



# TREATMENT OF DESCENDING NECROTIZING MEDIASTITIS ASSOCIATED WITH DEEP NECK INFECTIONS: AN ACCOUNT OF THE MOST CURRENT DATA

Antonio Mancini<sup>†</sup>, Angelo Michele Inchingolo<sup>†</sup>, Danilo Ciccarese<sup>\*</sup>, Francesco Inchingolo<sup>\*</sup> and Gianna Dipalma

Department of Interdisciplinary Medicine, University of Bari "Aldo Moro", Bari, Italy.

<sup>†</sup> These authors contributed equally as first authors

*\*Correspondence to:*

Francesco Inchingolo,  
Department of Interdisciplinary Medicine,  
University of Bari "Aldo Moro",  
70124 Bari, Italy.  
Email: [francesco.inchingolo@uniba.it](mailto:francesco.inchingolo@uniba.it)

Danilo Ciccarese,  
Department of Interdisciplinary Medicine,  
University of Bari "Aldo Moro",  
70124 Bari, Italy.  
Email: [danilo.ciccarese@uniba.it](mailto:danilo.ciccarese@uniba.it)

## ABSTRACT

Deep neck infections (DNIs) are life-threatening diseases that represent the crossroads between maxillofacial surgery and otorhinolaryngology. A DNS has the potential to extend into the mediastinum, where it can trigger a very dangerous disease known as descending necrotizing mediastinitis (DNM), which has a high mortality rate. This article reviews the etiology, bacteriology, clinical manifestations, diagnostic techniques, and management approaches for DNIs and DNM. We emphasize the importance of early diagnosis by imaging studies, such as computed tomography (CT), and a multidisciplinary approach utilizing antibiotics coupled with surgical intervention for the management of these life-threatening diseases.

**KEYWORDS:** *Deep neck infection, descending necrotizing mediastinitis, diagnosis, multidisciplinary treatment, surgical treatment*

## INTRODUCTION

Deep neck infections (DNIs) are significant emergencies that can easily spread to the mediastinum, a complication referred to as descending necrotizing mediastinitis (DNM). The cervical fascial planes play a crucial role in the spread of these DNIs (1,2). Thus, knowledge of the anatomy of these fascial planes is crucial for the successful diagnosis and treatment of DNIs.

DNIs can be caused by odontogenic infections, pharyngitis, salivary gland infections, trauma, or foreign bodies (3-5). These latter infections spread into the mediastinum when they overcome the natural resistance of the fascial planes. Cellulitis, abscess, and phlegmon are different stages of the infection and each has a different management approach (4,5).

Received: 21 October, 2024  
Accepted: 15 November, 2024

1972-6945 (2024)  
Copyright © by Biolife-Publisher

This publication and/or article is for individual use only and may not be further reproduced without written permission from the copyright holder. Unauthorized reproduction may result in financial and other penalties. Disclosure: all authors report no conflicts of interest relevant to this article.

The cervical fascia can be superficial and deep, creating retropharyngeal and parapharyngeal spaces, among others, that may become infected and serve as pathways for the dissemination of infection. Knowledge of such structures is important to the clinicians for the planning of appropriate treatment modalities and preventing further complications (5,6).

## DISCUSSION

### *Etiopathology*

Odontogenic and pharyngeal infections are the most common causes of DNIs, though there are other causes too, including trauma, cancer, and iatrogenesis following operations such as endoscopy (7-9). The infection usually extends to the salivary gland, buccal, pharyngeal, and dental mucosa from the initial site by hematogenous or lymphatic routes of dissemination (10,11). The submandibular space is reported to be the most commonly affected space, followed by the parotid and sublingual spaces in many cases (11,12).

DNI usually presents with a mixed bacterial flora comprising both aerobic and anaerobic bacteria (13-15). The most common pathogens found are *Streptococcus viridans* and *Staphylococcus aureus*, but in immunocompromised hosts, like in the case of diabetics, and human immunodeficiency virus (HIV) and cancer patients, *Klebsiella* and other Gram-negative bacteria are also prevalent (16,17). Because these diseases are serious and take root with rapid growth, affected patients must be admitted to a medical facility immediately. DNM is a condition that is present in 3% of DNI cases and comes with a mortality rate of 50%, which is due to inadequate drainage and delayed diagnosis following untreated DNIs (18-21) (Table I).

**Table I.** A literature review of the etiological factors of mediastinal involvement of deep neck infection (DNI).

	Mithos et al. 2007	Roccia et al. 2007	Ridder et al. 2010	Kocher et al. 2012	Celakovsky et al. 2014	Kimura et al. 2020	Gehrke et al. 2022	Ho et al. 2022
Odontogenic	63%	39,1%	11,1%	5,9%	40%	5%	15,56%	38%
Pharyngo-tonsillar	37%	60,9%	46,7%	82,3%	33,5%	33%	55,56%	19,1%
Retropharyngeal abscess	-	-	-	-	-	5 %	-	19,1%
Other	-	-	42%	-	6,5%	18%	28,88%	-
Unknown	-	-	-	11,8%	20%	39%	-	23,8%

### *Clinical Diagnosis*

The most common symptoms of deep throat infections include neck pain and swelling, although fever and odynophagia are frequent in diverse studies (22,23). Otagia, dysphonia, dysphagia, trismus, and dyspnea are some symptoms that are specific to the affected area. Pain, often dental or swallow-related, is an early symptom. Acute pain and dysphagia occur when the infection is at an advanced stage (24-26). Dysphonia and dyspnea may accompany pharyngeal and laryngeal infections with a sudden decline in the patient's status. More severe cases can also demand tracheotomy because limited mobility and trismus occur as a result of swelling and infiltration of neck tissues. Fever is also commonly present, although immunocompromised patients may show no fever despite significant invasion of tissue (26-28).

It is commonly observed that inadequately treated infections of the cervico-mediastinal may result in serious sequelae like dyspnea, spontaneous fistulization, and mediastinitis. Mediastinum infection produces symptoms such as erythema of the skin, respiratory distress, and toxic-septic shock. Other adverse effects include low oxygen saturation, tachycardia, hypertension or hypotension, and sweating (29,30). If left untreated, many fascial spaces involved in cervical suppuration can result in severe pneumonia, cardiac issues, and renal failure, all of which carry a significant mortality rate.

A physical examination is done by examining the neck and palpating it to observe soreness, edema, and erythema (31,32). A buccopharyngoscopy evaluates the airway, throat, and mouth. The examination can diagnose edema, erythema, and dental problems; the airway can only be evaluated with fiberoptic laryngoscopy. Tracheotomy or cricothyroidotomy may be necessary in severe cases due to the possibility of respiratory arrest, which causes symptoms such as dysphagia,

toxicity, fever, and stiff neck. If the infection extends to the mediastinum, mediastinitis with excruciating chest pain could lead to mortality (32-34).

Symptoms related to infections in the lateral pharyngeal spaces include fever, pain, rigidity, and possibly trismus, depending on the compartment involved. While the posterior compartment has mild swelling, the anterior compartment is infected, producing symptoms of infection and neck pain from muscle spasms. Common systemic manifestations are results of infection of the parotid gland (eg. swelling), while severe ones can include jugular vein thrombosis and erosion of the carotid artery, among others. If left untreated, necrotizing fasciitis is a dangerous infection which can spread along the plane of a tissue and cause tissue death and a sepsis systemically (35-37).

Dysphagia, chest pain, and fever are the early symptoms most frequently encountered in case of perforations from trauma of the crico-pharynx. Other signs such as subcutaneous emphysema, tachycardia, and respiratory distress may be encountered during the clinical examination (38-40). Although the initial symptoms in deep throat infection are moderate, diagnosis management must consider the presence of comorbid conditions such as diabetes, HIV, cancer, and autoimmune diseases. These are the groups of patients with increased risk of serious complications (41-45).

### *Paraclinical Diagnosis*

Laboratory tests include leukocyte count, C-reactive protein (CRP), erythrocyte sedimentation rate (ESR), and renal function, all of which are very important in delineating the extent of infection and guiding treatment. Imaging techniques are necessary to identify the extent of DNI, especially computed tomography (CT) (45-47). Compared with CT, magnetic resonance imaging (MRI) has the disadvantage of being more expensive and having longer scanning times. However, it yields high-resolution soft tissue images (48-50). CT scans also have their own shortcomings, with a false positive rate of about 10%, and a false negative rate of about 13%. MRI is preferred in complicated cases that might involve vascular abnormalities. Although chest X-rays are useful in the differential diagnosis of conditions like pneumonia and pleural effusion, CT remains the standard when mediastinal involvement is to be reviewed (50,51).

### *Treatment of Deep Neck Infection (DNI) Associated with Descending Necrotizing Mediastinitis (DNM)*

#### *1. Airway Management*

One of the major issues in DNI is the acute obstruction of the airways; this is particularly critical when more than one site is involved with the infection (52-54). Many times, tracheotomy or orotracheal intubation is required (55,56). When this is not possible by any of the above methods, emergency cricothyrotomy is the method of choice for securing the airway. Due to the inherent risks involved for the airway, the management of these conditions requires a multidisciplinary team of physicians specializing in ear, nose, and throat (ENT), anesthesia, and critical care (57-60).

#### *2. Medical Treatment*

As soon as possible, empirical antibiotic therapy should begin, taking a broad spectrum of bacteria in consideration, until the results of the culture allow for a more focused approach (61-63).

Generally used regimes include penicillin plus beta-lactamase inhibitors or cephalosporins plus metronidazole or clindamycin for anaerobic coverage. In cases suspected to involve methicillin-resistant *Staphylococcus aureus* (MRSA) or when the patient is immunocompromised, the addition of Vancomycin is suggested (64). Gentamicin is often added to cover for Gram-negative bacteria such as *Klebsiella* in diabetic patients. Dosages of antibiotics are modified once the culture and sensitivity results are available (65).

#### *3. Directed Antibiotic Therapy*

Short-term use of steroids is utilized to control airway inflammation and reduce edema (66). Generally, this is not advisable for diabetic patients due to the risk of hyperglycemia. Supportive care treatment in a critical care setting includes oxygen therapy, fluid control, and monitoring (66-68).

#### *4. Systemic Therapy*

As soon as possible, empirical antibiotic therapy must begin with a broad spectrum of bacteria in mind, until the results of the culture lead to a narrower approach (61-63).

#### *5. Surgical Treatment*

The appropriate surgical intervention in DNIs, which can be life-threatening, requires an understanding of the complex anatomy of the neck spaces (68,69).

The site of infection gives indications about the microorganisms involved and the source of infection, thereby helping aid empirical antibiotic therapy. The extension of infection to many cervical compartments necessitates aggressive surgical draining and debridement. Drainage usually entails incisions in the area of the sternocleidomastoid muscle (64); CT imaging studies are necessary to position the tissues that are involved. If the disease process involves the muscles of the peripharynx or perilarynx, drainage must be very carefully carried out with the fingers (70).

#### 6. Minimally Invasive Surgical Treatment

The role of minimally invasive procedures like drainage and radiologically guided aspiration versus non-surgical treatment remains controversial (70-72).

For patients who have a well-defined unilocular abscess with no compromised airway, the percutaneous echo-guided drainage is a useful treatment modality. Some of the advantages of minimally invasive procedures such as ultrasonography and CT-guided needle aspiration include less scarring and faster healing. This approach is more frequent in cases involving minors (73). However, in adults, if the surgery is delayed longer than two days after admission to the hospital, the morbidity and mortality rate is higher (74-76).

#### 7. Surgical Treatment by External Incisional Approach

The external cervical route is usually followed, especially in deep infections of the retropharyngeal space. Reasons to perform surgery include obstruction to the airway, septicemia, and failure of improvement after 48 hours of antibiotic treatment (77,78). Deep space infections are subjected to surgical drainage from the outside; single-space abscesses are treated with minimum surgery. Tracheal compression causing severe respiratory failure may require tracheotomy with local anesthetic (79-81).

Saline irrigation inhibits infection when the infected areas are left open for investigation and further debridement after the surgery. Extensive incisions and drainage may facilitate the creation of a more favorable site for healing in cellulitis cases which present various spaces. Some authors also propose that independent incisions be carried out only for drainage, as well as tracheotomy, to prevent the extension of infection to the mediastinum (82-84) (Table II).

**Table II.** Comparison of Surgical Techniques.

Technique	Indication	Advantage	Disadvantage
Open Drainage	Multi-space involvement	Thorough exploration	Invasive
Minimally invasive	Single, localized abscess	Faster recovery	Limited application

#### Multidisciplinary Approach and Complications

The management of DNIs, especially complicated cases or in patients with comorbidities, involves input from experts in ENT, maxillofacial surgery, infectious diseases, radiology, thoracic surgery, and intensive care (84,85). The experts almost all agree that surgical drainage is indicated in the effective management of these infections for the procurement of microbiological samples for further treatment (86,87).

Laryngeal infections can extend to involve the thyroid gland and may require surgical drainage. In Ludwig's angina, securing the airway and commencing antibiotics often requires tracheotomy. If there are no collections noted on a CT, then it is possible to manage this condition with antibiotic therapy alone; otherwise, surgery becomes necessary (87,88).

In cases of lateral pharyngeal infections, drainage is usually performed in order to avoid further complications. Retropharyngeal infections may need a transcervical approach or may require a joint ENT-thoracic team for their surgical management. Such infections tend to extend to vital structures such as the meninges or lungs. Therefore, the mortality rate associated with them can be quite high if there is any delay in management (88-90).

#### Descending Necrotizing Mediastinitis (DNM)

Failure to drain DNIs may lead to rapid deterioration and sepsis. DNM necessitates urgent multidisciplinary management involving ENT, thoracic surgery, and intensive care specialists. Early interventions, including airway protection and drainage, are critical to reduce complications (90-93). There are various approaches to mediastinal drainage; the video assisted thoracic surgery (VATS) approach is always preferred because it gives a very good view and is minimally traumatic, though at the cost of some risk of pleural contamination (53,93-97) (Table III).

**Table III.** Classification and Treatment of Deep Neck Infections (DNIs).

Type of infection	Description	Treatment
Cellulitis	Inflammation without suppuration	Antibiotics and monitoring
Abscess	Localized suppurative infection	Surgical drainage and antibiotic therapy
Phlegmon	Diffuse infection without boundaries	Surgical drainage and antibiotic therapy

## CONCLUSIONS

Most DNI and DNM lesions originate in the otorhinolaryngological (ORL) and oro-maxillofacial (OMF) areas. Mucosal contamination, which is mucosal damage due to endoscopic procedures or orotracheal intubations, results primarily from dental, buccal, pharyngeal, laryngeal, submaxillary, or parotid glandular inflammatory origins but also from iatrogenic sources. A lymphatic or hematogenous route spreads the infection.

Airway evaluation must be performed first, and any sign of respiratory distress or imminent airway compromise must be considered an emergency to be aggressively treated. This is followed by empirical broad range antibiotic treatment, with adjustments based on the reports of bacterial culture and sensitivity. Because of advances in anesthesia, antibiotics, surgery, diagnostic techniques, and intensive care protocols, the death rate among patients with DNI and DNM has been significantly reduced within the past 20 years.

Appropriate intensive and early therapy can be initiated from the findings in CT to prevent the infection from spreading into DNM. This may be followed by a dynamic CT follow-up at 48–72 hours after the initial surgical operation, which may show the need for a secondary procedure to drain any residual abscess or to remove devitalized tissue. When DNM is treated with DNI treatment by a multidisciplinary team, sepsis, death, and major sequelae are reduced.

### Author Contributions

Conceptualization, A.M., F.I., A.M.I., D.C. and G.D.; methodology, A.M.I., D.C., G.D., and A.M. software, F.I., G.D., A.M.I., D.C. and A.M.; validation, F.I., A.M.I., G.D., A.M. and D.C. formal analysis, A.D.I., A.M.I., A.M., D.C. and F.I.; investigation, A.M., D.C., A.M.I., F.I. and G.D.; resources, A.M., A.M.I., D.C., F.I. and G.D.; data curation, G.D., D.C., A.M., A.M.I. and F.I.; writing original draft preparation, A.M., A.M.I., D.C. and F.I.; writing review and editing, F.I., A.M., D.C., A.M.I. and G.D.; visualization, D.C., A.M., A.M.I., F.I. and A.M.I.; supervision, G.D, F.I., A.M., A.M.I. and D.C.; project administration, A.M., D.C., F.I., A.M.I. and G.D.. All authors have read and agreed to the published version of the manuscript.

### Conflict of interest

The authors declare that they have no conflict of interest.

## REFERENCES

- Hedge A, Mohan S, Eng W. Infections of the deep neck spaces. *Singapore medical journal*. 2012;53(5):305-311.
- Li RM, Kiemeny M. Infections of the Neck. *Emergency Medicine Clinics of North America*. 2019;37(1):95-107. doi:<https://doi.org/10.1016/j.emc.2018.09.003>
- Velhonoja J, Lääveri M, Soukka T, Irjala H, Kinnunen I. Deep neck space infections: an upward trend and changing characteristics. *European Archives of Oto-Rhino-Laryngology*. 2019;277(3):863-872. doi:<https://doi.org/10.1007/s00405-019-05742-9>
- Garner DH, Kortz MW, Baker S. Anatomy, Head and Neck: Carotid Sheath. In *StatPearls*; StatPearls Publishing: Treasure Island (FL), 2024.
- Reuter TC, Korell V, Pfeiffer J, Ridder GJ, Ketterer MC, Becker C. Descending necrotizing mediastinitis: etiopathogenesis, diagnosis, treatment and long-term consequences—a retrospective follow-up study. *European archives of oto-rhino-laryngology/European archives of oto-rhino-laryngology and head & neck*. 2022;280(4):1983-1990. doi:<https://doi.org/10.1007/s00405-022-07769-x>

6. Varghese J, Kirsch C. Magnetic Resonance Imaging of the Oral Cavity and Oropharynx. *Topics in Magnetic Resonance Imaging*. 2021;30(2):79-83. doi:<https://doi.org/10.1097/rmr.0000000000000282>
7. Misthos P, Katsaragakis S, Kakaris S, Theodorou D, Skottis I. Descending Necrotizing Anterior Mediastinitis: Analysis of Survival and Surgical Treatment Modalities. *Journal of oral and maxillofacial surgery : official journal of the American Association of Oral and Maxillofacial Surgeons*. 2007;65(4):635-639. doi:<https://doi.org/10.1016/j.joms.2006.06.287>
8. Ridder GJ, Maier W, Kinzer S, Teszler CB, Boedeker CC, Pfeiffer J. Descending Necrotizing Mediastinitis. *Annals of Surgery*. 2010;251(3):528-534. doi:<https://doi.org/10.1097/sla.0b013e3181c1b0d1>
9. Kocher G, Hokschi B, Caversaccio M, Wiegand J, Schmid R. Diffuse Descending Necrotizing Mediastinitis: Surgical Therapy and Outcome in a Single-Centre Series. *European journal of cardio-thoracic surgery : official journal of the European Association for Cardio-thoracic Surgery*. 2012;42:e66-72. doi:<https://doi.org/10.1093/ejcts/ezs385>.
10. Inchingolo AD, Inchingolo AM, Bordea IR, et al. SARS-CoV-2 Disease through Viral Genomic and Receptor Implications: An Overview of Diagnostic and Immunology Breakthroughs. *Microorganisms*. 2021;9(4):793. doi:<https://doi.org/10.3390/microorganisms9040793>
11. Kimura A, Miyamoto S, Yamashita T. Clinical predictors of descending necrotizing mediastinitis after deep neck infections. *The Laryngoscope*. 2019;130(11). doi:<https://doi.org/10.1002/lary.28406>
12. Gehrke T, Scherzad A, Hagen R, Hackenberg S. Deep neck infections with and without mediastinal involvement: treatment and outcome in 218 patients. *European Archives of Oto-Rhino-Laryngology*. 2021;279(3):1585-1592. doi:<https://doi.org/10.1007/s00405-021-06945-9>
13. Ho CY, Chin SC, Chen SL. Management of Descending Necrotizing Mediastinitis, a Severe Complication of Deep Neck Infection, Based on Multidisciplinary Approaches and Departmental Co-Ordination. *Ear Nose & Throat Journal*. 2022;103(9):014556132110685-014556132110685. doi:<https://doi.org/10.1177/01455613211068575>
14. Inchingolo F, Tatullo M, Abenavoli FM, et al. Comparison between traditional surgery, CO2 and Nd:Yag laser treatment for generalized gingival hyperplasia in Sturge-Weber syndrome: a retrospective study. *Journal of Investigative and Clinical Dentistry*. 2010;1(2):85-89. doi:<https://doi.org/10.1111/j.2041-1626.2010.00020.x>
15. Brajkovic D, Zjalic S, Kiralj A. Prognostic factors for descending necrotizing mediastinitis development in deep space neck infections—a retrospective study. *European Archives of Oto-Rhino-Laryngology*. 2021;279(5):2641-2649. doi:<https://doi.org/10.1007/s00405-021-07081-0>
16. Qu L, Xu H, Liang X, Cai X, Zhang W, Qian W. A Retrospective Cohort Study of Risk Factors for Descending Necrotizing Mediastinitis Caused by Multispace Infection in the Maxillofacial Region. *Journal of Oral and Maxillofacial Surgery*. 2020;78(3):386-393. doi:<https://doi.org/10.1016/j.joms.2019.11.017>
17. Ord R, Coletti D. Cervico-facial necrotizing fasciitis. *Oral Diseases*. 2009;15(2):133-141. doi:<https://doi.org/10.1111/j.1601-0825.2008.01496.x>
18. Inchingolo F, Tatullo M, Pacifici A, et al. Use of dermal-fat grafts in the post-oncological reconstructive surgery of atrophies in the zygomatic region: clinical evaluations in the patients undergone to previous radiation therapy. *Head & face medicine*. 2012;8(1). doi:<https://doi.org/10.1186/1746-160x-8-33>
19. Marioni G, Staffieri A, Saverio Giuseppe Parisi, et al. Rational Diagnostic and Therapeutic Management of Deep Neck Infections: Analysis of 233 Consecutive Cases. *Annals of Otolaryngology, Rhinology, and Laryngology*. 2010;119(3):181-187. doi:<https://doi.org/10.1177/000348941011900306>
20. Chi TH, Tsao YH, Yuan CH. Influences of Patient Age on Deep Neck Infection. *Otolaryngology–Head and Neck Surgery*. 2014;151(4):586-590. doi:<https://doi.org/10.1177/0194599814542589>
21. Hidaka H, Yamaguchi T, Hasegawa J, et al. Clinical and bacteriological influence of diabetes mellitus on deep neck infection: Systematic review and meta-analysis. Eisele DW, ed. *Head & Neck*. 2014;37(10):1536-1546. doi:<https://doi.org/10.1002/hed.23776>
22. Lee YQ, Kanagalingam J. Deep neck abscesses: the Singapore experience. *European Archives of Oto-Rhino-Laryngology*. 2010;268(4):609-614. doi:<https://doi.org/10.1007/s00405-010-1387-8>

23. Roccia F, Pecorari GC, Oliaro A, et al. Ten Years of Descending Necrotizing Mediastinitis: Management of 23 Cases. *Journal of oral and maxillofacial surgery : official journal of the American Association of Oral and Maxillofacial Surgeons*. 2007;65(9):1716-1724. doi:<https://doi.org/10.1016/j.joms.2006.10.060>
24. Taylor M, Patel H, Khwaja S, Rammohan K. Descending cervical mediastinitis: the multidisciplinary surgical approach. *European Archives of Oto-Rhino-Laryngology*. 2019;276(7):2075-2079. doi:<https://doi.org/10.1007/s00405-019-05471-z>
25. Endo S, Murayama F, Hasegawa T, et al. Guideline of surgical management based on diffusion of descending necrotizing mediastinitis. *The Japanese Journal of Thoracic and Cardiovascular Surgery*. 1999;47(1):14-19. doi:<https://doi.org/10.1007/bf03217934>
26. Stalfors J, Adielsson A, Ebenfelt A, Nethander G, Westin T. Deep neck space infections remain a surgical challenge. a study of 72 patients. *Acta Oto-laryngologica*. 2004;124(10):1191-1196. doi:<https://doi.org/10.1080/00016480410017864>
27. Inchingolo F, Hazballa D, Inchingolo AD, et al. Innovative Concepts and Recent Breakthrough for Engineered Graft and Constructs for Bone Regeneration: A Literature Systematic Review. *Materials*. 2022;15(3):1120. doi:<https://doi.org/10.3390/ma15031120>
28. Bambini F, Pellecchia M, Memè L, et al. Anti-Inflammatory Cytokines in Peri-Implant Soft Tissues: A Preliminary Study on Humans Using CDNA Microarray Technology. *European Journal of Inflammation*. 2007;5(3):121-127. doi:<https://doi.org/10.1177/1721727x0700500302>
29. Boscolo-Rizzo P, Marchiori C, Zanetti F, Vaglia A, Da Mosto MC. Conservative management of deep neck abscesses in adults: The importance of CECT findings. *Otolaryngology-Head and Neck Surgery*. 2006;135(6):894-899. doi:<https://doi.org/10.1016/j.otohns.2006.05.013>
30. Wang LF, Kuo WR, Tsai SM, Huang KJ. Characterizations of life-threatening deep cervical space infections: A review of one hundred ninety-six cases. *American Journal of Otolaryngology*. 2003;24(2):111-117. doi:<https://doi.org/10.1053/ajot.2003.31>
31. Gidley P, Ghorayeb B, Stiernberg C. Contemporary management of deep neck space infections. *Otolaryngology - Head and Neck Surgery*. 1997;116(1):16-22. doi:[https://doi.org/10.1016/s0194-5998\(97\)70345-0](https://doi.org/10.1016/s0194-5998(97)70345-0)
32. Vieira F, Allen SM, Stocks RMS, Thompson JW. Deep Neck Infection. *Otolaryngologic Clinics of North America*. 2008;41(3):459-483. doi:<https://doi.org/10.1016/j.otc.2008.01.002>
33. Brook I, Frazier EH. Clinical and microbiological features of necrotizing fasciitis. *Journal of clinical microbiology*. 1995;33(9):2382-2387. doi:<https://doi.org/10.1128/jcm.33.9.2382-2387.1995>
34. Deep Neck and Odontogenic Infections. ClinicalKey. Clinicalkey.com.au. Published 2019. <https://www.clinicalkey.com.au/#!/content/book/3-s2.0-B9780323611794000090>
35. Caccamese JF, Coletti DP. Deep Neck Infections: Clinical Considerations in Aggressive Disease. *Oral and Maxillofacial Surgery Clinics of North America*. 2008;20(3):367-380. doi:<https://doi.org/10.1016/j.coms.2008.03.001>
36. McGurk M. Diagnosis and treatment of necrotizing fasciitis in the head and neck region. *Oral and Maxillofacial Surgery Clinics of North America*. 2003;15(1):59-67. doi:[https://doi.org/10.1016/s1042-3699\(02\)00083-3](https://doi.org/10.1016/s1042-3699(02)00083-3)
37. Inchingolo F, Pacifici A, Gargari M, et al. CHARGE syndrome: an overview on dental and maxillofacial features. *European review for medical and pharmacological sciences*. 2014;18(15):2089-2093.
38. Schmidt SC, Strauch S, Rösch T, et al. Management of esophageal perforations. *Surgical Endoscopy*. 2010;24(11):2809-2813. doi:<https://doi.org/10.1007/s00464-010-1054-6>
39. Mureşan M, Mureşan S, Balmoş I, Sala D, Suci B, Torok A. Sepsis in Acute Mediastinitis – A Severe Complication after Oesophageal Perforations. A Review of the Literature. *The Journal of Critical Care Medicine*. 2018;5(2):49-55. doi:<https://doi.org/10.2478/jccm-2019-0008>
40. Kluge J. Die akute und chronische Mediastinitis [Acute and chronic mediastinitis]. *Der Chirurg; Zeitschrift für alle Gebiete der operativen Medizin*. 2016;87(6):469-477. doi:[10.1007/s00104-016-0172-7](https://doi.org/10.1007/s00104-016-0172-7)
41. Gasparro R, Sammartino G, Mariniello M, di Lauro AE, Spagnuolo G, Marenzi G. Treatment of periodontal pockets at the distal aspect of mandibular second molar after surgical removal of impacted third molar and application of L-PRF: a split-mouth randomized clinical trial. *Quintessence International (Berlin, Germany: 1985)*. 2020;51(3):204-211. doi:<https://doi.org/10.3290/j.qi.a43947>

42. Beka D, Lachanas VA, Dumas S, et al. Microorganisms involved in deep neck infection (DNIs) in Greece: detection, identification and susceptibility to antimicrobials. *BMC Infectious Diseases*. 2019;19(1). doi:<https://doi.org/10.1186/s12879-019-4476-3>
43. Hansen BW, Ryndin S, Mullen KM. Infections of Deep Neck Spaces. *Seminars in Ultrasound, CT and MRI*. 2020;41(1):74-84. doi:<https://doi.org/10.1053/j.sult.2019.10.001>
44. Sammartino G, Marenzi G, Howard CM, et al. Chondrosarcoma of the Jaw: A Closer Look at its Management. *Journal of Oral and Maxillofacial Surgery*. 2008;66(11):2349-2355. doi:<https://doi.org/10.1016/j.joms.2006.05.069>
45. Smith JL, Hsu JM, Chang J. Predicting deep neck space abscess using computed tomography. *American Journal of Otolaryngology*. 2006;27(4):244-247. doi:<https://doi.org/10.1016/j.amjoto.2005.11.008>
46. Eissa L, Mehanna AM. Biphasic CT imaging of deep neck infections (DNIs): how does dual injection mode helps in differentiation between types of collections? *The Egyptian Journal of Otolaryngology*. 2020;36(1). doi:<https://doi.org/10.1186/s43163-020-00047-9>
47. Inchingolo AD, Cazzolla AP, Di Cosola M, et al. The integumentary system and its microbiota between health and disease. *Journal of Biological Regulators and Homeostatic Agents*. 2021;35(2 Suppl. 1):303-321. doi:<https://doi.org/10.23812/21-2suppl1-30>
48. Lazor JB, Cunningham MJ, Eavey RD, Weber AL. Comparison of Computed Tomography and Surgical Findings in Deep Neck Infections. *Otolaryngology—Head and Neck Surgery*. 1994;111(6):746-750. doi:<https://doi.org/10.1177/019459989411100608>
49. Muñoz A, Castillo M, Melchor M, Gutiérrez RA. Acute Neck Infections: Prospective Comparison Between CT and MRI in 47 Patients. *Journal of Computer Assisted Tomography*. 2001;25(5):733-741. doi:<https://doi.org/10.1097/00004728-200109000-00011>
50. Crespo AN, Chone CT, Fonseca AS, Montenegro MC, Pereira R, Milani JA. Clinical versus computed tomography evaluation in the diagnosis and management of deep neck infection. *Sao Paulo Medical Journal*. 2004;122(6):259-263. doi:<https://doi.org/10.1590/s1516-31802004000600006>
51. Huang TT, Liu TC, Chen PR, Tseng FY, Yeh TH, Chen YS. Deep neck infection: Analysis of 185 cases. *Head & Neck*. 2004;26(10):854-860. doi:<https://doi.org/10.1002/hed.20014>
52. Parhiscar A, Har-El G. Deep Neck Abscess: A Retrospective Review of 210 Cases. *Annals of Otolaryngology, Rhinology & Laryngology*. 2001;110(11):1051-1054. doi:<https://doi.org/10.1177/000348940111001111>
53. Inchingolo F, Tatullo M, Marrelli M, et al. Combined occlusal and pharmacological therapy in the treatment of temporo-mandibular disorders. *European review for medical and pharmacological sciences*. 2011;15(11):1296-1300.
54. Gasparro R, Adamo D, Masucci M, Sammartino G, Mignogna MD. Use of injectable platelet-rich fibrin in the treatment of plasma cell mucositis of the oral cavity refractory to corticosteroid therapy: A case report. *Dermatologic Therapy*. 2019;32(5). doi:<https://doi.org/10.1111/dth.13062>
55. Miller LG, Perdreau-Remington F, Rieg G, et al. Necrotizing Fasciitis Caused by Community-Associated Methicillin-Resistant *Staphylococcus aureus* in Los Angeles. *New England Journal of Medicine*. 2005;352(14):1445-1453. doi:<https://doi.org/10.1056/nejmoa042683>
56. Inchingolo AD, Inchingolo AM, Malcangi G, et al. Effects of Resveratrol, Curcumin and Quercetin Supplementation on Bone Metabolism—A Systematic Review. *Nutrients*. 2022;14(17):3519-3519. doi:<https://doi.org/10.3390/nu14173519>
57. Ovassapian A, Tuncbilek M, Weitzel EK, Joshi CW. Airway Management in Adult Patients with Deep Neck Infections: A Case Series and Review of the Literature. *Anesthesia & Analgesia*. 2005;100(2):585-589. doi:<https://doi.org/10.1213/01.ane.0000141526.32741.cf>
58. Balzanelli MG, Distratis P, Dipalma G, et al. Sars-CoV-2 Virus Infection May Interfere CD34+ Hematopoietic Stem Cells and Megakaryocyte—Erythroid Progenitors Differentiation Contributing to Platelet Defection towards Insurgence of Thrombocytopenia and Thrombophilia. *Microorganisms*. 2021;9(8):1632-1632. doi:<https://doi.org/10.3390/microorganisms9081632>
59. Myers EN, Snyderman CH. *Operative Otolaryngology : Head and Neck Surgery*. Elsevier; 2018.
60. Potter JK, Herford AS, Ellis E. Tracheotomy versus endotracheal intubation for airway management in deep neck space infections. *Journal of Oral and Maxillofacial Surgery*. 2002;60(4):349-354. doi:<https://doi.org/10.1053/joms.2002.31218>
61. Huang TT, Tseng FY, Yeh TH, Hsu CJ, Chen YS. Factors affecting the bacteriology of deep neck infection: a retrospective study of 128 patients. *Acta Oto-Laryngologica*. 2006;126(4):396-401. doi:<https://doi.org/10.1080/00016480500395195>



62. Rzepakowska A, Rytel A, Krawczyk P, et al. The Factors Contributing to Efficiency in Surgical Management of Purulent Infections of Deep Neck Spaces. *Ear Nose & Throat Journal*. 2019;100(5):354-359. doi:<https://doi.org/10.1177/0145561319877281>
63. Termine N, Panzarella V, Ciavarella D, et al. Antibiotic prophylaxis in dentistry and oral surgery: use and misuse. *International dental journal*. 2009;59(5):263-270.
64. Ma C, Zhou L, Zhao JZ, et al. Multidisciplinary treatment of deep neck infection associated with descending necrotizing mediastinitis: a single-centre experience. *Journal of International Medical Research*. 2019;47(12):6027-6040. doi:<https://doi.org/10.1177/0300060519879308>
65. Ridder GJ, Technau-Ihling K, Sander A, Boedeker CC. Spectrum and Management of Deep Neck Space Infections: An 8-Year Experience of 234 Cases. *Otolaryngology–Head and Neck Surgery*. 2005;133(5):709-714. doi:<https://doi.org/10.1016/j.otohns.2005.07.001>
66. Mayor GP, Millan MS, Martinez-Vidal A. Is conservative treatment of deep neck space infections appropriate? *Head & Neck*. 2001;23(2):126-133. doi:[https://doi.org/10.1002/1097-0347\(200102\)23:2%3C126::aid-hed1007%3E3.0.co;2-n](https://doi.org/10.1002/1097-0347(200102)23:2%3C126::aid-hed1007%3E3.0.co;2-n)
67. Inchingolo F, Tatullo M, Marrelli M, et al. Clinical case-study describing the use of skin-perichondrium-cartilage graft from the auricular concha to cover large defects of the nose. *Head & Face Medicine*. 2012;8(1). doi:<https://doi.org/10.1186/1746-160x-8-10>
68. Nie W, Du L, Chen G, et al. Ultrasound-guided puncture drainage versus surgical incision drainage for deep neck space abscesses: a protocol for a systematic review with meta-analysis and trial sequential analysis. *BMJ Open*. 2024;14(1):e077631-e077631. doi:<https://doi.org/10.1136/bmjopen-2023-077631>
69. Yeow KM, Liao C, Hao SP. US-guided Needle Aspiration and Catheter Drainage as an Alternative to Open Surgical Drainage for Uniloculated Neck Abscesses. *Journal of Vascular and Interventional Radiology*. 2001;12(5):589-594. doi:[https://doi.org/10.1016/s1051-0443\(07\)61481-x](https://doi.org/10.1016/s1051-0443(07)61481-x)
70. Wheatley MJ, Stirling MC, Kirsh MM, Gago O, Orringer MB. Descending necrotizing mediastinitis: transcervical drainage is not enough. *The Annals of Thoracic Surgery*. 1990;49(5):780-784. doi:[https://doi.org/10.1016/0003-4975\(90\)90022-x](https://doi.org/10.1016/0003-4975(90)90022-x)
71. Gradon JD. Space-Occupying and Life-Threatening Infections of the Head, Neck, and Thorax. *Infectious Disease Clinics of North America*. 1996;10(4):857-878. doi:[https://doi.org/10.1016/s0891-5520\(05\)70330-5](https://doi.org/10.1016/s0891-5520(05)70330-5)
72. Chen C, Gao Q, Lu X. [Treatment of the pediatric neck abscess and surgery scar by high negative pressure drainage system]. *PubMed*. 2015;40(3):281-284. doi:<https://doi.org/10.11817/j.issn.1672-7347.2015.03.008>
73. Lawrence R, Bateman N. Controversies in the management of deep neck space infection in children: an evidence-based review. *Clinical Otolaryngology*. 2016;42(1):156-163. doi:<https://doi.org/10.1111/coa.12692>
74. Gu X, Chen W, Yuan K, Tan J, Sun S. The efficacy of artificial dermis combined with continuous vacuum sealing drainage in deep neck multiple spaces infection treatment. *Medicine*. 2021;100(5):e24367. doi:<https://doi.org/10.1097/md.00000000000024367>
75. Kleinberger AJ, Jumaily J, Spiegel JH. Safety of Modified Coronal Approach with Dissection Deep to Temporalis Fascia for Facial Nerve Preservation. *Otolaryngology–Head and Neck Surgery*. 2015;152(4):655-660. doi:<https://doi.org/10.1177/0194599814567869>
76. Cho IS, Lee MH, Cho J. Rare case of postoperative mediastinitis following thyroidectomy. *Korean Journal of Clinical Oncology*. 2021;17(1):52-55. doi:<https://doi.org/10.14216/kjco.21009>
77. Galio NJ. Peritonsillar Abscess. *American family physician*. 2017;95(8):501-506.
78. Mungul S, Maharaj S. Microbiology of paediatric deep neck space infection. *International Journal of Pediatric Otorhinolaryngology*. 2019;123:116-122. doi:<https://doi.org/10.1016/j.ijporl.2019.04.042>
79. Azar A, Alkheder A, Alsodi Z, Ayob H. Effective surgical drainage of a massive retropharyngeal abscess via an incision in the posterior wall of the oropharynx under local anesthesia. *International Journal of Surgery Case Reports*. 2024;123:110234-110234. doi:<https://doi.org/10.1016/j.ijscr.2024.110234>
80. Prado-Calleros HM, Jiménez-Fuentes E, Jiménez-Escobar I. Descending necrotizing mediastinitis: Systematic review on its treatment in the last 6 years, 75 years after its description. Eisele DW, ed. *Head & Neck*. 2016;38(S1):E2275-E2283. doi:<https://doi.org/10.1002/hed.24183>

81. Inchingolo F, Tatullo M, Abenavoli FM, et al. Severe Anisocoria after Oral Surgery under General Anesthesia. *International Journal of Medical Sciences*. 2010;7(5):314-318. doi:<https://doi.org/10.7150/ijms.7.314>
82. Daramola OO, Flanagan CE, Maisel RH, Odland RM. Diagnosis and treatment of deep neck space abscesses. *Otolaryngology–Head and Neck Surgery*. 2009;141(1):123-130. doi:<https://doi.org/10.1016/j.otohns.2009.03.033>
83. Bohanes T, Neoral C. Akutní mediastinitida [Acute mediastinitis]. *Rozhledy v chirurgii : měsíčník Československé chirurgické společnosti*. 2011;90(11):604-611.
84. Luan CW, Liu CY, Yang YH, et al. The Pathogenic Bacteria of Deep Neck Infection in Patients with Type 1 Diabetes, Type 2 Diabetes, and Without Diabetes from Chang Gung Research Database. *Microorganisms*. 2021;9(10):2059. doi:<https://doi.org/10.3390/microorganisms9102059>
85. Inchingolo AD, Dipalma G, Inchingolo AM, et al. The 15-Months Clinical Experience of SARS-CoV-2: A Literature Review of Therapies and Adjuvants. *Antioxidants*. 2021;10(6):881. doi:<https://doi.org/10.3390/antiox10060881>
86. Inchingolo F, Dipalma G, Paduanelli G, et al. Computer-based quantification of an atraumatic sinus augmentation technique using CBCT. *Journal of Biological Regulators and Homeostatic Agents*. 2019;33(6 Suppl. 2).
87. Villanueva-Fernández E, Casanueva-Muruáis R, Vivanco-Allende A, Llorente JL, Coca-Pelaz A. Role of steroids in conservative treatment of parapharyngeal and retropharyngeal abscess in children. *European Archives of Oto-Rhino-Laryngology*. 2022;279(11):5331-5338. doi:<https://doi.org/10.1007/s00405-022-07423-6>
88. Boscolo-Rizzo P, Stellin M, Muzzi E, et al. Deep neck infections: a study of 365 cases highlighting recommendations for management and treatment. *European archives of oto-rhino-laryngology*. 2012;269(4):1241-1249. doi:<https://doi.org/10.1007/s00405-011-1761-1>
89. Horowitz G, Ben-Ari O, Wasserzug O, Weizman N, Yehuda M, Fliss DM. The Transcervical Approach for Parapharyngeal Space Pleomorphic Adenomas: Indications and Technique. Amar S, ed. *PLoS ONE*. 2014;9(2):e90210. doi:<https://doi.org/10.1371/journal.pone.0090210>
90. Singhal G, Jain S, Sen K. Clinical Presentation and Microbiological Profile of Deep Neck Space Infections in Different Age Groups. *Indian Journal of Otolaryngology and Head & Neck Surgery*. 2020;74(Suppl 2):1870-1876. doi:<https://doi.org/10.1007/s12070-020-01869-y>
91. Montroy J, Breau RH, Cnossen S, et al. Change in Adverse Events After Enrollment in the National Surgical Quality Improvement Program: A Systematic Review and Meta-Analysis. Shamji M, ed. *PLOS ONE*. 2016;11(1):e0146254. doi:<https://doi.org/10.1371/journal.pone.0146254>
92. Conrad DE, Parikh SR. Deep Neck Infections. *Infectious Disorders - Drug Targets*. 2012;12(4):286-290. doi:<https://doi.org/10.2174/187152612801319348>
93. Nakazawa S, Shimizu K, Mogi A, Kuwano H. VATS segmentectomy: past, present, and future. *General Thoracic and Cardiovascular Surgery*. 2017;66(2):81-90. doi:<https://doi.org/10.1007/s11748-017-0878-6>
94. White A, Swanson SJ. Video-assisted thoracic surgery (VATS) segmentectomy: state of the art. *PubMed*. 2016;71(1):61-66.
95. Gonzalez M, Federici S, Perentes J. Uniportal VATS S9 segmentectomy: The ligamentum-based approach. *Multimedia manual of cardiothoracic surgery : MMCTS / European Association for Cardio-Thoracic Surgery*. 2021;2021. doi:<https://doi.org/10.1510/mmcts.2021.067>
96. Luciani C, Scacchi A, Vaschetti R, et al. The uniportal VATS in the treatment of stage II pleural empyema: a safe and effective approach for adults and elderly patients—a single-center experience and literature review. *World Journal of Emergency Surgery*. 2022;17(1). doi:<https://doi.org/10.1186/s13017-022-00438-8>
97. Chen IC, Hsu JH, Dai ZK. Management of descending necrotizing mediastinitis complicated with empyema by video-assisted thoracoscopic surgery. *Pediatric Pulmonology*. 2015;51(5):E17-E20. doi:<https://doi.org/10.1002/ppul.23375>