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1 Mature dendritic cells enriched in regulatory molecules may control regulatory T cells and
2 the prognosis of head and neck cancer

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4 Kiyoshi Minohara^{1,2*}, Masaki Imai^{1*}, Takuma Matoba^{1,2*}, James Badger Wing^{3,4}, Hiroaki
5 Shime¹, Mizuyu Odanaka¹, Ryuta Uraki^{1,5,6}, Daisuke Kawakita², Tatsuya Toyama⁷, Satoru
6 Takahashi⁸, Akimichi Morita⁹, Shingo Murakami², Naganari Ohkura^{10,11}, Shimon
7 Sakaguchi¹⁰, Shinichi Iwasaki², Sayuri Yamazaki¹

8

9 ¹Department of Immunology, Nagoya City University Graduate School of Medical Sciences,
10 Nagoya, Japan

11 ²Department of Otorhinolaryngology, Head and neck surgery, Nagoya City University
12 Graduate School of Medical Sciences, Nagoya, Japan

13 ³Laboratory of Human Immunology (Single Cell Immunology), Immunology Frontier
14 Research Center, Osaka University, Osaka, Japan.

15 ⁴Human Single Cell Immunology Team, Center for Infectious Disease Education and
16 Research (CiDER), Osaka University, Osaka, Japan.

17 ⁵Division of Virology, Institute of Medical Science, University of Tokyo, Tokyo, Japan.

18 ⁶The Research Center for Global Viral Diseases, National Center for Global Health and
19 Medicine Research Institute, Tokyo, Japan.

20 ⁷Department of Breast Surgery, Nagoya City University Graduate School of Medical
21 Sciences, Nagoya, Japan

22 ⁸Department of Experimental Pathology and Tumor Biology, Nagoya City University,
23 Graduate School of Medical Sciences, Nagoya, Japan

24 ⁹Department of Geriatric and Environmental Dermatology, Nagoya City University Graduate
25 School of Medical Sciences, Nagoya, Japan

26 ¹⁰Department of Experimental Immunology, World Premier International Research Center
27 Initiative, Immunology Frontier Research Center, Osaka University, Osaka, Japan

28 ¹¹Department of Frontier Research in Tumor Immunology, Center of Medical Innovation and
29 Translational Research, Graduate School of Medicine, Osaka University, Osaka, Japan

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31 * These authors contributed equally to this work.

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33 **Abstract**

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35 We previously reported that regulatory T (Treg) cells expressing CTLA-4 on the cell surface
36 are abundant in head and neck squamous cell carcinoma (HNSCC)¹. The role of expanded
37 Treg cells in the tumor microenvironment of HNSCC remains unclear. In this study, we
38 revealed that the tumor microenvironment of HNSCC is characterized by the high expression
39 of genes related to Treg cells, dendritic cells (DCs), and interleukin (IL)-17-related
40 molecules. Increased expression of *IL17A*, *IL17F*, or *IL23A* contributes to a favorable
41 prognosis of HNSCC. In the tumor microenvironment of HNSCC, *IL23A* and *IL12B* are
42 expressed in mature dendritic cells enriched in regulatory molecules (mregDCs)². The
43 mregDCs in HNSCC are a migratory and mature phenotype; their signature genes strongly
44 correlate with Treg signature genes in HNSCC. We also observed that *IL17A* was highly
45 expressed in Th17 cells and exhausted CD8⁺ T cells in HNSCC. These data suggest that
46 mregDCs in HNSCC may contribute to the prognosis by balancing Treg cells and effector T
47 cells that produce IL-17. Targeting mregDCs may be a novel strategy for developing new
48 immune therapies against HNSCC.

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50 **REFERENCES**

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52 on the cell surface with a proliferative gene profile are key features of human head
53 and neck cancer. *Int J Cancer*. 2019; 144: 2811-2822.
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