ELSEVIER

Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Letter to the Editor

To the Editor,

for 90% of saliva production [3].

glands [5].

Comments on "The tubarial salivary glands: First description of a potential new organ at risk for head-neck radiotherapy"

We read with interest the recent article by Valstar et al. report-

ing on the tubarial glands: a previously unreported new organ at

risk for radiotherapy [1]. There are several aspects that should be

considered before these glands are to be called a new organ. What gives a group of tissues in a given part of the body the features to

be considered as an organ? When it comes to glands, what distinguishes minor from major glands as adnexal organs from essential

organs? The evolution of a new organ was already studied in detail

via a placenta model recently [2], with organs such as major sali-

vary glands presenting their own enervation and vascularization

including a higher branching as opposed to minor salivary glands,

and it remains clear that the three major salivary glands account

ized by a continuous lining that encompasses the larynx, phar-

ynx, oral cavity, nasal cavities and auditory tubes. Underneath

this mucous membrane a high number of racemose mucous

glands can be seen, which are especially numerous at the upper

part of the pharynx around the orifices of the auditory tubes, in

which case they are obviously not called salivary glands [4].

Therefore, from an anatomical point of view, the presence of

seromucous glands in the adjacent connective tissue is very com-

mon even in the respiratory submucosa, which would be the case

with the tubarial glands, when they should not be called salivary

The mucous membrane of the aerodigestive tract is character-



Funding source

None.

Declaration of Competing Interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

References

- [1] Valstar MH, de Bakker BS, Steenbakkers RJHM, de Jong KH, Smit LA, Klein Nulent TJW, et al. The tubarial salivary glands: first description of a potential new organ at risk for head-neck radiotherapy. Radiother Oncol 2020:1–7. https://doi.org/10.1016/j.radonc.2020.09.034.
- [2] Griffith OW, Wagner GP. The placenta as a model for understanding the origin and evolution of vertebrate organs. Nat Ecol Evol 2017;1:1–10. https://doi.org/10.1038/s41559-017-0072.
- [3] Porcheri C, Mitsiadis TA. Physiology, pathology and regeneration of salivary glands. Cells 2019;8. https://doi.org/10.3390/cells8090976.
- [4] Berkovitz BKB, Evans BT, Hopkins C, McHanwell S. Upper aerodigestive tract. In: Standring S, editor. Gray's Anat. Anat. basis Clin. Pract.. Elsevier; 2015. p. 507–604.
- [5] Beule AG. Funktionen und Funktionsstörungen der respiratorischen Schleimhaut der Nase und der Nasennebenhöhlen. Laryngo-Rhino-Otol 2010;89(S 01):S15–34. https://doi.org/10.1055/s-0029-1246124.
- [6] Valentino W, Paknezhad H, Sataloff RT. Heterotopic salivary gland tissue in the larynx. Ear Nose Throat J 2020;99(8):NP84–5. https://doi.org/10.1177/0145561319853816.
- [7] Klein Nulent TJW, Valstar MH, de Keizer B, Willems SM, Smit LA, Al-Mamgani A, et al. Physiologic distribution of PSMA-ligand in salivary glands and seromucous glands of the head and neck on PET/CT. Oral Surg Oral Med Oral Pathol Oral Radiol 2018;125(5):478–86. https://doi.org/10.1016/j.goooo.2018.01.011.

Furthermore, heterotopic salivary gland disease - the presence of salivary gland tissue arising outside the normal distribution of the minor and major salivary glands tissue has been documented in the middle ear, larynx, lower neck, chest wall, sternoclavicular joint and brain [4]. The exact mechanism of salivary gland heterotopia is uncertain but as this is a known feature it seems plausible that the findings of "tubarial glands" may not be as unique as suggested [6]. Also, from a microscopic standpoint, the authors failed to demonstrate genuine lobular structures and ductal differentia-

tion, which are necessary to support the theory of a "previously

unnoticed pair of salivary glands in the nasopharynx" [3].

Last but not least, it comes to our attention that when the same research group published the PET/CT physiologic distribution of PSMA-ligand in salivary glands and seromucous glands of the head and neck, the tubarial seromucous glands did not seem to present a significant uptake [7]. Accordingly, it seems too early to state that this area may lead to significant clinical toxicity based on imaging findings from two patients [1].

Daniel Cohen Goldemberg*

Clinical Research Coordination, National Cancer Institute of Brazil (INCA), Rio de Janeiro, Brazil

* Address: Clinical Research Coordination, National Cancer Institute,
 COPQ/INCA, Rua André Cavalcanti, 37 – 5° andar-Anexo, Centro – Rio de Janeiro, RJ, Brazil.

E-mail address: daniel.cohen@inca.gov.br

Tiago Novaes Pinheiro

School of Health Sciences, Department of Dentistry, Amazonas State University, Manaus. Brazil

Alan Roger Santos-Silva

Oral Diagnosis Department, Semiology and Pathology Areas, Piracicaba Dental School, University of Campinas (UNICAMP), Piracicaba, Brazil

Andreia Cristina de Melo

Clinical Research Coordination, National Cancer Institute of Brazil (INCA), Rio de Janeiro, Brazil Jair Carneiro Leão

Oral Medicine Postgraduate Program Federal University of Pernambuco (UFPE), Recife, Brazil

Stefano Fedele

University College London (UCL) Eastman Dental Institute, London, UK
University College London (UCL), Eastman Dental Institute, NIHR
University College London Hospitals (UCLH), Biomedical Research
Centre, London, United Kingdom

Stephen Porter University College London (UCL) Eastman Dental Institute, London, UK Received 30 October 2020 Accepted 2 December 2020

Available online 11 December 2020