones. Chi-square and independent t-tests were used to test various hypotheses, considering the significant value lower than 0.05.

## Results

In the initial search, a total of 2,345 articles were identified from various databases. After removing duplicates, 1,987 articles remained. Following the screening of titles and abstracts, 1,634 articles were excluded as they did not meet the inclusion criteria. Subsequently, full-text assessment was performed on the remaining 353 articles, leading to the exclusion of an additional 311 articles.

Finally, after applying the eligibility criteria, 42 articles [23-64] were included in the systematic review for data extraction and analysis, as shown in Figure-1. The main characteristics of the cases are shown in Table-1.

Among 42 studies, 47 cases were included in the study. There were 25 cases of brain abscess. Amorim et al. describe an odontogenic brain abscess with hydrocephalus. Pallesen et al. encountered multiple brain abscesses attributed to Streptococcus intermedius and Staphylococcus warneri, leading to subsequent complications such as subdural empyema and focal epileptic seizures. Hollin et al. identified a parietal abscess, while Hollin & Gross observed a right thalamic abscess. Andersen and Horton and Strojnik et al. reported brain abscesses. Wong et al. noted an occipital lobe abscess following wisdom teeth extraction, later identified as esophageal squamous cell carcinoma. Chandy et al. reported a Pott's abscess with Streptococcus intermedius and Bacteroides melaninogenicus infection. Sakashita et al. documented a complex case involving subarachnoid and intraparenchymal abscesses, lung abscesses, intracerebral hemorrhage, fusiform aneurysms, cerebral infarction, and cerebral atrophy. Clancy et al. reported a brain abscess secondary to

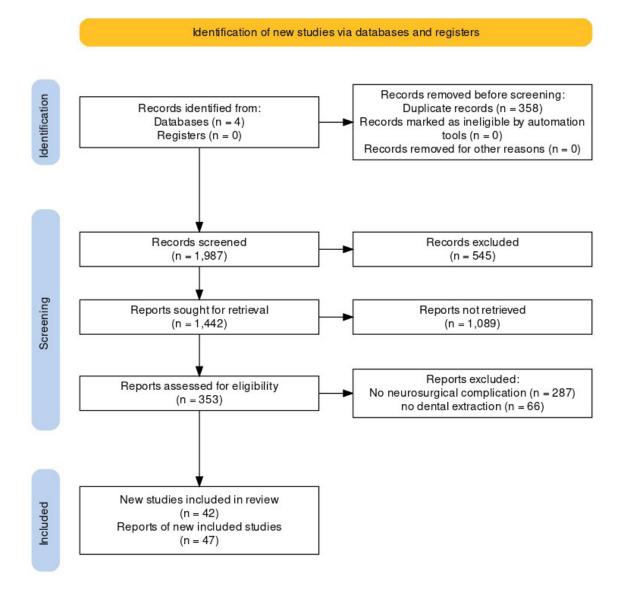


Figure 1. PRISMA flowchart of study showing steps taken from edibility assessment to study selection

Streptococcus mitis and A. meyeri infection. Brady et al. observed a brain abscess in conjunction with mitral regurgitation and left-sided weakness requiring rehabilitation. Clifton et al. documented brain abscesses along with non-convulsive status epilepticus due to hydrocephalus and hypertension. Funakoshi et al. reported an intracranial subdural abscess caused by A. meyeri and Fusobacterium nucleatum. Shibata et al. described brain abscesses secondary to chronic suppurative apical periodontitis, squamous cell carcinoma, and apical periodontitis after tooth extraction. The patient died in 6 months. Verma et al. encountered a medullary abscess secondary to tooth extraction with Streptococcus intermedius infection. Wu et al. reported odontogenic brain abscesses with septic embolic ischemic stroke. Hibberd et al. observed a temporoparietal intracerebral abscess, dental abscess, and Streptococcus anginosus infection. Heckmann et al. documented an epidural abscess secondary to dental extraction. Chang et al. identified a brain abscess caused by Streptococcus milleri group infection. Vargas et al. reported

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| Table 1. Characteristics of included studies | terist | fics of incl | luded studies  |  |   |   |  |
|--|--------|--------------|----------------|--|---|---|--|
|  | Age    | Gender       | Comorbidities  | Symptoms on ER Admission   | Extracted Tooth<br>(Tooth Identity)                 | CSF culture<br>organism                                       | Final Diagnosis  |
| Amorim et al.                                | 67     | female       | none           | headache, neck pain, no fever  | Upper left second<br>molar                          | ceftriaxone-sensitive<br>Streptococcus<br>intermedius         | OBA with hydrocephalus   |
| Kroppenstedt<br>et al.                       | 69     | male         | hypothyroidism | Headache, dizziness, fatigue,<br>blood pressure variations, left<br>thoracic pain  | Three teeth in the<br>lower left jaw                | none  | Pituitary macroadenoma   |
| Calderon-<br>Miranda                         | 26     | female       | missing        | Headache, nausea, emesis   | missing   | missing   | Subdural hematoma  |
| Pallesen et al.                              | 55     | male         | none           | Acute onset of weakness in the<br>left leg   | Professional tooth<br>cleaning                      | Streptococcus<br>intermedius and<br>Staphylococcus<br>warneri | Multiple BAs; subsequent<br>complications included<br>subdural empyema and focal<br>epileptic seizures |
| Hollin et al.<br>(a)                         | 19     | male         | missing        | Headaches, greenish foul<br>discharge from the right nostril,<br>fever (2 days)  | Two carious right<br>upper molar teeth<br>extracted | Staphylococcus<br>CJWXU.S,<br>nonhemolytic<br>streptococcus   | subdural empyema and<br>diffuse leptomeningitis  |
| Hollin et al.<br>(b)                         | 31     | female       | missing        | Headaches, painful swelling in<br>the right jaw  | Right lower molar                                   | sterile   | Subdural empyema   |
| Hollin et al.<br>(c)                         | 36     | male         | none           | Personality changes, Dysarthria,<br>focal convulsions, weakness,<br>numbness of the right side,<br>insomnia, lethargy, generalized<br>malaise, fever | A right upper<br>premolar tooth                     | missing   | parietal abscess   |
| Hollin &<br>Gross (a)                        | 25     | male         | missing        | Headaches, malaise,<br>drowsiness, confusion   | Fourteen upper teeth                                | missing   | Subdural empyema   |
| Hollin &<br>Gross (b)                        | 38     | male         | missing        | Headaches, fèver, mental<br>changes  | Infected tooth                                      | sterile   | Right thalamic abscess   |
|  |        |              |                |  |   |   |  |



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| Continue of Table         | e 1. Châ | tracteristics | Continue of Table 1. Characteristics of included studies  |   |  |  |   |
|---------------------------|----------|---------------|---|---|--|--|---|
| Martines et<br>al.        | 18       | male          | post-traumatic<br>splenectomy                             | Dysarthria, lethargy, purulent<br>rhinorrhea, fever   | 6th dental element<br>of the left side | Bacteroides,<br>alpha-hemolytic<br>Streptococci                                  | Subdural empyema<br>secondary to sinusitis with                                       |
| Andersen and<br>Horton    | 70       | male          | Hepatitis A (1983)  | Left shoulder, neck, and chest<br>numbness; "heaviness" without<br>pain; altered sensation in left<br>upper chest and arm.  |  | gram-positive<br>anaerobic coccus<br>and a Gram-negative<br>anaerobic rod        | BA  |
| Wong et al.               | 37       | male          | none  | Complaints of headache, visual<br>disturbances, throbbing pain in<br>the whole head, blurred vision,<br>colored lights, "blotches and<br>fuzzy spots," photophobia,<br>phonophobia, vertigo, and chills | Left upper and<br>lower wisdom teeth   | missing  | Occipital lobe abscess, later<br>identified as esophageal<br>squamous cell carcinoma. |
| Wohl et al.               | 44       | male          | migraine  | Severe headache, visual<br>disturbances, confusion, fever   | Missing                                | Microaerophilic<br>streptococci  | Migraine complicated by vascular infarction   |
| Nair et al.               | 40       | male          | missing   | Swelling in the right<br>temporal region for 4 months;<br>Holocranial headache for 15<br>days; Sinuses over swelling<br>with pus discharge 24h after<br>admission                                       | Ycs                                    | missing  | Calvarial tuberculosis;<br>Osteomyelitis of right<br>parietal bone;                   |
| Chandy et al.             | 21       | male          | respiratory disease<br>(sepsis and right<br>lung empyema) | Fever, frontal headache,<br>scalp and forehead swelling,<br>left-sided rhinorrhea, nasal<br>congestion.   | Left maxillary molar<br>(tooth no. 15) | Streptococcus<br>intermedius<br>and Bacteroides<br>melaninogenicus<br>infection. | Pott's abscess  |
| Cariati et al.            | 46       | male          | missing   | Temporomandibular pain,<br>swelling, fever  | Tooth 38                               | gram-positive Cocos  | Bacterial meningitis  |
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|  | Left maxillary<br>posterior region   | missing  | missing  |
|--|--|--|--|
|  | Swelling on the left side of<br>the face; diplopia; periorbital<br>ecchymosis; left eye symptoms | Diplopia, pain in the back of the<br>right eye, headache | Severe right hemiparesis, more pronounced in the leg |
| Continue of Table 1. Characteristics of included studies | missing  | Hypertension,<br>Diabetes                                | none   |
| tracteristics  | male   | male   | male   |
| e <b>1.</b> Cha  | 58   | 62   | 12   |
| Continue of Table  | Reddy et al.   | Sakashita et<br>al.                                      | Strojnik et al.                                      |
| •  |  |  |  |
| J.2024;13:e35<br>www.gmj.ir                              | 570  |  |  |

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| Complications from<br>Cavernous Sinus<br>Thrombosis; Death due | Subarachnoid and |
|--|------------------|
|--|------------------|

missing

th due to

cerebral infarction, cerebral

atrophy

BA

Streptococcus betahaemolyticus group

Streptococcus

intermedius,

species, and gram-F, Fusobacterium

negative rods

left middle cerebral artery,

fusiform aneurysms in the

intracerebral hemorrhage, lung abscess, massive

Fusobacterium sp.;

Pavimonas micra

intraparenchymal abscess,

Subarachnoid hemorrhage

missing

Upper lateral incisor

Bleeding from red and swollen

diastema formation, extrusion, gingivae, loosening of teeth,

Hypertension

female

58

Okada et al.

periodontal pocket formation

Cause of death:

BA

Streptococcus mitis

and A. meyeri

cocci initially, later

Left lower molar

Left retro-orbital headache,

right hemisensory loss, unsteady gait

Chronic right-sided hearing impairment

female

55

Clancy et al.

Gram-positive

hemorrhagic venous

missing

Left second upper premolar: 25

Left-sided toothache, swelling,

**Diabetes Mellitus** 

male

55

Reddy et al.

9

fever, frontal headaches

infarction

left temporo-frontal

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| in        |
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| Continue of Tabl          | e 1. Cha | racteristics | Continue of Table 1. Characteristics of included studies        |  |   |  |  |
|---------------------------|----------|--------------|---|--|---|--|--|
| Brady et al.              | 68       | male         | none  | Sudden onset slurred speech,<br>left-sided facial droop, and left<br>upper limb weakness. VII nerve<br>palsy. Poor dentition. Pan-<br>systolic murmur. | Non-traumatic loss<br>of a tooth one week<br>before admission | missing  | BA   |
| Clifton et al.            | 56       | male         | Hypertension,<br>cholecystectomy,<br>obstructive sleep<br>apnea | Mental changes, dry cough,<br>intermittent fever, tunnel vision,<br>memory lapse, headache, neck<br>and back pain, nausea, vomiting                    | Left the second<br>molar                                      | Gram-positive<br>anaerobic<br>streptococcal ns   | BAs; Non-convulsive status<br>epilepticus  |
| Funakoshi<br>et al.       | 57       | female       | Hypertension  | Headache, Left arm numbness<br>and weakness  | Dental problems<br>requiring tooth<br>extractions             | A. meyeri and<br>Fusobacterium<br>nucleatum  | Intracranial subdural abscess  |
| Yoshii et al.             | 54       | male         | none  | Severe headache and malaise,<br>no nausea or vomiting  | Second and third<br>molars of the left<br>lower jaw           | Peptostreptococcus<br>tetradius,<br>Streptococcus<br>milleri,<br>Streptococcus<br>salivarius,<br>Capnocytophaga spp. | Bacterial meningitis, later<br>complicated by a right<br>subdural empyema                        |
| Shibata et al.<br>(a)     | 62       | male         | Esophageal cancer,<br>Type 2 diabetes<br>mellitus               | Headache, fever, motor aphasia,<br>right hemiparesis   | Right maxillary<br>second premolar and<br>second molar        | S. intermedius   | BAs; patient deceased 6<br>months after surgery.   |
| Shibata et al.<br>(b)     | 68       | male         | Advanced non-<br>small-cell lung<br>cancer                      | Left hemiparesis and fever   | missing   | S. intermedius   | BA secondary to squamous<br>cell carcinoma and apical<br>periodontitis after tooth<br>extraction |
| Verma et al.              | 68       | male         | none  | Malaise, numbness in feet,<br>lower limb weakness, choking,<br>respiratory distress  | Right upper jaw<br>tooth                                      | Streptococcus<br>intermedius infection   | Medullary abscess secondary<br>to tooth extraction,  |
| Wu et al.                 | 32       | male         | none  | Progressive headache left upper<br>limb weakness, left facial palsy  | missing   | sterile  | OBAs with septic embolic ischemic stroke   |
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| Singh et al.              | 47       | male   | Polycystic kidney<br>disease (PCKD),<br>Hypertension<br>(HTN)   | Toothache, Right eye pain,<br>Orbital swelling, Fever,<br>Dyspnea                              | Tooth extraction<br>10 days before<br>symptom onset                         | sterile                               | Mucormycosis, Kluyvera<br>intermedia, Pseudomonas<br>aeruginosa sepsis, Acute<br>infarcts, Thrombosis,<br>Cavernous sinus thrombosis |
|---------------------------|----------|--------|---|--|---|---------------------------------------|--|
| Naganawa<br>et al.        | 76       | male   | Hyperthyroidism,<br>Myocardial<br>Infarction,<br>Chronic Subdural<br>Hematoma, Aortic<br>Dissection, Chronic<br>Kidney Disease,<br>Hypertension,<br>Chronic DIC | Spontaneous pain in the upper<br>front teeth region  | Maxillary left<br>central and lateral<br>incisors, Right<br>central incisor | missing                               | Death due to Intracranial<br>Hemorrhage associated with<br>Aortic Dissection and DIC   |
| Choi et al.               | 39       | male   | History of<br>Behçet's disease,<br>particularly NBD   | Hypesthesia of the left face and<br>extremity - Ataxia -Memory<br>disturbance - Disorientation | Molar tooth<br>extraction   | Sterile                               | hemorrhagic infarction   |
| Hibberd et al.            | 11       | male   | none  | 2-week history of dull<br>continuous headache, 1-week<br>history of nausea and vomiting        | Lower left second<br>primary molar<br>(tooth 75)                            | Streptococcus<br>anginosus (day 2)    | Temporoparietal<br>intracerebral abscess   |
| Prabhu et al.             | 70       | male   | Uncontrolled<br>diabetes mellitus   | Altered sensorium, vomiting,<br>decreased oral intake, right<br>facial swelling                | Right tooth   | mucormycosis                          | neural invasion  |
| Hobson et al.             | 35       | female | none  | Severe headache, facial<br>swelling, mental status changes                                     | Left maxillary third<br>molar   | Suggestive of<br>bacterial meningitis | Acute meningoencephalitis,<br>subdural empyema,<br>intraparenchymal<br>hemorrhage, neurologic<br>deficits                            |
| Continue in the next page | e next p | age    |   |  |   |                                       |  |



| Continue of Tabl          | e 1. Cha | uracteristics | Continue of Table 1. Characteristics of included studies |  |  |   |  |
|---------------------------|----------|---------------|--|--|--|---|--|
| Heckmann<br>et al.        | 77       | female        | none   | Neck pain, pronounced neck<br>stiffness  | Fractured first<br>premolar in the left<br>maxilla                   | Streptococcus<br>intermedius (Milleri)  | Epidural abscess   |
| Chang et al.              | 6.7      | male          | Ebstein's anomaly,<br>intellectual<br>disabilities       | Sudden vomiting, loss of<br>consciousness, facial spasm  | Left lower<br>deciduous central<br>and lateral incisors              | Streptococcus<br>milleri group and<br>Methicillin-resistant<br>Staphylococcus<br>aureus | BA   |
| Chang et al.              | 63       | female        | none   | Right frontal headaches,<br>puffiness of the right eye,<br>fever, diplopia, high fever,<br>right-eye ptosis, limitation of<br>eyeball movement, stiff neck,<br>positive meningeal signs,<br>muscle weakness, sensation<br>responsive to pain stimulation,<br>leucocytosis, neutrophilia.         | Tooth extraction   | Fusobacterium<br>nucleatum  | Bacterial meningitis and<br>ischemic changes.<br>Subsequent left-side<br>hemiplegia. |
| Vargas et al.             | 18       | male          | none   | Headache, vomiting, aphasia,<br>weakness in left extremities,<br>behavior and mood alterations,<br>fever   | Multiple teeth were<br>extracted three<br>months before<br>admission | Arcanobacterium<br>haemolyticum   | BA   |
| Ng et al.                 | 33       | male          | ecstasy adiction<br>visited prostitutes                  | Acute confusion; semiconscious<br>(GCS: 10/15); urinary<br>incontinence; fever; upper<br>respiratory tract symptoms;<br>expressive dysphasia; right-<br>sided pyramidal signs; apical<br>pansystolic murmur; no<br>Kussmaul's breathing; neck<br>stiffness; no Kernig's or<br>Brudzinski's signs | Yes (two weeks<br>before admission)                                  | sterile   | Meningitis and Brain infarct;<br>Endocarditis with mitral<br>valve vegetation        |
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| Combined<br>mucormycosis and<br>aspergillosis                     | Fusobacterium<br>nucleatum and<br>Staphylococcus<br>epidermidis | Group A<br>Streptococcus (GAS) BA                           | P. alactolyticus and Bacterial meningitis<br>MTB                         | Streptococcus BA with intracerebral anginosus left hemiparesis | Abscess in the right<br>cerebellar hemisphere<br>sigmoid diverticulitis<br>with an adjacent abscess.pus drainage)Complete recovery after<br>surgical drainage. |   |
|---|---|---|--|--|--|---|
| Right first premolar  | 10 teeth  | Front milk tooth<br>(exact identity ) Stre                  | Dental extraction<br>performed 2 days<br>before the onset of<br>headache | missing  | missing  | erculosis   |
| Loss of consciousness, swelling,<br>tenderness of the right cheek | Acute confusion, fever, and aphasia                             | Fever, severe headache, neck<br>stiffness, nausea, vomiting | Progressive headache, fever<br>>39°C, neck stiffness                     | Shortness of breath and fever<br>following tooth extraction    | Dizziness, worsening<br>headaches, blurry vision   | <b>OBA:</b> Odontogenic brain abscess, ; <b>BA:</b> Brain Abscess, <b>MTB:</b> Mycobacterium tuberculosis |
| Ketoacidotic<br>diabetic coma                                     | Hereditary<br>haemorrhagic<br>telangiectasia<br>(HHT)           | none  | missing  | missing  | Ductal carcinoma<br>in situ,<br>hypothyroidism,<br>diverticulosis  | ess, ; <b>BA:</b> Brain Absc  |
| female  | female  | female  | male   | male   | female   | orain absc  |
| 50  | 63  | 9   | 44   | 78   | 56   | genic l   |
| Alfano et al.   | Corre et al.  | Hayashi et al.  | Liao et al.  | Lin et al.   | Al Moussawi<br>et al.  | <b>OBA:</b> Odonto§   |

a brain abscess caused by Arcanobacterium haemolyticum. Corre et al. linked hereditary haemorrhagic telangiectasia (HHT) with neurological complications of dental extraction. Hayashi et al. documented a brain abscess with meningitis due to Group A Streptococcus (GAS) infection. Lin et al. reported a Streptococcus anginosus brain abscess with intracerebral hematoma. Al Moussawi et al. encountered an abscess in the right cerebellar hemisphere originating from Streptococcus intermedius, coupled with sigmoid diverticulitis and an adjacent abscess, ultimately achieving complete recovery after surgical drainage.

There were 13 cases of meningitis. The reported cases depict a range of meningitis presentations with diverse etiologies and complications. Hollin et al. (a) and Hollin et al. (b) documented subdural empyema with diffuse leptomeningitis, the former linked to tooth extraction complications. Hollin & Gross (a) reported subdural empyema secondary to tooth extraction, highlighting dental procedures as potential sources of intracranial infections. Martines et al. observed subdural empyema secondary to sinusitis with Bacteroides and alpha-hemolytic Streptococci infection, underscoring the significance of sinus-related complications. Nair et al. encountered calvarial tuberculosis with osteomyelitis of the right parietal bone, confirming TB infection through positive Mantoux and TB interferon gamma tests. Cariati et al. reported bacterial meningitis of dental origin, emphasizing the oral-healthrelated nature of the infection. Yoshii et al. documented bacterial meningitis, later complicated by a right subdural empyema. Prabhu et al. reported invasive zygomycosis (mucormycosis) with extensive angioinvasion and neural invasion, illustrating a rare but severe form of fungal meningitis. Hobson et al. observed acute meningoencephalitis with additional complications such as left pterygoid muscle abscess, subdural empyema, intraparenchymal hemorrhage, and resulting neurologic deficits. Chang et al. reported bacterial meningitis secondary to Fusobacterium nucleatum, complicated by ischemic changes in the brain and subsequent left-side hemiplegia. Ng et al.

documented meningitis and septic emboli, brain infarction, and endocarditis with mitral valve vegetation, showcasing the systemic impact of the infection. Alfano et al. described combined mucormycosis and aspergillosis of the rhinocerebral region, emphasizing the potential for multiple fungal infections. Liao et al. reported bacterial meningitis with coinfection of P. alactolyticus and Mycobacterium tuberculosis (TB), highlighting the coexistence of different pathogens in meningitis cases. These cases collectively underscore the diverse etiologies, complications, and severity associated with meningitis, emphasizing the importance of prompt diagnosis and appropriate management.

The cases reported 8 cerebrovascular accidents (CVAs) with distinct etiologies and complications. Kroppenstedt et al. documented a pituitary macroadenoma with a secondary infection post-tooth extraction, emphasizing the potential complications associated with dental procedures. Calderon-Miranda observed a subdural hematoma, highlighting the intracranial consequences of traumatic injuries. Wohl et al. reported migraine complicated by vascular infarction, showcasing the association between migraines and cerebrovascular events. Reddy et al. reported complications from cavernous sinus thrombosis (CST) leading to death, underscoring the severity of this condition. et al. identified Okada subarachnoid hemorrhage as the cause of death, indicating a rupture of blood vessels into the space surrounding the brain. Reddy et al. (b) documented a rhino-orbital infection from a dental source with cavernous sinus extension causing left temporo-frontal hemorrhagic venous infarction, illustrating the potential for localized infections to impact venous structures. Singh et al. encountered a complex case involving mucormycosis, Kluyvera intermedia, Pseudomonas aeruginosa sepsis, acute infarcts, thrombosis, and cavernous sinus thrombosis, highlighting the multifactorial nature of cerebrovascular complications. Naganawa et al. reported death due to intracranial hemorrhage associated with aortic dissection and disseminated intravascular coagulation (DIC), emphasizing the systemic impact of vascular disorders. Choi *et al.* documented Behçet's disease, particularly neuro-Behçet's disease (NBD), illustrating the association between inflammatory conditions and cerebrovascular involvement.

#### Descriptive statistics

In this study involving 47 participants (34 males, 13 females), the mean age for male participants was  $45.17 \pm 20.60$  years, while the mean age for female participants was  $49.54 \pm 19.63$  years. The independent t-test revealed no statistically significant difference in mean age between genders (P = 0.5134).

most common symptom overall The was headache, reported by 55.32% of all participants. Fever was presented in 38.3% of the cases. Laterality symptoms, such weakness, hemiparesis, and sensory as disturbances, were noted in 25.53% of individuals. Neck pain or neck rigidity followed, with a prevalence of 14.89%, while symptoms encompassing dizziness, fatigue, malaise, and vertigo were observed in 19.15% of cases. Nausea and vomiting were reported in 14.89% of cases. When examining gender differences, the prevalence of neck pain/neck rigidity was significantly higher in females (30.77%) compared to males (8.82%), with a P-value of 0.042. Nausea and vomiting

| Table 2. | Symptoms | comparison a | among genders |
|----------|----------|--------------|---------------|
|          |          |              |               |

also showed a notable gender difference, with 30.77% of females experiencing these symptoms compared to 8.82% of males (P = 0.028). While fever was a prevalent symptom in both genders, there was no significant difference observed (P = 0.632).

Other rare symptoms are as follows: gait disturbances, facial droop/spasm, cardiac symptoms, focal convulsions, rhinorrhea/ sinusitis, insomnia, personality changes, and blood pressure variations.

Among males, 29.41% had no underlying diseases, while 38.46% of females fell into the same category, resulting in an overall rate of 31.91%. Hypertension was reported in 11.76% of males and 15.38% of females, with a combined prevalence of 17.65%. Diabetes was less prevalent, with 5.88% of males and no cases reported among females. Among the patients, the distribution of various disorders was as follows: hypothyroidism was reported in 2 individuals, and post-traumatic splenectomy and hepatitis A conditions were each identified in 1 patient. Additionally, respiratory diseases were observed in 3 cases. Cancers, kidney diseases, and congenital disorders each affected 2 patients. Cardiac stroke/cerebral hemorrhages, diseases. neurological diseases, addiction, and surgical diseases were each reported in 1 patient.

|  | male       | female    | Total      | P-value |
|--|------------|-----------|------------|---------|
| n  | 34         | 13        | 47         | -       |
| Headache   | 18(52.94%) | 8(61.54%) | 26(55.32%) | 0.41    |
| Neck pain/neck rigidity                                      | 3(8.82%)   | 4(30.77%) | 7(14.89%)  | 0.042   |
| Dizziness/Fatigue/malaise/vertigo                            | 8(23.53%)  | 1(7.69%)  | 9(19.15%)  | 0.254   |
| Nausea & vomiting  | 3(8.82%)   | 4(30.77%) | 7(14.89%)  | 0.028   |
| Fever  | 14(41.18%) | 4(30.77%) | 18(38.3%)  | 0.632   |
| Laterality (weakness, hemiparesis, sensory)                  | 11(32.35%) | 1(7.69%)  | 12(25.53%) | 0.103   |
| Chest pain   | 1(2.94%)   | 0(0%)     | 1(2.13%)   | 0.548   |
| Difficulty in speaking/slurred speech/<br>Dysarthria/aphasia | 4(11.76%)  | 2(15.38%) | 6(12.77%)  | 0.665   |
| Numbness in any part of the body                             | 5(14.71%)  | 0(0%)     | 5(10.64%)  | 0.159   |
| Cough/respiratory  | 4(11.76%)  | 0(0%)     | 4(8.51%)   | 0.214   |
| Memory disturbances  | 4(11.76%)  | 0(0%)     | 4(8.51%)   | 0.214   |
| Mental state changes (confusion)                             | 10(29.41%) | 2(15.38%) | 12(25.53%) | 0.387   |
| Visual disturbances  | 7(20.59%)  | 0(0%)     | 7(14.89%)  | 0.088   |

The results of various cultures from different studies revealed a diverse spectrum of microbial isolates. Streptococcus intermedius was identified in multiple cases, either alone or in combination with other pathogens such as Staphylococcus warneri, Streptococcus betahaemolyticus group F, and Fusobacterium species. Staphylococcus CJWXU.S and nonhemolytic streptococci were observed in separate cases. Gram-positive anaerobic cocci and Gram-negative anaerobic rods were detected together in one case. Microaerophilic streptococci were reported, as well as Grampositive cocci initially, later identified as Streptococcus mitis and A. meyeri. Peptostreptococcus tetradius, Streptococcus milleri, Streptococcus salivarius, and Capnocytophaga spp. were identified together in a distinct case. Other findings included Fusobacterium nucleatum, Arcanobacterium haemolyticum, and Methicillin-resistant Staphylococcus aureus. Notably, sterile cultures were reported in several cases, while Streptococcus anginosus was specifically mentioned in one case, suggesting a potential association with bacterial meningitis.

Time from tooth extraction to emergency room visit was examined for 40 observations, indicating a mean duration of 19.73 days, with a standard deviation of 31.01 days. The range spanned from a minimum of 2 days to a maximum of 180 days.

In cases where data is available, 29.79% (14 cases) of dental extractions are from the upper jaw, 12.77% (6 cases) to the lower jaw, and 4.26% (2 cases) involve both the upper and lower jaws simultaneously.

The regression analysis conducted on death among cases of neurosurgical complications after tooth extraction reveals that age and

gender do not significantly influence the likelihood of death, as indicated by odds ratios of 1.07 (95% CI: 0.98-1.17) and 0.86 (95% CI: 0.08-9.11), respectively, with P-values of 0.108 and 0.901. However, a statistically significant association is observed between the number of preexisting diseases and death, with an odds ratio of 2.15 (95% CI: 1.08-4.29) and a P-value of 0.03, suggesting that each additional preexisting disease increases the odds of death by approximately 2.15 times. Conversely, the time from tooth extraction to ER symptoms does not significantly impact the likelihood of death, with an odds ratio of 1.01 (95% CI: 0.98-1.04) and a P-value of 0.718. hypertension shows a statistically significant association with death, as evidenced by an odds ratio of 9.75 (95% CI: 1.07-89.2) and a P-value of 0.044. This suggests that individuals with hypertension are at significantly higher odds of death compared to those without hypertension.

The quality of included studies was assigned as 16 studies with high quality, 15 with intermediate, and 16 studies with low quality, as shown in Table-4.

## Discussion

Our study aimed to investigate the characteristics and outcomes of patients experiencing neurosurgical complications following tooth extraction. This study demonstrates typical cases of post-tooth extraction neurosurgical complications. While being indicated, we cannot delay dental care in any case, there should be caution regarding male patients who have more than one underlying disease and need upper jaw dental manipulation, while female

**Table 3.** Logistic regression of the predictors of death among patients with neurosurgical complications after tooth extraction

|   | OR   | lower<br>95%CI | upper<br>95%CI | P-value |
|---|------|----------------|----------------|---------|
| age                                       | 1.07 | 0.98           | 1.17           | 0.108   |
| gender                                    | 0.86 | 0.08           | 9.11           | 0.901   |
| number of preexisting diseases            | 2.15 | 1.08           | 4.29           | 0.03    |
| time from tooth extraction to ER symptoms | 1.01 | 0.98           | 1.04           | 0.718   |
| hypertension                              | 9.75 | 1.07           | 89.2           | 0.044   |

and lower jaw incidents are also possible. Risk factors of post-dental extraction shortterm complications might include traumatic extraction, tobacco use, oral contraceptives, female gender, and preexisting infections [61-63]. Additionally, hemorrhage after extraction could be linked to the expertise level of practitioners and patient-specific factors like bleeding disorders [62-64]. Poor oral hygiene, smoking, and underlying systemic conditions are associated with suppurative alveolitis, while post-extraction pain may result from factors like extraction complexity, insufficient pain control, and individual pain tolerance [62-64]. Moreover, postoperative infections are more prevalent in individuals with compromised immune systems and inadequate oral hygiene practices [61-64]. But, in our study, male gender was more prominent. However, the incident of post-dental extraction neurosurgical complication is not a short-term outcome and mostly happened after 2 weeks of the dental extraction. So, it seems that the risk factors and pathophysiological nature of these complications are far away from the classic complications of dental extraction.

In our study, hypertension and multiple preexisting conditions were significant predictors of death. The relationship between hypertension and neurosurgical complications can be complex. There is a case report of the sudden increase in blood pressure and intracerebral hemorrhage in a normotensive patient and death in the dentistry room [65]. Research suggests a link between trigeminal nerve stimulation, including methods like trigeminal nerve combing or proprioceptive stimulation, and alterations in arterial blood pressure [66, 67]. The involvement of trigeminal nerve inputs in governing cerebral blood flow also suggests potential implications for blood pressure regulation [68]. It's generally advised to avoid emergency dental procedures in patients with severely elevated blood pressure (>180/110 mmHg) due to increased risks [69].

Most vases were gram-positive Cocos bacteria (22 cases, 46.8%) as the source of infection and abscess, but rare cases of Capnocytophaga spp. (Flavobacteriia), Peptostreptococcus tetradius (Clostridia) in the study of Yoshii *et al.* [40], MTB and aspergillosis in Liao

et al. study [58], and Arcanobacterium haemolyticum in Vargas et al. study [52] were seen. Most observed co-infection was a coincidence of isolates of gram-negative anaerobic rods and gram-positive Bacilli in 5 cases. Gram-positive cocci bacteria, like Streptococcus intermedius and Anaerococcus prevotii, have been associated with brain abscesses stemming from dental origins [70]. These microbes reside naturally in the oral cavity and can lead to infections if introduced into the bloodstream, frequently due to dental procedures or infections [71]. Anaerococcus prevotii, characterized as a gram-positive coccus, has emerged as a potential pathogen responsible for brain abscesses, some of which are linked to dental sources [72]. Similarly, Parvimonas micra, another grampositive anaerobic coccus prevalent in the oral mucosa, has been linked to cerebral abscesses, often arising from dental infections [73].

Maurer et al. reported a case of meningitis that didn't require neurosurgical intervention and CSF culture was sterile [74]. In our study there were 7 cases in which no infective source was isolated from abscess or CSF cultures. A seldom-seen event, meningitis caused by Capnocytophaga spp. is infrequent but warrants consideration in individuals with underlying health issues or predisposing factors, like compromised immune systems [75]. Also, it was reported after a dog bite [76]. Another study showed that lower jaw wisdom tooth extraction causes more cases of infection than upper jaw [77]. However, in our study, most cases were experiencing the incident after the upper jaw manipulation. The reason behind this difference could be attributed to the diverse pathophysiology of the complications. However, it's necessary to acknowledge the limitations of this study. Firstly, the relatively small sample size of cases limits the generalizability of the findings. Also, we cannot estimate the prevalence of this condition in public as no data is available from any cohort and the mentioned risk factors are in fact the classic representation post-tooth extraction neurosurgical of complications. Furthermore, the lack of a control group hinders the ability to establish causality or determine the true prevalence of complications following tooth extraction.

#### Conclusion

In conclusion, this systematic review and individual patient meta-analysis showed characteristics and outcomes of neurosurgical complications following tooth extraction. The study revealed a variety of complications including brain abscess, meningitis, cerebrovascular accidents, and others, with notable gender differences in symptom presentation. Headache and fever emerged as the most common symptoms, showing their importance in prompt evaluation and management, particularly in male patients with pre-existing conditions undergoing upper jaw teeth extraction.

#### **Conflict of Interest**

The authors have no conflicts of interest relevant to this article to disclose.

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