Infections of the vertebral column can result in vertebral osteomyelitis with or without epidural abscess. Vertebral osteomyelitis is a relatively rare condition and comprises 2%–7% of all hematogenous bone infections, but their incidence appears to be increasing in recent years compared with the first half of the 20th century. Only 6.5% of these infections are located in the cervical region, the least commonly affected area of the spine. Spinal epidural abscess is also a rare diagnosis with an incidence of approximately 0.2–2 cases per 10,000 admissions. This condition is defined as a pyogenic infection between the vertebral body and dura mater covering the spinal cord. The epidural space can be seeded with infection by a variety of mechanisms, including hematogenous spread (50%), direct extension from adjacent infection (33%), inoculation from spinal procedures (15%), and other mechanisms. Although *Staphylococcus aureus* causes approximately two-thirds of spinal abscesses, other infecting pathogens, including gram-positive and -negative bacteria and anaerobic bacteria, are possible causative agents.

The rate of misdiagnosis at presentation for vertebral osteomyelitis and epidural abscess is still high (50%). The wide variation in clinical presentations makes it difficult to conduct any large-scale observational studies. Furthermore, such a study would raise ethical concerns, and, consequently, the decision making and management have been largely dependent on empirical experience, animal studies, and literature case reports.

Because of the proximity of the oropharynx (a naturally contaminated region) to the spinal structures of the craniocervical junction, it is possible that small mucosal lacerations in the oropharynx caused by unstable traumatic craniocervical injuries may become contaminated and lead to secondary infection and osteomyelitis. In this report, the authors describe the case of a previously healthy and immunocompetent patient who developed a large retropharyngeal abscess with spinal osteomyelitis after a high-energy craniocervical injury. This unusual report of osteomyelitis with a delayed presentation after a high-energy traumatic injury of the craniocervical junction highlights the possibility of direct injury to a specific area in the oropharyngeal mucosa adjacent to the osteoligamentous structures of the craniocervical junction, an overall underrecognized complication of unstable craniocervical injuries.
We propose in this paper that, because of the proximity of the oropharynx mucosa, a naturally contaminated region, and the osteoligamentous structures of the cranio cervical junction, especially in the area adjacent to the anterior arch of C-1, a secondary infection leading to retropharyngeal abscess formation and resultant contiguous osteomyelitis with epidural abscess formation might be caused by the introduction of microorganisms into the RPS due to minor mucosal lacerations. As an illustration, we report the case of a previously healthy and immunocompetent patient who developed a large retropharyngeal abscess with spinal osteomyelitis and epidural abscess after a high-energy cervical trauma with a subsequent unrecognized unstable cranio cervical injury. To our knowledge, this is the first report that specifically highlights this proposed pathophysiological mechanism.

Case Report

This 50-year-old man with no significant prior medical history was involved in a motor vehicle accident in which he struck a deer at approximately 30 miles per hour. He did not seek medical attention at the time of the accident since he had only mild neck pain. After 5 days his neck pain worsened and he underwent a session of chiropractic manipulations.

The following day the patient developed fevers as high as 102.8°F (39.3°C) and went to an urgent care facility. Because of associated nasal congestion at the time, he was prescribed Bactrim for a supposed sinus infection. His clinical picture worsened with fever, chills, drenching sweats, worsening neck pain, neck weakness, and a subjective truncal weakness.

Around 40 days after the accident and with the progression of his symptoms, the patient went back to the urgent care facility and was advised to go to the emergency department, but he decided not to follow the recommendation. Shortly after, while at home, he developed a sudden onset of weakness in all 4 extremities and was brought by ambulance to an outside hospital for evaluation.

Workup revealed a C-2 fracture with spinal cord compression and findings suggestive of osteomyelitis with epidural abscess (Fig. 1). CT angiography demonstrated vertebral artery compression and occlusion on the right side by the fractured segment level (C2–3). Blood cultures grew methicillin-sensitive S. aureus (MSSA), and the patient was started on a regimen of cefazolin. Additionally, at this same hospital, the patient underwent surgery for posterior decompression and cervical stabilization performed with the placement of 2 short screws, a transforminal lumbar interbody fusion (TLIF) cage between the spinous process of the oropharynx mucosa, and wiring that led to coronal deformity (Fig. 2A–D).

On postoperative Day 9, while still in the hospital, the patient’s neck slouched forward while he was seated and he developed sudden-onset shortness of breath and quadriplegia. The patient passed out, and when he was laid down he was able to regain consciousness and breathe. The patient was then transferred to our emergency department with Grade 0/5 strength in all extremities, with the exception of minimal strength in the left leg. A new imaging examination with cervical MRI demonstrated worsening of the epidural abscess extending from the clivus to C3–4 levels and diffuse cord edema from the cranio cervical junction to C-6 along with C1–2 subluxation with a left lateral mass displacement of 9 mm. Severe bone irregularity was noted, suggesting severe osteomyelitis in the occipital condyles and C1–2 vertebral. There was also enhancement extending into the paraspinal soft tissues consistent with abscess with a ring-enhancing collection from the occipital region to the C-6 level. The patient was then admitted to the neurosurgery department for further workup.

During the hospitalization, the patient developed several complications related to dysautonomia, including hypoxia, acute respiratory insufficiency, and a postoperative deep vein thrombosis in the right upper extremity. For the MSSA bacteremia and spine osteomyelitis, the antibiotic regimen was changed to intravenous Unasyn and intravenous vancomycin for a total period of 6 weeks. According to infectious disease recommendations, the patient should be initially treated with 7 days of intravenous antibiotic therapy before being submitted to surgical reintervention to reduce the risks of infection during insertion of the new hardware. On the 5th day after admission, the patient was placed in cervical traction with the weight increased in a slow fashion (increase of 5 lbs every 12 hours with sequential radiographs) to obtain reduction of the deformity (Fig. 3).
After obtaining significant reduction of the deformity, the patient underwent surgical intervention 9 days after admission. The old construct was removed, and an occiput to C-4 arthrodesis with fusion was performed with open reduction and correction of the deformity between the occiput and C-2 (Fig. 4). An intraoperative CT angiogram utilizing the O-Arm was performed to ensure that the remaining vertebral artery was patent.

Postoperative radiographs and CT scans demonstrated optimal reduction of both the sagittal and coronal imbalance (Fig. 5). Following surgery, the patient was kept in an external halo orthosis for 8 weeks after discharge. The halo was removed after the findings of upright radiographs with and without the halo were similar and a CT scan demonstrated fusion in progress. At the last follow-up 5 months after surgery, the patient had regained full strength in all his extremities (Fig. 6A–C). The most recent MRI study demonstrated reduction in the size of the abnormal T2 signal in the spinal cord (Fig. 6D).

**Discussion**

The understanding of the anatomy of the RPS and its relationship with the osteoligamentous structures in the cervical area is essential to understand our proposed pathophysiological mechanism.

The RPS is bordered anteriorly by buccopharyngeal fascia and posteriorly by prevertebral fascia. The RPS can be divided into an anterior and a posterior compartment by the thin alar fascia that comes from the deep cervical fascia. The anterior compartment extends from the skull base to the T-4 vertebral body. The posterior compartment extends inferiorly to the level of the diaphragm. Head and neck infections might spread to the posterior mediastinum through the RPS. The RPS can be further divided relative to the hyoid bone, with the suprahyoid space containing retropharyngeal lymph nodes and fat, while the infrahyoid area contains only fat.48

The RPS has a very close relationship with the cranio-cervical junction, a complex articulation between the cranium and upper cervical spine (C1–2) with several bony and ligamentous structures, including the transverse atlantal ligament, paired alar ligaments, and the apical ligament of the dens. Lesions at this articulation can lead to hematoma formation into the RPS. The main anatomical structures that could be sources of hematoma when injured are the longus capitis and longus colli muscles, the anterior longitudinal ligament, and the vertebral artery branches.

In particular, the anterior arch of C-1 is easily accessible through the retropharyngeal mucosa, as can be seen in our cadaveric endoscopic endonasal dissection (Fig. 7). We performed this dissection on a de-identified cadaveric specimen at the Anatomy Laboratory Toward Visuospatial Surgical Innovations in Otolaryngology and Neurosurgery at the Wexner Medical Center at The Ohio State University (OSU), which is certified by regulatory agencies in dealing with the use of human tissues and cadaveric studies. Our dissection highlights the very close relationship with the oropharynx mucosa, an area inhabited not only by normal flora but also potentially by virulent...
bacteria. Thus, minor mucosal lacerations might provide a possible path for bacterial invasion and subsequent abscess development in the RPS given a setting of a high-energy craniocervical injury, as demonstrated in our dynamic illustration (Fig. 8)\(^3,4^8\).

In patients with degenerative changes in the spine, the risk of a mucosal tear might be even greater. Robinson et al. described a case of perforation of the posterior pharyngeal wall with abscess formation by an anterior marginal osteophyte at the time of hyperextension injury of the cervical spine. The abscess communicated with the epidural space via the disrupted intervertebral disc, leading to the delayed onset of upper airway obstruction and tetraparesis.\(^3^5\)

Another interesting case reported by Wadie et al. describes a patient with a sewing pin impaction in the laryngeal introitus, with late development of cervical spondylodiscitis after an uneventful removal of the foreign body. The authors concluded that the patient had a microperforation of the posterior pharyngeal wall that was not visible during the endoscopy procedure performed at his initial evaluation.\(^5^0\) This conclusion supports that even minor trauma to the posterior oropharyngeal wall might be a source of infection to the retropharyngeal space and, consequently, the spine.

Additionally, cases in the literature confirm that the forces acting in a whiplash injury could be strong enough to cause esophageal perforation.\(^4^9\) In 1960, Morrison described a patient in whom hyperextension of the cervical spine had ruptured the esophagus; the patient died of me-
diastinitis and bronchopneumonia 2 weeks after injury. In 1976, Splener and Benfield reported on a patient with esophageal trauma in whom the delayed diagnosis and treatment contributed to the development of an abscess that required surgical drainage. In this context, it is also feasible that a microrupture to the fragile retropharyngeal mucosa specifically located in the area adjacent to the anterior arch of C-1 could occur after whiplash injuries.

Moreover, esophageal perforation after radiation and chemotherapy is described as a possible cause of spinal abscess, and one of the proposed mechanisms is via a direct extension through a fistula formed between the pharynx and the prevertebral fascia due to radiation necrosis with posterior extension to the spine, potentially causing osteomyelitis and abscess formation. Even small traumatic lesions such as the one caused by a fish bone are described as possible sources of infection to adjacent areas.

Retropharyngeal hematoma, which is a known possible complication of trauma described in the literature, is usually treated conservatively with observation. It is possible...
that if there is a microlesion to the oropharyngeal wall with bacteria inoculation, the hematoma might function as a medium of culture for the bacteria to grow and lead to an abscess with posterior involvement of the cervical spine and epidural space.  

In regard to retropharyngeal abscesses, the most common cause is bacterial infection that originates from the nasopharynx, middle ear, sinuses, or tonsils. Because the RPS lymph nodes degenerate during adolescence, there is a decreased incidence of spontaneous retropharyngeal abscess in adults. Nonetheless, if left untreated, these abscesses can lead to additional complications, such as thrombosis, airway obstruction, pericarditis, and rarely vertebral osteomyelitis. When vertebral osteomyelitis occurs, serious consequences such as vertebral bone destruction and epidural abscess may ensue.

To date, we could find no description in the literature about oral mucosal perforation in the thinner portion of this structure located in the area adjacent to the anterior arch of C-1 as a cause of retropharyngeal abscess and subsequent osteomyelitis. In addition, whereas patients who develop retropharyngeal abscess usually have some degree of immunodeficiency, our patient was young and immunocompetent.  

Infection Progression and Craniocervical Instability

There are strong clinical and biomechanical studies supporting that soft-tissue injuries of the cervical spine can lead to instability, which could have been the cause of our patient’s unstable craniocervical injury that was previously not recognized. Subacute instability of the cervical spine has been reported in patients presenting with neck pain and normal findings on radiographs who subsequently developed clinical instability, most likely due to ligamentous injury. In vitro investigations have demonstrated that transection or injury to the anterior longitudinal ligament (ALL) and anterior annulus resulted in increased flexibility under extension loading, implying that injury to these components could lead to instability. Ivancic et al. found that the ruptured ligaments were associated with significantly greater dynamic intervertebral extension, peak ligament strain, and joint laxity compared with the uninjured ligaments.

We also agree that these lacerations in the ALL could...
serve as the path through which the infection could progress to osteomyelitis and epidural abscess. Harris and Yeakley reviewed the MR images obtained in 8 adults with acute hyperextension-dislocation of the cervical spine. The images were obtained to evaluate damage to the spinal cord. All 8 patients had disruption of the ALL and of the annulus of the intervertebral disc.11 Similar injuries have also been discovered at surgery and autopsy.7,19 Whiplash simulations using cadavers and monkeys have also produced ALL tears and anterior disc detachments.10,41,54 Along these lines, a meta-analysis by Reihausa et al. verified that closed trauma might play a role in spinal epidural abscess since about 10% of patients with spinal epidural abscess of unknown origin had a history of extraspinal or spinal trauma.30

Moreover, destruction of bone caused by the osteomyelitis itself contributes to the craniocervical instability. The vascular spread of bacteria and microthrombosis due to pus creates an increased intraosseous pressure that impairs the blood flow to the vertebrae and intervertebral discs.22,51,52 This may lead to ischemic necrosis and destruction of the vertebrae and intervertebral discs with consequent spinal deformity. Another proposed mechanism for bone destruction is through the production of destructive enzymes by bacterial agents such as *S. aureus*. The most important of these proteolytic enzymes is hyaluronidase, which is used by the bacteria as an invasive mechanism into the connective tissue. It causes destruction of structures such as the annulus fibrosus, leading to biomechanical instability and deformity.18,44 In a cohort published by Srinivasa et al., the overall risk of developing a deformity with osteomyelitis was 44% (21/48). Of these 21 deformities, 14 involved kyphosis only, 5 involved subluxation only, and 2 involved both kyphosis and subluxation.44

Infectious lesions within the RPS and extending to the spine are difficult to diagnose clinically; the first symptoms to appear are unspecific, and a high level of suspicion is necessary to make the diagnosis. For instance, our patient presented initially with vague symptoms that progressed to weakness, falls, 30-lb weight loss, spasms, headaches, finger numbness, fevers, and chills. As in our case, patients present with more specific symptoms only at a later stage when the degree of spread is advanced, and neurological deficits and the destruction of the vertebral bodies due to osteomyelitis could be already permanent.

Despite spinal epidural abscess (SEA) being a rare condition, delay in diagnosis may lead to permanent neurological deficit, sepsis, and even death. The mortality of SEA has been reported to be as high as 16%.10,34,37 A study by Tang et al. in 2002 revealed that the initial accurate diagnosis rate was as low as 26%.46 Unfortunately, as many patients are initially misdiagnosed, they will have a poor prognosis; therefore, these patients should be assessed in detail.

According to the available literature, the main symptom of the SEA is pain, followed by common signs of infection, such as fever and malaise. The classic diagnosis of SEAs with mechanical pain, tenderness, fever, and neurological abnormalities is likely only enough to capture a maximum of 10% of patients. According to a study by Davis et al. the typical clinical presentation of retropharyngeal abscess is acute to subacute onset of neck pain, with or without associated dysphagia and odynophagia, which might lead to weight loss—as in our case—and low-grade fever.6 Less frequently, patients might present with clearerer symptoms, such as airway distress, high-grade fever (> 101°F), drooling, trismus, limited neck motion, head tilt or torticollis, and noisy breathing.31

In this unclear clinical setting the key diagnostic tools are the imaging findings. Plain radiographs are more likely to show normal findings in the acute phase, although soft-tissue changes may be present. Bony changes on plain radiographs are usually present in chronic infections. CT is also helpful in evaluating the amount of bone involvement. Nonetheless, contrast-enhanced MRI is the modality of choice for identifying the spinal infection, because of its excellent visualization of the abscess itself, and the infection foci in the vertebrae, disc space, or paraspinal region.4,22,38

In our case the patient had not only an unstable craniocervical injury but also the presence of a purulent bacterial empyema or phlegmon in the epidural space overlying the spinal cord. Therefore, this case was considered a surgical emergency with the intent to decrease the likelihood of neurological deficits caused by infection progression. Studies in the literature decades ago demonstrated in rabbits that diffuse thrombosis of the underlying vasculature is the probable cause of neurological decline in this condition.8,9

Although surgical treatment of neurologically intact patients with vertebral osteomyelitis alone or minimal epidural abscess is quite controversial, vertebral osteomyelitis and epidural abscess associated with neurological deficits are considered surgical emergencies.1,6,24,42,53 Since local inflammation leading to thrombophlebitis or venous congestion is thought to be a possible mechanism of neurological injury, surgical debridement is useful to remove the inciting organisms and factors to reduce the risk of vascular-related neurological complications.8

Aggressive treatment of the prevertebral abscess is the key to decrease mortality; the most recommended management would be surgical drainage, followed by systemic antibiotics. The patient’s neurological status just after surgery is the main prognosis tool.39 About 5% of the patients with spinal epidural abscess die, usually of progressive sepsis, consequent meningitis, and complications due to underlying diseases.6 In our case, the patient experienced a severe neurological deficit while he was in the hospital, which allowed prompt management and recovery of his neurological function despite the complications encountered.10

Conclusions

Retropharyngeal abscess leading to spinal osteomyelitis is usually a very challenging clinical situation, especially in the setting of craniocervical instability and secondary deformity. By presenting an illustrative surgical case, this article highlights the relationship between a routine anatomical finding (the short distance between the oropharyngeal mucosa and the anterior arch of C-1) and the possibility of osteomyelitis occurring after high-en-
ergy craniocervical trauma. The pathophysiology of such injury would involve direct disruption of the oropharyngeal mucosa by the unstable osseous structures, leading either to a direct communication with the oropharyngeal space or to the formation of a retropharyngeal hematoma that may become infected. Ultimately, early diagnosis and treatment of vertebral osteomyelitis is highly dependent on physicians’ awareness of such potential complications of craniocervical junction trauma, which, in our opinion, has been highly underrecognized in the spinal literature. Finally, this article emphasizes the necessity of further combined histological/biomechanical studies to evaluate the effects of progressive levels of craniocervical instability upon the integrity of the oropharyngeal mucosa.

References

Disclosure
The authors report no conflict of interest concerning the materials or methods used in this study or the findings specified in this paper.

Author Contributions
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