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Recent findings on the impact of ErbB receptors status on prognosis and therapy of head and neck squamous cell carcinoma

Camilla Palumbo¹. Monica Benvenuto^{1,2}. Chiara Focaccetti¹. Loredana Albonici¹, Loredana Cifaldi^{1,1}, Alessandra Rufini^{2,4}, Daniela Nardozi⁵, Valentina Angiolini⁵, Arianna Bei⁶, Laura Masuelli⁵ and Roberto Bei**

Department of Clinical Sciences and Translational Medicine. University of Rome 'Tor Vergata'. Rome: Italy. Saint Camillus International University of Health and Medical Sciences, Rome, Italy, "Academic Department of Pediatrics (DPUO). Ospedale Pediatrico Bambino Gesia, Istituto di Ricovero e Cura a Carattere Scientifico. BRCCSI, Rome, Italy, "Department of Biomedicine and Prevancion, University of Rome "Tor Vergata", Rome. Rate "Department of Experimental Medicine, Sapienta University of Rome, Rome, Italia "Medical School, Linkersity of Rome "for Vergata", flome, Italy

Head and neck squamous cell carcinoma (HNSCC) is the sixth most common cancer type. has often an aggressive course and is poorly responsive to current therapeutic approaches, so that 5-year survival rates for patients diagnosed with advanced disease is lower than 50%. The Epidermal Growth Factor Receptor (EGFR) has emerged as an established oncogene in HNSCC. Indeed, although HNSCCs are a heterogeneous group of cancers which differ for histological, molecular and clinical features, EGFR is overexpressed or mutated in a percentage of cases up to about 90%. Moreover, aberrant expression of the other members of the ErbB receptor family, ErbB2, ErbB3 and Erb84, has also been reported in variable proportions of HINSCCs. Therefore, an increased expression/activity of one or multiple ErbB receptors is found in the vast majority of patients with HNSCC. While aberrant Erb8 signaling has long been known to play a critical role in tumor growth, angiogenesis, invasion, metastatization and resistance to therapy, more recent evidence has revealed its impact on other features of cancer cells' biology, such as the ability to evade antitumor immunity. In this paper we will review recent findings on how ErbB receptors expression and activity, including that associated with non-canonical signaling mechanisms, impacts on prognosis and therapy of HNSCC.

head and neck squamous cell carcinoma, EGFR, ErbB receptors, Cetuximab, immune checkpoint inhibitors

1. Introduction

Head and neck (HN) cancers include a heterogeneous group of cancers which differ for histological, molecular and clinical features (1-3). More than 90% of HN cancers are squamous cell carcinomas (HNSCCs) arising from the mucosal epithelium of the oral cavity, pharynx and larynx, while much less common HN subtypes originate from the salivary glands, sinuses, muscles or nerves in the head and neck! (2-4).

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Table of Contents Receptors Recent Findings

- 1. Understanding the eBook Receptors Recent Findings
 - The Rise of Digital Reading Receptors Recent Findings
 - Advantages of eBooks Over Traditional Books
- 2. Identifying Receptors Recent Findings
 - Exploring Different Genres
 - o Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
- 3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Receptors Recent Findings
 - User-Friendly Interface
- 4. Exploring eBook Recommendations from Receptors Recent Findings
 - Personalized Recommendations
 - $\circ\,$ Receptors Recent Findings User Reviews and Ratings
 - $\circ\,$ Receptors Recent Findings and Bestseller Lists
- 5. Accessing Receptors Recent Findings Free and Paid eBooks
 - Receptors Recent Findings Public Domain eBooks
 - Receptors Recent Findings eBook Subscription Services
 - Receptors Recent Findings Budget-Friendly Options

- 6. Navigating Receptors Recent Findings eBook Formats
 - o ePub, PDF, MOBI, and More
 - Receptors Recent Findings Compatibility with Devices
 - Receptors Recent Findings Enhanced eBook Features
- 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Receptors Recent Findings
 - Highlighting and Note-Taking Receptors Recent Findings
 - Interactive Elements Receptors Recent Findings
- 8. Staying Engaged with Receptors Recent Findings
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Receptors Recent Findings
- 9. Balancing eBooks and Physical Books Receptors Recent Findings
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Receptors Recent Findings
- 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
- 11. Cultivating a Reading Routine Receptors Recent Findings
 - Setting Reading Goals Receptors Recent Findings
 - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Receptors Recent Findings
 - Fact-Checking eBook Content of Receptors Recent Findings
 - Distinguishing Credible Sources
- 13. Promoting Lifelong Learning
 - Utilizing eBooks for Skill Development
 - Exploring Educational eBooks
- 14. Embracing eBook Trends
 - Integration of Multimedia Elements

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