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Radiographic aids in oral and maxillofacial surgery – Recent Advances

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Abstract

The current scenario of radiological imaging of the maxillofacial region is a far cry from its humble beginnings. Rapid strides have been made over the years, with Radiology having made an immense impact on the practice of maxillofacial surgery. These advances in imaging, play a vital role not only in the diagnosis of maxillofacial disease conditions but also in the planning and implementation of appropriate and timely treatment. Since Radiology has a crucial role to

play in the practice of maxillofacial surgery it is essential that an OMFS should have basic understanding of the core concepts of different imaging techniques, their similarities and differences and should also be able to comprehend how imaging helps both diagnosing the clinical condition of a patient as well as management of the same. In this review article we have reviewed the current advances in the field of radiology and its uses in the field of diagnosis. **Key words:** Recent advances, PET CT, SPECT, DEXA, Narrow Band Imaging.

Introduction

The last decade has witnessed a significant change in the field of technology. The evolving technology aids the field of oral and maxillofacial surgery from diagnosis, treatment planning, to the surgical procedure itself.

The success rate of any surgical procedure depends upon pre-operative imaging. We have witnessed a huge leap from intraoral periapical radiographs to laser surface scanning in terms of imaging. The conventional radiographs used to show images in 2-dimensional plane while the recent techniques provide images in the 3dimensional plane which enables us to see the area of interest upto its finest detail.

Computed tomography (CT) scan

Sir Godfrey Hounsfield discovered the CT scan in 1967, and the first scan was performed in Wimbledon, England in 1971, but it was not publicised for another year. The scan can also be contrast enhanced in order to visualize the soft tissue.

CT scan is used to identify bone fracture along with the presence of bleed in extradural. subdural. intraventricular, subarachnoid and intracerebral compartment. It also plays a vital role in the detection of cerebral blood flow volume and the location of oedema. Its biggest advantage is that with all the resuscitation equipment the scan can be performed without any difficulties and it has a shorter scanning period of approximately 5 minutes. The only problem in the CT scan is that the small extra cerebral lesion in the posterior cranial fossa cannot be detected¹. Stengel D et al conducted a study to find out the accuracy of the CT scan in patients with major blunt trauma and find out that the sensitivity of pan CT scan was nearly about 84.6% in the head and neck region².

Contrast enhanced computed tomography is generally used to detect the carcinoma of the head and neck region. Its sensitivity in the case of primary tumor is about 68% and 63 % in case of recurrent carcinoma. Its specificity in the case of primary tumor is about 69% and 80 % in case of the recurrent carcinoma³⁻⁴.

In CT angiography a special dye is injected in the blood vessel to detect aneurysm, atherosclerosis, blood clot and arteriovenous malformation.

Magnetic resonance imaging (MRI)

Magnetic resonance imaging is based on the behaviour of the positively charged nuclear particles. MRI is noninvasive and the radiation dose is less as it uses nonionizing radiation. It is used as a diagnostic tool in soft tissue neoplasia, TMJ evaluation, malignant involvement of the lymph nodes, perineural invasion, and soft tissue swelling. But MRI cannot be used to detect the bone related diseases and it has a long scanning time. The sensitivity and specificity of MRI is dependent on the level of malignancy. If the malignancy level is greater than 10mm the MRI has a sensitivity of the 81.5% and specificity of 55.2% if the malignancy level is greater than 15mm then MRI has a sensitivity of the 78.6% and specificity of the 78.8%⁵.

MRI spectroscopy is the recent advancement in this imaging system which is used to evaluate the presence of a specific metabolite. MRI spectroscopy differentiates the non-malignant tissue from the malignant ones and also detects the post radiation changes. But its disadvantages are its cost and longer scan time. The longer scan time is generally due to the large number of images and the complexity of reconstruction.

Diffusion weighted imaging (DWI) came into reality in the year 1990. The diffusion weighted imaging is the work of Stejskal, Tanner and Le Bihan⁶. Our body mostly consists of water. This technique uses the random

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Brownian movement of water molecule within the voxel tissue. The DWI demonstrates the heterogenicity in terms of apparent diffusion coefficient (ADC) within the solid tumors. Hyper cellular tumor and pathological lymph node shows low apparent diffusion coefficient while oedema, inflammatory and necrotic process shows high apparent diffusion coefficient. DWI is used to detect acute brain is chaemia, white matter disease, haematoma and paediatric brain development and aging. The diffusion weighted imaging plays an immense role in the diagnosis of head and neck pathology⁷.

According to Wang et al there is significant reduction in the apparent diffusion coefficient value in neoplasm with the benign salivary gland tumor and cyst of the head and neck region having an ADC value of $1.22 \times 10^{-3} \text{ mm}^2/\text{s}$. The ADC value of necrosis helps in differentiating lymphadenitis from the metastasis. The ADC value for lymphadenitis is nearly about 0.89 ± 0.21 but in case of the malignancy it is nearly about 1.46 ± 0.46 . It has a predictive value of 86%, a sensitivity of 84% and specificity of 91%⁸⁻⁹.

²³Na in MRI oncology

²³Na shows quadrupolar nucleus which is also used for the detection of cancer. It is found that the sodium concentration of the malignant neoplasm is increased by 60% in comparison to the normal glandular tissue¹⁰.

Single photon emitted computed tomiography (SPECT)

SPECT produces images in the same manner as CT and MRI but with thin slices through a particular organ. It is used to detect cardiac perfusion, and brain, liver and bone malignancy. Thalium 201 SPECT shows a sensitivity of 95% in detecting squamous cell carcinoma¹¹.

Positron emission tomography computed tomography (PET CT)

In PET CT the metabolic or biochemical activity of the body is corelated with the anatomic imaging by CT scan. In this imaging modality intravenous bolus of 18 fluorodeoxyglucose is given 1 to 2 hours before then CT scan is performed at 2 to 3 mm slices and hypermetabolic regions are detected and colour coded. It is useful in diagnosing the distant metastasis, osteomyelitis of the jaw, metastatic masses of unknown origin and acute inflammatory lesions. 18 F-FDG PET CT shows a sensitivity of 87–90 % and a specificity of $80–93 \%^{12}$.

Ultrasound

Ultrasound uses the sound waves which is converted to the image form. For the head and neck region 7 to 12 MHz is used for diagnosis. Shorter the wavelength, better the resolution of the image. In comparison to the other modalities ultrasound is non-invasive, rapid, inexpensive, widely available and there is no harmful radiation produced. It is used to diagnose inflammation, salivary gland pathology, arteriovenous malformation, lymph node status and neoplasm¹³. The staging system with the ultrasound shows accuracy of nearly about 90%¹⁴. Now-a-days ultrasound guided biopsy is also performed as a diagnostic procedure.

Bone scan

Active bone formation in the body can be evaluated by bone scintigraphy method. The radioisotope used is Tc-99methylene diphosphate. Its uptake depends upon the osteoblastic and osteoclastic activity. Osteoblastic activity shows increased uptake of the radioisotope while the osteoclastic activity shows decreased uptake of the radioisotope. The hot spot area appears black as there is accumulation of the radionuclide due to increased bone growth. Cold spot area appears lighter or white because of the decreased metabolic activity. The bone scan has 3 phases i.e., perfusion phase, blood pooling

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phase and bony phase. It is used for the detection of malignancy, is chaemia, necrosis, hemangioma and radiation therapy¹⁵.

Dentascan (DS)

Dentascan is another modification of the CT scan software programme in which maxilla and mandible can be visualized in all three planes¹⁶. The Dentascan was initially developed in the mid-1980 but it gained popularity recently. It provides a better evaluation of the osseous maxilla and mandible which is found to be useful in cases of rehabilitation with dental implants¹⁷⁻¹⁹. Dentascan is also used to evaluate the invasion of carcinoma in the bone. Brocken Brough JM conducted a study to determine the diagnostic accuracy of the Dentascan and found that it shows a sensitivity of 95%, specificity of 79%, positive predictive value of 87% and negative predictive value of 92%. Although it shows highest evidence of diagnosis the problem with it is its cost which is nearly about 1650 dollars²⁰.

Dual-energy x-ray absorptiometry (DEXA)

DEXA scan is generally used to evaluate the bone mineral density. According to WHO "T" score is the score which is determined to evaluate the mean standard deviation from the reference population and patient's average bone mineral density²¹⁻²².

- Greater than or equal to -1.0: normal
- Less than -1.0 to greater than -2.5: osteopenia
- Less than or equal to -2.5: osteoporosis
- Less than or equal to -2.5 plus fragility fracture: severe osteoporosis

The advantage of DEXA scan is that it is fast, reliable and needs little exposure²³. Jackson W used the DEXA scan and studied the change in total body composition of a patient with head and neck cancer undergoing chemotherapy and found that there is substantial loss of muscle and fat body mass seen in the patients undergoing concurrent chemoradiotherapy 24 .

Narrow band imaging system (NBI)

The recent advancement in the field of cancer detection is the narrow band imaging. If the tumour is less than or equal to 1 centimetre in size, it is very difficult to detect²⁵. This technique is based on the principle that it alters the spectral characteristics of illuminating light by limiting the bandwidth of an optical filter in the light source²⁶. The NBI contains red, green and blue (RGB) wavelength in sequential video endoscope. The optical filter present in the NBI system allows narrow band light to pass a wavelength of 415nm which only passes superficially through the mucosa and enhances its vasculature. Then a longer wavelength of 515nm is used which reveals the submucosal vessel as it passes the deeper tissue²⁷. Blue light has less scattering and penetration in comparison to the red light so the blue light enhances the image resolution²⁸. The peak absorption spectrum of the hemoglobin corresponds to the blue light which also enhances the visualization of the submucosal vessel. A monochromatic charge coupled device (CCD) enhances the reflection, and an image processor builds a composite pseudo color image that is presented on a high-definition video screen 27 . Thus, the lesion which is initially not detected through the white light emitted by the endoscope is now visible in the blue light filter of the narrow band imaging based on the neo angiogenesis and increased vascularity of tumor. Integrating this optical technology with magnifying endoscopy (NBI-ME), which allows the endoscopist to zoom in on the mucosa by merely pressing a specialised button, increases the diagnostic usefulness of this optical technology. The advantage of the NBI is that it is not a cumbersome procedure, easy to perform and can also detect the lesion less than equal to

1 centimetre in dimension²⁹. NBI is also used to detect the primary tumor³⁰. Hayashi et al. studied 46 patients with cervical lymph node metastases due to SCC. In none of the patients could a standard clinical examination or white light naso endo scopy detect a lesion. All of them were then subjected to NBI with magnifying endoscopy, which revealed 26 worrisome lesions. All of them were subjected to a biopsy, and 16 of them were determined to have SCC³¹. Zhou H et al conducted a metanalysis where he included 25 studies with 6187 lesion and found the sensitivity, specificity, positive predictive value, negative predictive value and odds ratio of 88.5%, 95.6%, 12.33, 0.11 and 121.26 respectively in cases of narrow band imaging³².

Fluorescent lifetime imaging microscopy (FILM)

Fluorescent Lifetime Imaging Microscopy (FILM) is a potential imaging modality that extracts various suggestive factors from autofluorescence signals, such as intensity, lifetime, and wavelength, to distinguish malignant from healthy tissue³³⁻³⁴. This method is mostly used for the intraoperative procedure³⁵.

When development of the oral cancer occurs, there is change in the stratified squamous epithelium, lamina propria and connective tissue due to morphological, biochemical and functional alterations³⁶. During the development of precancerous and cancerous lesion there is reduction in the nicotinamide adenine dinucleotide (NADH) and flavin adenine dinucleotide (FAD)³⁷⁻³⁸. The fluorescent intensity of the NADH to FAD is called as the optical redox ratio³⁹. A decrease in the optical redox ratio is frequently blamed for increasing cellular metabolic activity, which is a hallmark of neoplastic cell transformation⁴⁰. As a consequence, evaluating NADH, FAD, and collagen autofluorescence for optical biomarkers of oral epithelial cancer could be useful. Duran-Sierra E conducted a study to differentiate the oral epithelial dysplasia, SCC and normal healthy cell and concluded that the auto fluorescent lifetime imaging microscopy provides is a better method to differentiate dysplasia, neoplasia and normal healthy cell³⁶.

Cone beam computed tomography (CBCT)

In the last 10-year CT scan is replaced by CBCT for a particular region of bone. Detailed reconstruction is found in CBCT as compared to CT scan because of its small voxel size, good spatial resolution and detailed reconstruction of the image⁴¹. CBCT is used to diagnose chronic sinusitis, implant placement sites, maxillofacial pathology, orthodontic and 3 dimensional cephalometry, post operative aesthetic evaluation in the orthognathic patients, determination of the canal in the tooth, osteomyelitis and salivary gland pathologies. The advantage of CBCT scan is that it provides 3dimensional data setup, has potential for generating 2dimensional image like lateral cephalogram, high resolution, easy handling, less disturbance from the metal artefacts, compatible with the digital imaging and communication in medicine and also has low radiation dose. The disadvantage of the CBCT is that low contrast range, limited detector size cause limited field of view, limited soft tissue information and increased noise due to the scatter radiation and along with loss of resolution 42 .

Temporomandibular joint arthroscopy

Dr. Ohnishi was the first to perform the TMJ arthroscopy in the year 1974 but he reported in the Japanese literature in the year 1975⁴³. It is a technique used for the visualization of the internal joint structures directly and to perform surgical procedures under visual control with the guidance of the arthroscope. It can be used for the diagnostic and therapeutic purposes. The advantages of the TMJ arthroscopy are that no significant incision, reduced incidence of facial nerve injury and the ability of diagnostic along with

therapeutic abilities but it requires a steep learning curve to perform.

Conclusion

This paper demonstrates how developments in technology have enabled diagnostic imaging procedures to provide deeper insight into physiological structure and function for medicinal applications. We must also look at the evidence before implementing new radiological procedures, taking into account the influence on clinical results, health economics, and radiation protection concerns.

Reference

 Bruce DA. Imaging after head trauma: why, when and which. Childs Nerv Syst. 2000 Nov;16(10-11):755 9.

2. Stengel D, Otters Bach C, Matthes G, Weigeldt M, Grundei S, Rademacher G, Tittel A, Mutze S, Ekkernkamp A, Frank M, Schmucker U, Seifert J. Accuracy of single-pass whole-body computed tomography for detection of injuries in patients with major blunt trauma. CMAJ. 2012 May 15;184(8):869-76.

3. Som PM. Lymph nodes of the head and neck. Radiology. 1987;165(3):593–600.

4. Di Martino E, Nowak B, Hassan HA, et al. Diagnosis and Staging of Head and Neck Cancer: A Comparison of Modern Imaging Modalities (Positron Emission Tomography, Computed Tomography, Color-Coded Duplex Sonography) With Pan endoscopic and Histopathologic Findings. Arch Otolaryngol Head Neck Surg. 2000;126(12):1457–1461.

5. Guenzel T, Franzen A, Wiegand S, Kraetschmer S, Jahn LG, Mironczuk R, Thomas W And Schro T. The value of PET compared to MRI in malignant head and neck tumors. Anticancer Research March 2013; 33 (3) 1141-1146. 6. Le Bihan D. Diffusion MRI: what water tells us about the brain. EMBO Mol Med. 2014; 6:569–573.

7. Baliyan V, Das CJ, Sharma R, Gupta AK. Diffusion weighted imaging: Technique and applications. World J Radiol. 2016;8(9):785-798.

8. Wang J, Takashima S, Takayama F, Kawakami S, Saito A, Matsushita T, Momose M, Ishiyama T. Head and neck lesions: characterization with diffusion-weighted echo-planar MR imaging. Radiology. 2001; 220:621–630.

9. Kato H, Kanematsu M, Kato Z, Teramoto T, Mizuta K, Aoki M, Makita H, Kato K. Necrotic cervical nodes: usefulness of diffusion-weighted MR imaging in the differentiation of suppurative lymphadenitis from malignancy. Eur J Radiol. 2013;82: e28–e35.

10. Ouwerkerk R, Jacobs MA, Macura KJ, Wolff AC, Stearns V, Mezban SD, et al. Elevated tissue sodium concentration in malignant breast lesions detected with non-invasive 23Na MRI. Breast Cancer Res Treat. 2007; 106:151–160.

11. Valdes Almos et al. Thallium-201 SPECT in the Diagnosis of Head and Neck Cancer J NucÃ-Med 1997; 38:873-879.

12. Schöder H, Fury M, Lee N, Kraus D. PET monitoring of therapy response in head and neck squamous cell carcinoma. J Nucl Med. 2009;50 Suppl 1:74S–88.

13. Kotecha S, Bhatia P, Rout PG. Diagnostic ultrasound in the head and neck region. Dent Update. 2008 Oct;35(8):529-30, 533-4.

14. Koischwitz D, Gritzmann N. Ultrasound of the neck. Radiol Clin North Am 2000; 38: 1029–1045.

 Balaji SM, Balaji PP. Radiodiagnosis. Textbook of Oral & Maxillofacial Surgery, 3rd edition. New Delhi: Elsevier;2018. Au-Yeung KM, Ahuja AT, Ching AS, Metreweli C.
Dentascan in oral imaging. Clinical radiology. 2001 Sep 1;56(9):700-13.

17. Yanagisawa K, Abrahams JJ, Friedman CD. Dentascan: a new imaging method for the maxilla and mandible. New England Otolaryngological Society, Boston, MA. 1990 Oct.

18. Vining E, Friedman CD, Abrahams JJ, Lowlicht R. Diagnosis of oroantral fistula using Dentascan. Poster presentation at the American Academy of Otolaryngology Head and Neck Surgery, San Diego, CA. 1990 Sep.

19. Abrahams JJ, Levine B. Expanded applications of Dentascan (multiplanar computerized tomography of the mandible and maxilla). Int J Periodont Rest Dent 1990;10: 465-471.

20. Brocken Brough JM, Petruzzelli GJ, Lomasney L. Dentascan as an accurate method of predicting mandibular invasion in patients with squamous cell carcinoma of the oral cavity. Archives of otolaryngology–head & neck surgery. 2003 Jan 1;129(1):113-7.

21. Krugh M, Langaker MD. Dual Energy X-ray Absorptiometry. [Updated 2021 Jul 26]. In: Stat Pearls [Internet]. Treasure Island (FL): Stat Pearls Publishing; 2022 Jan

22. Expert Panel on Musculoskeletal Imaging: Ward RJ, Roberts CC, Bencardino JT, Arnold E, Baccei SJ, Cassidy RC, Chang EY, Fox MG, Greenspan BS, Gyftopoulos S, Hochman MG, Mintz DN, Newman JS, Reitman C, Rosenberg ZS, Shah NA, Small KM, Weissman BN.ACR Appropriateness

Criteria® Osteoporosis and Bone Mineral Density. J Am Coll Radiol. 2017 May;14(5S): S189-S202.

23. Ramos RL, Armán JA, Galeano NA, Hernández AM, Gómez JG, Molinero JG. Dual energy X-ray

absorptimetry: fundamentals, methodology, and clinical applications. Radiología (English Edition). 2012 Sep 1;54(5):410-23.

24. Jackson W, Alexander N, Schipper M, Fig L, Feng F, Jolly S. Characterization of changes in total body composition for patients with head and neck cancer undergoing chemoradiotherapy using dual-energy x-ray absorptiometry. Head & neck. 2014 Sep;36(9):1356-62.

25. Watanabe A, Tsujie H, Taniguchi M, Hosokawa M, Fujita M, Sasaki S. Laryngoscopic detection of pharyngeal carcinoma in situ with narrowband imaging. Laryngoscope 2006; 116:650-4.

26. Gono K, Obi T, Yamaguchi M, Ohyama N, Machida H, Sano Y, et al. Appearance of enhanced tissue features in narrow-band endoscopic imaging. J Biomed Opt 2004; 9:568-77.

27. Uedo N, Ishihara R, Iishi H, Yamamoto S, Yamamoto S, Yamada T, et al. A new method of diagnosing gastric intestinal metaplasia: narrow-band imaging with magnifying endoscopy. Endoscopy 2006; 38:819-24.

28. Kara MA, Bergman JJGHM. Autofluorescence imaging and narrow-band imaging for the detection of early neoplasia in patients with Barrett's esophagus. Endoscopy 2006; 38:627-31.

29. Piazza C, Dessouky O, Peretti G, Cocco D, De Benedetto L, Nicolai P. Narrow-band imaging: a new tool for evaluation of head and neck squamous cell carcinomas. Review of the literature. Acta otorhinolaryngologica italica. 2008 Apr;28(2):49.

30. Tan NC, Herd MK, Brennan PA, Puxeddu R. The role of narrow band imaging in early detection of head and neck cancer. British Journal of Oral and Maxillofacial Surgery. 2012 Mar 1;50(2):132-6.

31. Hayashi T, Muto M, Hayashi R, Minashi K, YanoT, Kishimoto S, Ebihara S. Usefulness of narrow-band

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imaging for detecting the primary tumor site in patients with primary unknown cervical lymph node metastasis. Japanese journal of clinical oncology. 2010 Jun 1;40(6):537-41.

32. Zhou H, Zhang J, Guo L, Nie J, Zhu C, Ma X. The value of narrow band imaging in diagnosis of head and neck cancer: a meta-analysis. Scientific reports. 2018 Jan 11;8(1):1-1.

33. Elson D, Requejo-Isidro J, Munro I, Reavell F, Siegel J, Suhling K, Tadrous P, Benninger R, Lanigan P, McGinty J, Talbot C. Time-domain fluorescence lifetime imaging applied to biological tissue. Photochemical & Photobiological Sciences. 2004;3(8):795-801.

34. Cubeddu R, Comelli D, D'Andrea C, Taroni P, Valentini G. Time-resolved fluorescence imaging in biology and medicine. Journal of Physics D: Applied Physics. 2002 Apr 16;35(9): R61.

35. Sun Y, Phipps JE, Meier J, Hatami N, Poirier B, Elson DS, Farwell DG, Marcu L. Endoscopic fluorescence lifetime imaging for in vivo intraoperative diagnosis of oral carcinoma. Microscopy and Microanalysis. 2013 Aug;19(4):791-8.

36. Duran-Sierra E, Cheng S, Cuenca-Martinez R, Malik B, Maitland KC, Cheng YL, Wright J, Ahmed B, Ji J, Martinez M, Al-Khalil M. Clinical label-free biochemical and metabolic fluorescence lifetime endoscopic imaging of precancerous and cancerous oral lesions. Oral oncology. 2020 Jun 1; 105:104635.

37. Müller MG, Valdez TA, Georgakoudi I, Backman V, Fuentes C, Kabani S, Laver N, Wang Z, Boone CW, Dasari RR, Shapshay SM. Spectroscopic detection and evaluation of morphologic and biochemical changes in early human oral carcinoma. Cancer: Interdisciplinary International Journal of the American Cancer Society. 2003 Apr 1;97(7):1681-92.

38. Shah AT, Demory Beckler M, Walsh AJ, Jones WP, Pohlmann PR, Skala MC. Optical metabolic imaging of treatment response in human head and neck squamous cell carcinoma. PloS one. 2014 Mar 4;9(3): e90746.

39. Chance B, Schoener B, Oshino R, Itshak F, Nakase Y. Oxidation-reduction ratio studies of mitochondria in freeze-trapped samples. NADH and flavoprotein fluorescence signals. Journal of Biological Chemistry. 1979 Jun 10;254(11):4764-71.

40. Zhang Z, Blessington D, Li H, Busch TM, Glickson JD, Luo Q, Chance B, Zheng G. Redox ratio of mitochondria as an indicator for the response of photodynamic therapy. Journal of biomedical optics. 2004 Jul;9(4):772-8.

41. Stratemann SA, Huang JC, Maki K, Miller AJ, Hatcher DC. Comparison of cone beam computed tomography imaging with physical measures. Dent maxillofacial Radiology. 2008 Feb;37(2):80-93.

42. De Vos W, Casselman J, Swennen G. Cone-beam computerized tomography (CBCT) imaging of the oral and maxillofacial region: a systematic review of the literature. International journal of oral and maxillofacial surgery. 2009 Jun 1;38(6):609-25.

43. Murakami K. Rationale of arthroscopic surgery of the temporomandibular joint. J Oral Biol Craniofac Res. 2013 Sep-Dec;3(3):126-34.