Management of Penetrating Skull Base Injury: A Single Institutional Experience and Review of the Literature

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Received 7 May 2017; Accepted 18 June 2017; Published 30 July 2017

Background. Penetrating skull base injury (PSBI) is uncommon among head injuries, presenting unique diagnostic and therapeutic challenges. Although many cases of PSBIs have been reported, comprehensive understanding of its initial diagnosis, management, and outcome is still unavailable. Materials and Methods. A retrospective review was performed for patients treated in neurosurgical department of Changzheng Hospital for PSBIs. Presurgical three-dimensional (3D) Slicer-assisted reconstructions were conducted for each patient. Then we reviewed previous literature about all the published cases of PSBIs worldwide and discussed their common features. Results. A total of 5 patients suffering PSBIs were identified. Penetrating points as well as the surrounding neurovascular structures were clearly visualized, assisting in the presurgical planning of optimal surgical approach and avoiding unexpected vascular injury. Four patients underwent craniotomy with foreign bodies removed successfully and 1 patient received conservative treatment. All of them presented good outcomes after proper management. Conclusion. Careful physical examination and radiological evaluation are essential before operation, and angiography is recommended for those with suspected vascular injuries. 3D modeling with 3D Slicer is practicable and reliable, facilitating the diagnosis and presurgical planning. Treatment decision should be made upon the comprehensive evaluation of patient’s clinicoradiological features and characteristics of foreign bodies.

1. Introduction

Penetrating injuries of the skull base caused by foreign bodies are relatively uncommon, representing about 0.4% of head injuries [1, 2]. Various foreign bodies have been reported in penetrating skull base injuries (PSBIs), including wood, bamboo, metallic fragments, and toothbrushes. PSBIs could present unique diagnostic and therapeutic challenges. Patients may be initially asymptomatic but subject to serious events for several days, months, or even years after the injuries [3]. It is not difficult to detect most of intracranial foreign bodies by head computed tomography (CT) scan. However, the relationships between foreign bodies and its surrounding structures are hard to decide, which is the prerequisite for the management of these injuries [4, 5].

Treatment for PSBIs includes the surgical retraction of foreign bodies, prevention of infection, management of vascular injuries, reconstruction of skull base, and so forth [3, 5–7]. Due to the low incidence, there is no sufficient literature about the diagnosis and management of these patients. The aim of our study is to present patients of PSBIs treated in our department and review related literature in order to highlight the proper management of PSBIs and improve prognosis in the long run.

2. Materials and Methods

A retrospective review was conducted for patients who were treated in our hospital between January 2010 and September 2016 for PSBIs. This study was performed in accordance with the Declaration of Helsinki (1964) and approved by the investigational review board of Changzheng Hospital. Informed consent was available for each patient. Presurgical head CT scans were performed for all patients. Moreover, digital subtraction angiography (DSA) was conducted on suspicion of vascular injuries and magnetic resonance imaging (MRI) was performed in case of nonmetallic objects. Data concerning patients’ demographics, mechanisms of
injury, medical managements, complications, and prognosis were collected by two authors (D. F. Zhang and J. G. Chen).

In order to visualize the location of foreign body and its relationship with surrounding structures, three-dimensional (3D) Slicer-assisted reconstructions were conducted by a professional neuroradiologist (K. W. Han) according to presurgical imaging data. During the reconstruction, all the Digital Imaging and Communications in Medicine (DICOM) images were imported into 3D Slicer (3D Slicer 4.0–4.4; Surgical Planning Laboratory, Harvard University, USA). Segmentation of skull, foreign body, and cerebral artery was first performed with built-in modules in Slicer. Individual models of each structure were created, which could be rotated and viewed from any perspective (Figure 1(g); Figures 2(g)–2(i); Figure 3(g)).

3. Results

A total of 5 patients with PSBIs were identified. There were 4 males and 1 female aged 29–75 years. They were all victims of tumble or work-related accident. Four of them underwent surgical retrieval of foreign bodies and 1 patient received conservative treatment. Penetrating points as well as the surrounding neurovascular structures were clearly visualized in 3D models, assisting in the presurgical planning of optimal surgical approach and avoiding unexpected vascular injury. Details regarding patients’ demographics, locations of foreign bodies, and treatment were listed in Table 1.

3.1. Representative Case 1. This 75-year-old female was admitted to her local hospital with complaints of headache and dizziness for 3 days. She was conscious and neurologically intact with a slightly elevated body temperature. Head CT scan revealed a low density 4 cm long foreign body extending from the left orbit to superior orbital fissure and posteriorly to the left temporal lobe (Figures 1(a)–1(c)). Brain tissue surrounding the foreign body was swollen and signs of abscess were indicated (Figures 1(d)–1(f)). The patient recalled that while she was walking in a bamboo garden, she tripped and fell forward, striking her left forehead on a bamboo stick. She did not feel any discomfort except the pain on her left upper eyelid. Symptoms of headache and fever emerged 5 days later and she was taken to the hospital by her family 8 days after the injury. Anti-infective therapies were given before she was transferred to our hospital. Careful physical examination revealed a slight skin scar on her left upper eyelid. Head DSA was performed later with no signs of vascular injuries although the bamboo stick was adjacent to the left middle cerebral artery (MCA) in the 3D reconstruction model (Figure 1(g)). Anti-infectious treatment was administrated for 12 days to control the brain abscess before surgery. During operation, an orbitozygomatic approach was adopted and the pterional craniotomy was first performed. Then superior orbital fissure was revealed after removing the great wing of sphenoid. We explored the abscess in the temporal lobe. After yellowish pus in the abscess was removed, the distal end of the bamboo stick was then visualized. We opened the dura and orbital fascia along the bamboo stick to expose its full length (Figure 1(h)). The stick was removed completely under direct visualization (Figure 1(i)) and dura was sutured in a water tight fashion. The postoperative course was uncomplicated. Broad-spectrum antibiotics were given until she was discharged free of symptoms 10 days after operation.

3.2. Representative Case 2. This 32-year-old man was hit by the fragments of a burst grinding wheel on his left cheek during working hours. He was taken to the local hospital complaining of headache and kept neurologically intact on physical examination. Head CT scan suggested a short piece of metal fragment locating right between the left maxillary and ramus mandibulae and a long piece sticking in the left temporal lobe. Both pieces presented to be highly dense on the CT scan (Figures 2(a)–2(c)). The patient was then transferred to our department for further treatment 5 days later. DSA suggested no obvious vascular injury (Figures 2(d) and 2(e)), while axial CT scan revealed close relationship between the bone fragment and branch of MCA (Figure 2(f)). The exact location of foreign body could be visualized clearly on the 3D reconstruction model (Figures 2(g)–2(i)). Conservative management was given to control the infection and surgery was performed 15 days after the injury. During operation, short piece of the foreign body was first removed through an intraoral incision. Then a temporozygomatic approach was performed to remove the long piece, which was visualized at the infratemporal fossa after the zygomatic arch was detached and retracted downwardly with temporal muscle. Several pieces of bone fragment were found around the foreign body. After the lateral portion of infratemporal fossa was drilled off, the object was removed in a retrograde fashion. During this process, yellowish pus was drained into infratemporal fossa through the intracranial trajectory of foreign body. This trajectory was not explored and defect of dura was sealed with muscle flap (Figures 3(a)–3(e)). Postoperative skull radiography showed complete removal of the foreign body (Figure 3(f)). A small bone fragment was left in place due to its close relationship with branch of MCA, which was found to be stenotic during follow-up (Figure 3(g)). The patient was discharged 1 week after operation with sporadic focal epilepsy which was controlled well with carbamazepine, and he recovered well without abscess formation at three-month follow-up (Figures 3(h) and 3(i)).

4. Discussion

4.1. Literature Review. Clinioradiological features of PSBIs in previous literatures were reviewed and summarized in Table 2 [2, 3, 8–24]. Most of the subjects were males (66%, 21/32) with an average age of 24 years old. 41% (13/32) of the patients were children under 10 years of age. As for the mechanism of injury, children injuries were all caused by tumbling or falling, while the injury of 4 adults resulted from suicidal or homicidal attempt, which was absent in our case series. The most common foreign body was metallic (12/32), followed by wooden (9/32) and plastic (9/32). One patient was injured by a wild deer’s antler and another one was attacked by bear paws, both of which had no retained foreign bodies. As a weak area of skull, the orbit was the most
common penetrating point, followed by oral or nasal cavity and maxillofacial region.

4.2. Diagnosis of PSBIs. After careful physical examination, proper radiological examination on the basis of patients' condition is necessary. Although the importance of head CT scan in the management of PSBIs has been emphasized in previous literatures, several instructions should be noticed. Firstly, the density of foreign bodies on CT scan varies according to their types. For example, metal presents as high density, while wood or plastics are of isodensity or low density and difficult to identify. Secondly, the density of some foreign bodies would change over time. For example, bamboo is of low density on initial CT scan, but it would be of high density...
later, which frequently leads to misdiagnosis (Case 1) [25]. Therefore, MRI is a useful supplement to CT in the detection of nonmetallic foreign bodies.

Angiography such as the CT angiography, MR angiography, or DSA is highly recommended for patients in suspicion of artery injuries or traumatic aneurysms [7, 26, 27]. As demonstrated in Case 2, foreign bodies may not cause immediate vascular injury sometimes but lead to cerebral vasospasm or stenosis in the long run, highlighting the importance of angiography in the setting of PSBIs.
| Case | Age and gender | Mechanism of injury | Type and size of foreign body | Penetrating pathway | Symptoms and concurrent conditions | DSA findings | Interval to operation | Operation | Surgical approach | Residual fraction | Antibiotics used | Symptoms at discharge |
|------|----------------|---------------------|-------------------------------|---------------------|-----------------------------------|--------------|----------------------|-----------|---------------------|-----------------|-----------------|---------------------|
| 1    | 75/female      | Accident            | Bamboo, 7 cm                  | Left orbit-superior orbital fissure-anterior skull base-left temporal lobe | Headache, dizziness, brain abscess | Negative     | 20 days              | Yes       | Orbitozygomatic approach and pterional craniotomy | No               | Metronidazole, ceftazidime, vancomycin, linezolid, biapenem | Free of symptom |
| 2    | 32/male        | Accident            | Grinding wheel, 6.5 cm        | Left maxilla-infratemporal fossa-middle cranial fossa, cistern of lateral sulcus | Headache, brain abscess | Negative     | 19 days              | Yes       | Temporozygomatic approach | No               | Linezolid         | Epilepsy            |
| 3    | 42/male        | Accident            | Electrodrill, 2.2 cm          | Left eyebrow-anterior cranial base-left frontal lobe | Headache, cerebral contusion | Not conducted | No operation | No       | —                 | Yes             | Linezolid, biapenem | Slight headache    |
| 4    | 29/male        | Accident            | Screw, 5.4 cm                 | Right orbit-anterior cranial base-right frontal lobe | Headache, blurred vision, anterior skull base fracture, oculomotor nerve injury | Not conducted | 2 days               | Yes       | Frontotemporal approach | No               | Vancomycin, linezolid | Improved vision, oculomotor nerve injury |
| 5    | 40/male        | Accident            | Hot projective oil paint, 3.4 cm | Right orbit-anterior cranial base-right frontal lobe | Headache, blindness, right eye penetrating injury, frontal hematoma | Not conducted | 4 days               | Yes       | Subfrontal approach | Yes             | Vancomycin, ceftriaxone | Blindness           |
| Author and year         | Age and gender | Side/size | Material                  | Mechanism of injury | Penetrating pathway | Symptom          | Surgical approach                     | Infection | Outcome         |
|-------------------------|----------------|-----------|---------------------------|---------------------|---------------------|------------------|----------------------------------------|-----------|-----------------|
| Matsumoto et al., 1998 | 3/female       | Left/3 cm | Plastic chopstick          | Falling             | Optic canal         | Blindness        | Craniotomy                            | NK        | Blindness       |
| Matsumoto et al., 1998 | 57/male        | Right/2 cm | Wooden chopsticks          | Stick into chopstick| Optic canal         | Blindness        | Craniotomy                            | NK        | Blindness       |
| Hebecker, 2009          | 32/male        | Left/no foreign body       | Wild deer’s antler      | Accident            | Left orbit floor/lateral orbit Preauricular area to posterior fossa | Coma            | Bifrontal intradural neurosurgical approach | No        | Free of symptom |
| Ishikawa et al., 2000   | 4/male         | Left/6 cm | Wooden chopstick           | Falling             | Preauricular area to posterior fossa | Fever, headache | Suboccipital craniectomy                | Brain abscess | Ataxia and dysmetria |
| Matsuyama et al., 2001  | 1/male         | Right/2.5 cm | Wooden chopsticks          | Falling             | Superior orbital fissure | Swollen eye     | Transcranial approach                  | Meningitis | Free of symptom |
| Maruya et al., 2002     | 56/female      | Left/1 cm | Bamboo                    | Tumbling            | Temporal (lateral orbit) | Drowsiness       | Left frontotemporal craniotomy and orbitozygomatic osteotomy Frontal craniectomies to repair CSF fistulas | No        | Lateral gaze in the left eye |
| Hayashi et al., 2003    | 71/female      | Right/no foreign body       | Bear claws             | Animal assault      | Frontal sinus       | Lacerated face, CSF fistulas            | No        | Free of symptom |
| Nishio et al., 2004     | 6/female       | Right/2.5 cm | Wooden chopstick          | Falling             | Frontal (orbital roof) | Fever, headache, neck stiffness Crying | Craniotomy | Brain abscess |
| Kim et al., 2005        | 1/male         | Right/2 cm | Metallic chopstick        | Falling             | Frontal (orbital roof) |                      | Craniotomy | Free of symptom |
| Tsao et al., 2006       | 45/male        | Left/2.5 cm * 2.5 cm | Plastic chair glide | Be assaulted | Sinonasal cavity, anterior skull base | Ptosis, impaired vision | Transnasal endoscopic exploration and removal of the foreign body | No | Anosmia and diplopia |
| De Tommasi et al., 2006 | 20/male        | Right/10 cm | Screw driver              | Falling             | Right maxilla       | Scotoma in the left eye, CSF leakage | Left pterional approach | NK | Restored vision |
| Park, 2006              | 9/female       | Right/NK  | Wooden chopstick          | Falling             | Frontal (orbital roof) | Swollen eye       | Craniotomy | No                     |
| Park, 2006              | 1/male         | Left/NK   | Metallic chopstick        | Falling             | Frontal (orbital roof) | Swelling, left hemiparesis | NK | No                     |
| Park, 2006              | 5/male         | Left/NK   | Metallic chopstick        | Falling             | Frontal (orbital roof) | Swollen eye       | NK | No                     |
| Author and year | Age and gender | Side/size | Material | Mechanism of injury | Penetrating pathway | Symptom | Surgical approach | Infection | Outcome |
|----------------|----------------|-----------|----------|---------------------|---------------------|---------|-------------------|-----------|---------|
| Park, 2006     | 2/male         | Left/NK   | Metallic chopstick | Falling | Frontal (orbital roof) | Swollen eye | NK | No | Free of symptom |
| Nitsch et al., 2007 | 22/male       | Right/2 cm | Nail-gun | Accident | Right upper jaw to temporal lobe | Intracerebral haematoma | Simple extraction of the nail without a craniotomy | NK | NK |
| Hiraishi, 2007 | 5/female       | Left/3.5 cm | Plastic chopstick | Falling | Frontal (orbital medial) | Fever, headache, neck stiffness | Craniotomy | Meningitis, brain abscess | Hyposmia |
| Kawada, 2009   | 5/female       | Left/3 cm  | Wooden chopstick | Falling | Frontal (orbital roof) | Swollen eye | Craniotomy | Brain abscess | Free of symptom |
| Mitilian, 2009 | 4/male         | Right/11 cm | Wooden chopstick | Falling | Superior orbital fissure | Mild confusion, vomiting | Craniotomy | Brain abscess | Mild dysmetria of the arm |
| Wieland et al., 2010 | 58/male     | Left-sided/NK | Wooden stick | Accident | Maxillofacial to ethmoid roof | Endoscopic repair of the skull base defect | Intracranial infection | NK |
| Hettige et al., 2010 | 38/female   | Left/NK   | Plastic chopstick | Stumble | Posterior wall of the oropharynx to jugular foramen into the posterior fossa | Nystagmus, left 9th, 10th, and 12th cranial nerves palsy, quadrantanopia | No | NK |
| Sweeney et al., 2011 | 31/male       | Right/NK  | Knife | Suicidal attempt | Lower jaw to anterol skull base | Pain | Combined right periorbital and interhemispheric craniotomies | No | Left 9th, 10th, and 12th cranial nerves palsy |
| Sweeney et al., 2011 | 21/female     | Right/NK  | Knife | Homicidal attempt | Left orbit to middle skull base | Emesis, seizure | Left occipital craniotomy, retrosigmoid craniectomy | No | NK |
| Yonezawa, 2011  | 28/male        | Left/NK   | Plastic chopstick | Falling | Frontal (orbital medial) | NK | Right frontoparietal craniotomy | No | Free of symptom |
| Arslan et al., 2012 | 13/male      | Right/18 cm | Iron bar | Falling | Right orbit and superior orbital fissure | Coma (GCS3) | Removal of chopstick without craniotomy | No | Died |
| Shin et al., 2012 | 38/male       | Left/14 cm | Plastic chopstick | Falling | Superior orbital fissure | Blindness, swelling, numbness around eye | No | Blindness |
| Katayama et al., 2013 | 18/male       | Right/12 cm | Metal rod | Falling | Frontal (subzygomatic bone) | Vomiting | No | No neurological deficit |
| Wang et al., 2013 | 35/male       | Left/18 cm | Steel  | Falling | Left maxillary sinus | Limited movement of neck | Craniotomy | No | Left facial palsy |
Table 2: Continued.

| Author and year          | Age and gender | Side/size | Material            | Mechanism of injury | Penetrating pathway | Symptom               | Surgical approach                      | Infection | Outcome                      |
|--------------------------|----------------|-----------|---------------------|---------------------|---------------------|-----------------------|----------------------------------------|-----------|------------------------------|
| Skoch et al., 2013       | 35/female      | Medial/NK | Electric toothbrush | Accident            | Temporal (lateral orbit) | Blurred vision, pain | Modified frontotemporal orbitozygomatic craniotomy | No        | Improved pain and vision     |
| Deveer et al., 2013      | 43/male        | Right/5.6 cm | Plastic pen        | Falling             | Right orbit         | No                    | No                                     | NK        | Free of symptom              |
| Yamasaki et al., 2013    | 4/female       | Left/6.5 cm | Plastic chopstick  | Falling             | Frontal (orbital roof) | Swelling              | NK                                     | NK        | Free of symptom, encephalitis |
| Williams et al., 2014    | 56/male        | Left/NK   | Speargun            | Suicidal attempt    | Submandibular region, oral cavity | Coma                   | Large craniotomy                      | Cerebritis | Died                         |

ACA, Anterior Cerebral Artery; CSF, Cerebrospinal Fluid; GCS, Glasgow Coma Scale; ICA, internal carotid artery; NK, not known.
Figure 3: Photography showed the entry point (black triangle, △) of foreign body on face and incision for a frontotemporoorbitozygomatic approach (a). Short piece of foreign body (simple arrow, ←) in the face was exposed and removed by a maxillofacial surgeon (b). The infratemporal fossa was opened to expose the long piece of foreign body (c), and it (hollow arrow, ⇑) was removed in a retrograde fashion. During the removing process, yellowish pus (asterisk, ∗) was drained into the infratemporal fossa (d). Photography displayed short piece (simple arrow, ←) and long piece (hollow arrow, ⇑) of the foreign body (e). Postoperative skull radiography suggested complete removal of the foreign body (f). The small bone fragment (hollow arrowhead, ◀) was left in place due to its close relationship with branch of MCA (g). Three-month follow-up MRI revealed no abscess formation (h). The patient recovered uneventfully (i). MCA, middle cerebral artery; MRI, magnetic resonance imaging.

3D modeling has been frequently used in the presurgical planning of PSBIs in previous studies [8, 11, 15–17, 19, 23, 24, 28]. 3D models can be viewed in 360 degrees, rotated, and studied from any perspective [29–31], which facilitate the diagnosis and operation to some extent [30, 31]. 3D Slicer is a free, open source software that can be used for segmentation and 3D modeling with high accuracy and reliability [30, 32]. It has been used in the management of various diseases including intracranial aneurysms, trigeminal neuralgia, and intracerebral hematomas [29, 30, 33]. PSBIs in our case series were reconstructed using 3D Slicer, in which penetrating points as well as the surrounding neurovascular structures
were clearly visualized. It contributed to the presurgical planning of optimal surgical approach and avoiding unexpected vascular injury. Our findings confirmed the feasibility and reliability of 3D Slicer in the modeling of foreign objects and adjacent neurovascular structures. Moreover, the segmentation and modeling procedure using 3D Slicer allowed higher quality of visualization, better view of objects and richer information than workstation reconstruction (Figures 1(g), 2(g)–2(i), and 3(g)) [30]. To our knowledge, this is the first study to visualize PSBIs with 3D Slicer, granting preoperative surgical plan for proper approaches.

4.3. Treatment of PSBIs. Owing to the low incidence of PSBIs, prospective or controlled study is difficult to conduct on the limited cases. Temporary management of PSBIs constantly depends on the experience of different institutions [7, 15]. Despite the availability of some complicated guidelines for penetrating brain injury, they were mostly based on the data of scattered cases without systematic summary for PSBIs [3, 8–10]. Thus, we discussed the treatment of PSBIs based on related cases we treated and previously reported.

Operation is the major strategy for the treatment of PSBIs. Indications for surgery are retained objects, CSF leakage, fracture displacement, intracranial hemorrhage, and vascular injury [34–38]. The purpose of surgery is to remove foreign objects, decompress brain tissue, and reconstruct skull base. Generally, operation is suggested in 12 hours after PSBIs [39, 40]. However, delayed operation is not recommended until full physical and radiological examination are performed since premature surgery might lead to fatal results. Artery injury is one of the common concurrent conditions that should be identified before surgery, the presence of which will be a disaster for emergent operation [16].

In PSBIs with vascular injuries, protection of injured arteries through preoperative endovascular occlusion or intraoperative artery control would be helpful in the removal of foreign bodies [41]. In contrast, in PSBIs without vascular injuries, foreign bodies could be removed directly [7]. For some typical cases, the metal foreign bodies or bone fragments are adjacent to important structures and difficult to extract but cause no obvious symptoms. These foreign bodies could be retained in the brain (Case 2).

Surgical approach for skull base injury should be individualized according to the penetrating trajectory, location of foreign body, and accompanying vascular and brain injuries. In current study, different modifications of frontotemporo-orbitozygomatic approach were adopted to manage the anterior and middle skull base injuries. By removing the superior and lateral bony orbit, we could deal with most of foreign bodies penetrating from the orbit into frontal lobe. The removal of zygomatic arch enables inferior displacement of temporalis muscle, allowing exploration of undersurface of temporal lobe. Generally, principles for skull base surgery can also be applied to PSBIs. Proximal vascular control should be first guaranteed to prevent intraoperative hemorrhage. And direct visualization of foreign object should be achieved before its removal. Usually, it is necessary to drill away the bone of skull base to expose the foreign bodies. Thorough debridement along the exposed trajectory as well as careful reconstruction of the skull base is of great significance to prevent postoperative infection and CSF leakage. However, aggressive debridement for deep seated debris should be avoided, which may be associated with increased disability and mortality [20].

Infection is the main complication of PSBIs with a reported overall rate of 64–70% and mortality rate of 14–57% [42–44]. Organic foreign body like wood or bamboo is not only the carrier, but also the best medium for infection such as brain abscess, meningitis, and cerebritis [36, 45]. In this way, organic foreign bodies should be totally retrieved, while some other deep seated foreign bodies such as small metal or bone fragments could be retained since total extraction would cause more damage. In the absence of sufficient data and definitive guidelines, the type, timing, and duration of antibiotic use remains uncertain, especially when the result of CSF culture is negative. In recent publications, prophylactic use of broad-spectrum antibiotic was suggested within 7–14 days after the injury [16, 39], while others indicated that antibiotic therapy should be administrated according to the findings of CSF culture [46]. We recommended prophylactic use of antibiotics and proper adjustment according to CSF culture results, especially for wooden objects.

5. Conclusions

PSBI is a rare disease with various injury mechanisms and complicated traumatic conditions. Lots of difficulties regarding the diagnosis and management of PSBIs remain to be solved. Based on our experience and review of previous studies, we suggest full physical examination and radiological evaluation before surgery. Preoperative 3D modeling with 3D Slicer could help visualize penetrating pathway and surrounding neurovascular structures in detail, granting free view from any angle and selection of optimal approach. However, caution is needed in interpreting our findings because of the limited cases. Further large scale prospective studies are required to identify the effect of preoperative 3D reconstruction on the prognosis of PSBIs, as well as the guideline for the management of PSBIs.

Conflicts of Interest

The authors declare that there are no conflicts of interest regarding the publication of this article.

Authors’ Contributions

Danfeng Zhang performed data collection, data analysis, and writing of the article. Jigang Chen collaborated in literature search and study design; Kaiwei Han collaborated in figure generation and study design; Mingkun Yu collaborated in study design, data analysis, and editing of the article; Lijun Hou collaborated in literature research, figure generation, and editing of the article.
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