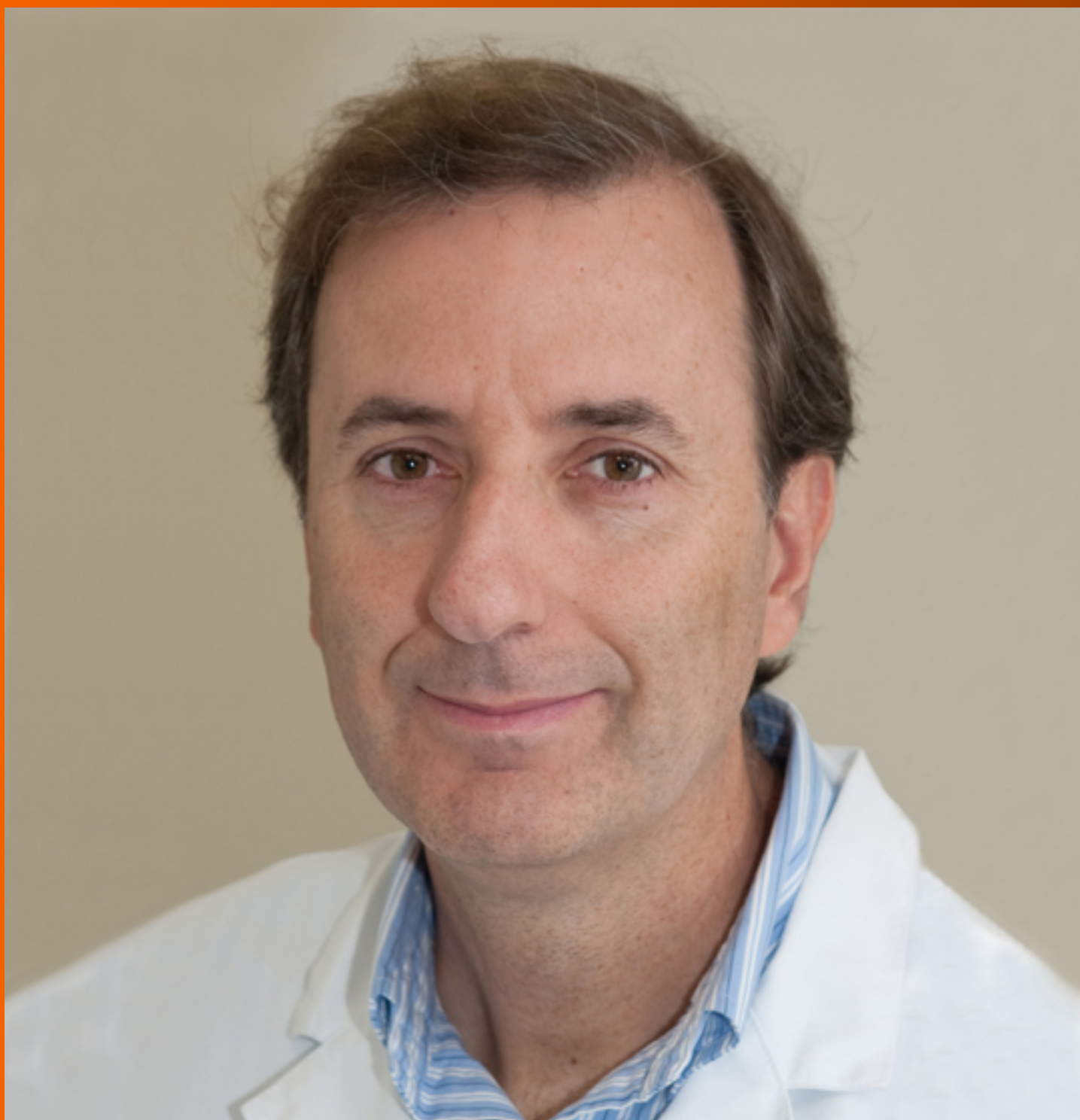


World Journal of *Clinical Cases*

World J Clin Cases 2019 March 6; 7(5): 548-690



**ORIGINAL ARTICLE****Retrospective Study**

- 548** Clinical presentation and early predictors for poor outcomes in pediatric myocarditis: A retrospective study
Rodriguez-Gonzalez M, Sanchez-Codez MI, Lubian-Gutierrez M, Castellano-Martinez A

Observational Study

- 562** Safety of an improved patent ductus arteriosus occluder for transcatheter closure of perimembranous ventricular septal defects with abnormally attached tricuspid chordae tendineae
He L, Du YJ, Cheng GS, Zhang YS

META-ANALYSIS

- 572** Adiponectin gene polymorphisms and risk of gestational diabetes mellitus: A meta-analysis
Huang LT, Wu SL, Liao X, Ma SJ, Tan HZ
- 585** Maternal serum level of resistin is associated with risk for gestational diabetes mellitus: A meta-analysis
Hu SM, Chen MS, Tan HZ
- 600** Docetaxel, cisplatin, and 5-fluorouracil compared with epirubicin, cisplatin, and 5-fluorouracil regimen for advanced gastric cancer: A systematic review and meta-analysis
Li B, Chen L, Luo HL, Yi FM, Wei YP, Zhang WX

CASE REPORT

- 616** Sustained complete response to erlotinib in squamous cell carcinoma of the head and neck: A case report
Thinn MM, Hsueh CT, Hsueh CT
- 623** Exercise-induced anaphylaxis with an Ayurvedic drug as cofactor: A case report
Losa F, Deidda M, Firinu D, Martino MLD, Barca MP, Giacco SD
- 628** Diagnostic detection with cardiac tomography and resonance of extremely rare coronary anomaly: A case report and review of literature
Schicchi N, Fogante M, Giuseppetti GM, Giovagnoni A
- 636** Fatal meningococcal meningitis in a 2-year-old child: A case report
Mularski A, Žaba C
- 642** Perioperative topical ascorbic acid for the prevention of phacoemulsification-related corneal endothelial damage: Two case reports and review of literature
Lee CY, Chen HT, Hsueh YJ, Chen HC, Huang CC, Meir YJJ, Cheng CM, Wu WC

- 650** Application of computer-assisted navigation in treating congenital maxillomandibular syngnathia: A case report
Lin LQ, Bai SS, Wei M
- 656** Concomitant paraganglioma and thyroid carcinoma: A case report
Lin B, Yang HY, Yang HJ, Shen SY
- 663** Rare empty sella syndrome found after postoperative hypotension and respiratory failure: A case report
Guo P, Xu ZJ, Hu CE, Zheng YY, Xu DF
- 668** Use of tunnel endoscopy for diagnosis of obscure submucosal esophageal adenocarcinoma: A case report and review of the literature with emphasis on causes of esophageal stenosis
Liu S, Wang N, Yang J, Yang JY, Shi ZH
- 676** Intrauterine cystic adenomyosis: Report of two cases
Fan YY, Liu YN, Li J, Fu Y
- 684** Melanotic Xp11-associated tumor of the sigmoid colon: A case report
Wang G, Li GG, Zhu SM, Cai BJ, Yu PJ, Zhang CW

ABOUT COVER

Editorial Board Member of *World Journal of Clinical Cases*, Manel Sabate, MD, PhD, Associate Professor, Interventional Cardiology Department, Clinic University Hospital, Barcelona 08036, Spain

AIMS AND SCOPE

World Journal of Clinical Cases (*World J Clin Cases*, *WJCC*, online ISSN 2307-8960, DOI: 10.12998) is a peer-reviewed open access academic journal that aims to guide clinical practice and improve diagnostic and therapeutic skills of clinicians.

The primary task of *WJCC* is to rapidly publish high-quality Case Report, Clinical Management, Editorial, Field of Vision, Frontier, Medical Ethics, Original Articles, Meta-Analysis, Minireviews, and Review, in the fields of allergy, anesthesiology, cardiac medicine, clinical genetics, clinical neurology, critical care, dentistry, dermatology, emergency medicine, endocrinology, family medicine, gastroenterology and hepatology, etc.

INDEXING/ABSTRACTING

The *WJCC* is now indexed in PubMed, PubMed Central, Science Citation Index Expanded (also known as SciSearch®), and Journal Citation Reports/Science Edition. The 2018 Edition of Journal Citation Reports cites the 2017 impact factor for *WJCC* as 1.931 (5-year impact factor: N/A), ranking *WJCC* as 60 among 154 journals in Medicine, General and Internal (quartile in category Q2).

RESPONSIBLE EDITORS FOR THIS ISSUE

Responsible Electronic Editor: Ying-Na Bian

Proofing Editorial Office Director: Jin-Lei Wang

NAME OF JOURNAL

World Journal of Clinical Cases

ISSN

ISSN 2307-8960 (online)

LAUNCH DATE

April 16, 2013

FREQUENCY

Semimonthly

EDITORS-IN-CHIEF

Dennis A Bloomfield, Sandro Vento

EDITORIAL BOARD MEMBERS

<https://www.wjgnet.com/2307-8960/editorialboard.htm>

EDITORIAL OFFICE

Jin-Lei Wang, Director

PUBLICATION DATE

March 6, 2019

COPYRIGHT

© 2019 Baishideng Publishing Group Inc

INSTRUCTIONS TO AUTHORS

<https://www.wjgnet.com/bpg/gerinfo/204>

GUIDELINES FOR ETHICS DOCUMENTS

<https://www.wjgnet.com/bpg/GerInfo/287>

GUIDELINES FOR NON-NATIVE SPEAKERS OF ENGLISH

<https://www.wjgnet.com/bpg/gerinfo/240>

PUBLICATION MISCONDUCT

<https://www.wjgnet.com/bpg/gerinfo/208>

ARTICLE PROCESSING CHARGE

<https://www.wjgnet.com/bpg/gerinfo/242>

STEPS FOR SUBMITTING MANUSCRIPTS

<https://www.wjgnet.com/bpg/GerInfo/239>

ONLINE SUBMISSION

<https://www.f6publishing.com>

Sustained complete response to erlotinib in squamous cell carcinoma of the head and neck: A case report

Mie Mie Thinn, Chung-Tzu Hsueh, Chung-Tsen Hsueh

ORCID number: Mie Mie Thinn (0000-0001-7605-2526); Chung-Tzu Hsueh (0000-0002-7574-9863); Chung-Tsen Hsueh (0000-0002-1409-0030).

Author contributions: Hsueh CT performed clinical examination and collected data; Thinn MM, Hsueh CT and Hsueh CT designed the report, analyzed the data and wrote the paper; all authors read and approved the final manuscript.

Informed consent statement: Informed written consent was obtained from the surviving spouse of deceased patient for publication of this report and any accompanying images.

Conflict-of-interest statement: The authors declare that they have no conflict of interest.

CARE Checklist (2016) statement: The authors have read the CARE Checklist (2016), and the manuscript was prepared and revised according to the CARE Checklist (2016).

Open-Access: This article is an open-access article which was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: <http://creativecommons.org/licenses/by-nc/4.0/>

Mie Mie Thinn, Division of Hematology and Medical Oncology, Loma Linda Veterans Administration Medical Center, Loma Linda, CA 92357, United States

Chung-Tzu Hsueh, Department of Dentistry, Cathay General Hospital, Taipei City 106, Taiwan

Chung-Tsen Hsueh, Division of Medical Oncology and Hematology, Department of Medicine, Loma Linda University, Loma Linda, CA 92357, United States

Corresponding author: Chung-Tsen Hsueh, MD, PhD, Professor, Division of Medical Oncology and Hematology, Department of Medicine, Loma Linda University, 11175 Campus Street, CSP 11015, Loma Linda, CA 92357, United States. chsueh@llu.edu

Telephone: +1-909-5588107

Fax: +1-909-5580219

Abstract

BACKGROUND

Squamous cell carcinoma of head and neck (SCCHN) is the fifth most common cancer worldwide. Inhibition of epidermal growth factor receptor signaling has been shown to be a critical component of therapeutic option. Herein, we report a case of durable complete response to erlotinib.

CASE SUMMARY

An 81-year-old Caucasian male who presented with metastatic poorly differentiated squamous cell carcinoma of right cervical lymph nodes (levels 2 and 3). Imaging studies including (18)F-fluorodeoxyglucose positron emission tomography/computed tomography (CT) and contrast-enhanced CT scan of neck and chest did not reveal any other disease elsewhere. Panendoscopic examination with random biopsy did not reveal malignant lesion in nasopharynx, oropharynx, and larynx. He underwent modified neck dissection and postoperative radiation. Within 2 mo after completion of radiation, he developed local recurrence at right neck, which was surgically removed. Two mo after the salvage surgery, he developed a second recurrence at right neck. Due to suboptimal performance status and his preference, he started erlotinib treatment. He achieved partial response after first 2 mo of erlotinib treatment, then complete response after total 6 mo of erlotinib treatment. He developed severe skin rash and diarrhea including *Clostridium difficile* infection during the course of erlotinib treatment requiring dose reduction and eventual discontinuation. He remained in complete remission for more than two years after discontinuation of erlotinib.

CONCLUSION

We report a case of metastatic SCCHN achieving durable complete response from

[ses/by-nc/4.0/](#)

Manuscript source: Invited manuscript

Received: December 19, 2018

Peer-review started: December 20, 2018

First decision: January 5, 2019

Revised: January 12, 2019

Accepted: January 29, 2019

Article in press: January 29, 2019

Published online: March 6, 2019

erlotinib. Patient experienced skin rash and diarrhea toxicities which were likely predictors of his treatment response.

Key words: Squamous cell carcinoma of head and neck; Epidermal growth factor receptor; Erlotinib; Complete response; Skin rash; Tyrosine kinase inhibitor; Case report

©The Author(s) 2019. Published by Baishideng Publishing Group Inc. All rights reserved.

Core tip: We present a patient with recurrent/metastatic squamous cell carcinoma of head and neck who had durable complete response after completion of 6-mo erlotinib treatment. Patient experienced severe skin rash and diarrhea toxicities from erlotinib. The severity of these adverse effects has been shown to be predictors of treatment response from inhibitors of epidermal growth factor receptor.

Citation: Thinn MM, Hsueh CT, Hsueh CT. Sustained complete response to erlotinib in squamous cell carcinoma of the head and neck: A case report. *World J Clin Cases* 2019; 7(5): 616-622

URL: <https://www.wjgnet.com/2307-8960/full/v7/i5/616.htm>

DOI: <https://dx.doi.org/10.12998/wjcc.v7.i5.616>

INTRODUCTION

Squamous cell carcinoma of head and neck (SCCHN) is the fifth most common cancer worldwide with a global yearly incidence of more than 600000 new cases and around 300000 annual deaths in 2012^[1]. Early-stage disease is managed with either surgery or radiation. Two-third of patients present with locally advanced disease, and are generally treated with multimodality therapy, which commonly includes chemotherapy^[2]. Tumor control and survival in SCCHN remain unsatisfactory. Even for those who have achieved complete response after initial treatment, the incidence of local recurrence is 50%-60% and the incidence of distant metastases is 20%-30%. Systemic therapy is the mainstay for metastatic disease and unsalvageable recurrence to improve survival and quality of life.

With recent advances in cancer biology, there appear to be common molecular events in SCCHN that are biologically significant in cell survival and invasion, and could be used for therapeutic development such as epidermal growth factor receptor (EGFR). Overexpression of EGFR and its ligand have been reported in 80% to 90% of SCCHN tumors compared with levels in normal mucosa of patients without cancer^[3]. Increased EGFR expression has been reported to be a predictor of worse survival in SCCHN patients receiving surgery and chemotherapy^[4,5]. The two targeting strategies for inhibition of EGFR are small-molecule tyrosine kinase inhibitors (TKIs) and monoclonal antibodies directed against the receptor. Cetuximab (a monoclonal antibody against EGFR) has been approved by United States Food and Drug Administration as initial treatment of locally advanced SCCHN in combination with radiation therapy, as first-line treatment of recurrent or metastatic SCCHN in combination with platinum-based therapy plus 5-fluorouracil, and as a single agent for recurrent or metastatic SCCHN progressing from prior platinum-based therapy^[2]. Adding cetuximab to platinum-based chemotherapy with 5-fluorouracil as first-line treatment of recurrent or metastatic SCCHN significantly prolonged the median overall survival^[6]. Herein, we report a case of sustained complete response to erlotinib, a TKI of EGFR, as first-line treatment for recurrent SCCHN.

CASE PRESENTATION

Chief complaints

An 81-year-old Caucasian male presented with right neck mass.

History of present illness

He had no history of cigarette smoking or alcohol use.

History of past illness

He had an extensive history of multiple skin lesions removed from his scalp, arms, chest and back over 10-15 years prior to presentation. Most of the skin lesions were squamous cell carcinoma except one lesion was in-situ melanoma. Other significant past medical history included diabetes mellitus type II, hypertension, hypothyroidism and remote cerebrovascular accident with mild residual dysarthria and right central vision loss.

Personal and family history

Family history was pertinent for mother died of liver cancer at age 86.

Physical examination upon admission

Physical examination was unremarkable except right neck lymphadenopathy.

Imaging examinations

Imaging studies including (18)F-fluorodeoxyglucose positron emission tomography/computed tomography (CT) and contrast-enhanced CT (CECT) scan of neck and chest revealed right cervical lymphadenopathy. The panendoscopic examination of ENT field did not reveal any abnormality, and the random biopsy of nasopharynx, oropharynx, and larynx showed no evidence of malignancy. Excisional biopsy of right neck mass turned out to be metastatic poorly differentiated squamous cell carcinoma. He underwent right modified neck dissection with seventeen lymph nodes removed from levels 2, 3, 4 and 5. Two of the two lymph nodes from level 2 had evidence of metastatic carcinoma with the largest focus being 2.5 cm and extracapsular extension being present. Three of the five lymph nodes from level 3 were positive for metastatic carcinoma with largest focus being 0.7 cm and extracapsular extension being present. None of the lymph nodes from level 4 or 5 were found to have any metastatic carcinoma. Shortly after postoperative intensity-modulated radiotherapy to right neck with total dose of 66 Gy, he developed local recurrence at right neck and underwent salvage surgery.

FINAL DIAGNOSIS

Two mo after salvage surgery, he was found to have palpable skin nodules over the right neck. CECT scan of the neck showed interval increase in the soft tissue thickening and diffuse subcutaneous thickening of the right neck. Focal skin irregularity and interval development of submental, right submandibular adenopathy as well as adenopathy at the right inferior parotid compatible with recurrent/metastatic SCCHN were noted (Figure 1).

TREATMENT

Treatment options were discussed including platinum based chemotherapeutic regimen and targeted therapy. Due to patient's preference and literature support (see discussion), erlotinib 150 mg orally per day was started. He tolerated the treatment well initially, and achieved partial response after 2 mo of treatment. Subsequently, he developed grade 3 toxicities including skin rash and diarrhea with *Clostridium difficile* infection requiring dose reduction of erlotinib from 150 mg to 100 mg daily.

OUTCOME AND FOLLOW-UP

He eventually had complete response after total 6 mo of treatment (Figure 2). Erlotinib was discontinued due to intolerance. He remained free of recurrent disease for more than two years. Subsequently he succumbed to death due to postoperative complication with respiratory failure after resection of an ulcerating skin lesion at right clavicular head.

DISCUSSION

Our patient presented with SCCHN of unknown primary with cervical lymph node metastasis. His primary site of origin has never been identified over the course of the subsequent workup and follow-up. Retrospective analyses indicate SCCHN of unknown primary represents about 3% of newly diagnosed SCCHN^[7]. Most SCCHN of unknown primary may represent clinically occult oropharyngeal cancer. The

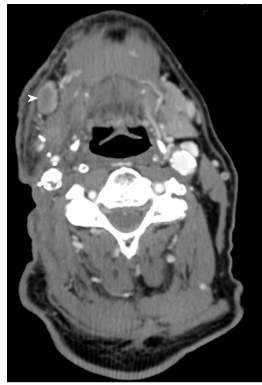


Figure 1 Contrast-enhanced computed tomography scan of neck prior to erlotinib treatment. Development of right neck focal skin irregularity and right submandibular lymphadenopathy (white arrowhead) consistent with metastases were noted.

management of SCCHN of unknown primary is aimed at curative for most patients; cervical lymphadenopathy indicates locally advanced disease and is amenable for multimodality treatment such as surgery and radiotherapy.

Our patient developed regional recurrence shortly after initial surgery with neck dissection and post-operative radiotherapy. Despite of salvage surgery, he developed recurrent disease at prior surgical site two mo later. Due to suboptimal performance status and patient's preference, he received systemic therapy with erlotinib, which is an EGFR TKI. EGFR TKIs inhibit EGFR and downstream signaling leading to apoptosis. Advantages of TKIs include the ease of oral administration. As shown in [Table 1](#), erlotinib, lapatinib, gefitinib and afatinib have been studied in phase II/III trials^[8-11]. Most of them showed promising activities in SCCHN, but failed to demonstrate improved survival compared to chemotherapy. A multicenter phase II study evaluated erlotinib in the treatment of recurrent or metastatic SCCHN showed the overall objective response rate of 4.3% in 115 patients. Forty-seven percent of patients received erlotinib at 150 mg daily throughout the entire study, 6% had dose escalations, and 46% required dose reductions and/or interruptions. Stable disease for a median duration of 16.1 wk was noted in 38% of patient. The median progression-free survival was 9.6 wk, and the median overall survival was 6 mo. Better overall survival was observed in patients who developed grade 2 or higher rash. Rash was the most common adverse event, observed in 79% of patients, followed by diarrhea, which was seen in 37% of patients. Most of the adverse events were mild to moderate^[8].

Retrospective analyses of clinical trials investigating EGFR TKIs in SCCHN have shown skin rash and diarrhea are common toxicities and severity of these side effects correlates with therapeutic responses^[12]. Skin rash has been shown to a predictor of response to EGFR TKI in patients with non-small cell lung cancer, likely due to skin injury caused by inhibition of EGFR signaling in epidermal cells^[13]. Many factors may affect severity of skin rash including genetic variations in EGFR and metabolism of EGFR TKI^[14]. Higher drug levels may result from polymorphisms in metabolizing enzymes such as cytochrome P450 family. The severe skin rash and diarrhea toxicities seen in our case are likely predictors of good treatment response from erlotinib.

In non-small cell lung carcinoma, patients with activating mutations in the EGFR tyrosine kinase domain are sensitive to EGFR TKIs. However, the incidence of this type of EGFR mutations is low and fails to predict sensitivity to EGFR TKIs in SCCHN^[15]. Van Allen *et al*^[16] reported a patient with locally advanced SCCHN achieving a near-complete pathological response after 13 d of neoadjuvant erlotinib treatment. After surgical resection, histologic evaluation revealed 2 residual foci (approximately 2 mm each) of invasive, moderately differentiated squamous cell carcinoma within the primary site but there was no evidence of lymph node metastasis. The patient did not receive adjuvant therapy and had no evidence of disease recurrence 24 mo postoperatively. Whole exome sequencing of the pretreatment tumor revealed a MAPK1 E322K somatic mutation, which was further implicated in mediating erlotinib sensitivity. This response to erlotinib occurred in the context of an activating somatic MAPK1 E322K mutation, which led to increased EGFR ligand production and EGFR activation in preclinical studies^[17].

Recent adoption of next-generation sequencing in genomic testing of tumor has provided opportunity to advance precision medicine. Dumbrava *et al*^[18] reported a case of metastatic SCCHN who progressed on a phase II clinical trial with erlotinib,

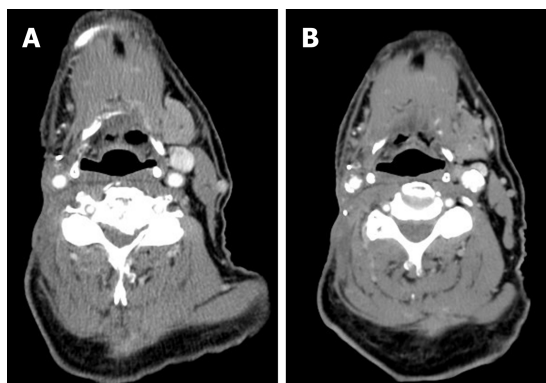


Figure 2 Contrast-enhanced computed tomography scan of neck. A: After completion of 6-mo erlotinib treatment; B: Seven mo after discontinuation of erlotinib treatment. Both showed resolution of right neck focal skin irregularity and right submandibular lymphadenopathy as shown in [Figure 1](#).

docetaxel and cisplatin. Patient's tumor was subsequently biopsied and found to have amplification of fibroblast growth factor genes by next-generation sequencing. Patient was treated with fibroblast growth factor receptor inhibitor on clinical trial and achieved complete response for about 9 mo^[18].

CONCLUSION

We report a case of durable complete response to erlotinib in SCCHN. Patient experienced skin rash and diarrhea toxicities which were likely predictors of his treatment response. The use of next-generation sequencing in genomic profiling of tumor samples may select patients who will likely respond to targeted therapy.

Table 1 Outcome of single-agent epidermal growth factor receptor tyrosine kinase inhibitor studies in recurrent/metastatic squamous cell carcinoma of head and neck

Agent	Phase/publication	Control arm	RR (%)	OS (mo)
Erlotinib	II/Soulieres ^[8]	None	4.3	6
Lapatinib	II/de Souza ^[11]	None	0	9.6-5.2 (lower on prior EGFR inhibitor exposure)
Gefitinib	III/Stewart ^[9]	Methotrexate	2.7 (250 mg/d); 7.6 (500 mg/d)	5.6 (250 mg/d); 6 (500 mg/d)
Afatinib	III/Machiels ^[10]	Methotrexate	10	6.8

RR: Response rate; OS: Overall survival; EGFR: Epidermal growth factor receptor.

REFERENCES

- 1 Ferlay J, Soerjomataram I, Dikshit R, Eser S, Mathers C, Rebelo M, Parkin DM, Forman D, Bray F. Cancer incidence and mortality worldwide: sources, methods and major patterns in GLOBOCAN 2012. *Int J Cancer* 2015; **136**: E359-E386 [PMID: 25220842 DOI: 10.1002/ijc.29210]
- 2 Marur S, Forastiere AA. Head and Neck Squamous Cell Carcinoma: Update on Epidemiology, Diagnosis, and Treatment. *Mayo Clin Proc* 2016; **91**: 386-396 [PMID: 26944243 DOI: 10.1016/j.mayocp.2015.12.017]
- 3 Grandis JR, Tweardy DJ. Elevated levels of transforming growth factor alpha and epidermal growth factor receptor messenger RNA are early markers of carcinogenesis in head and neck cancer. *Cancer Res* 1993; **53**: 3579-3584 [PMID: 8339264]
- 4 Rubin Grandis J, Melhem MF, Gooding WE, Day R, Holst VA, Wagener MM, Drenning SD, Tweardy DJ. Levels of TGF- α and EGFR protein in head and neck squamous cell carcinoma and patient survival. *J Natl Cancer Inst* 1998; **90**: 824-832 [PMID: 9625170 DOI: 10.1093/jnci/90.11.824]
- 5 Etienne MC, Pivot X, Formento JL, Bensadoun RJ, Formento P, Dassonville O, Francoual M, Poissonnet G, Fontana X, Schneider M, Demard F, Milano G. A multifactorial approach including tumoural epidermal growth factor receptor, p53, thymidylate synthase and dihydropyrimidine dehydrogenase to predict treatment outcome in head and neck cancer patients receiving 5-fluorouracil. *Br J Cancer* 1999; **79**: 1864-1869 [PMID: 10206306 DOI: 10.1038/sj.bjc.6690297]
- 6 Vermorken JB, Mesia R, Rivera F, Remenar E, Kawecki A, Rottey S, Erfan J, Zabolotnyy D, Kienzer HR, Cupissol D, Peyrade F, Benasso M, Vynnychenko I, De Raucourt D, Bokemeyer C, Schueler A, Amellal N, Hitt R. Platinum-based chemotherapy plus cetuximab in head and neck cancer. *N Engl J Med* 2008; **359**: 1116-1127 [PMID: 18784101 DOI: 10.1056/NEJMoa0802656]
- 7 Strojanc P, Ferlito A, Medina JE, Woolgar JA, Rinaldo A, Robbins KT, Fagan JJ, Mendenhall WM, Paleri V, Silver CE, Olsen KD, Corry J, Suárez C, Rodrigo JP, Langendijk JA, Devaney KO, Kowalski LP, Hartl DM, Haigentz M, Werner JA, Pellitteri PK, de Bree R, Wolf GT, Takes RP, Genden EE, Hinni ML, Mondin V, Shaha AR, Barnes L. Contemporary management of lymph node metastases from an unknown primary to the neck: I. A review of diagnostic approaches. *Head Neck* 2013; **35**: 123-132 [PMID: 22034046 DOI: 10.1002/hed.21898]
- 8 Soulieres D, Senzer NN, Vokes EE, Hidalgo M, Agarwala SS, Siu LL. Multicenter phase II study of erlotinib, an oral epidermal growth factor receptor tyrosine kinase inhibitor, in patients with recurrent or metastatic squamous cell cancer of the head and neck. *J Clin Oncol* 2004; **22**: 77-85 [PMID: 14701768 DOI: 10.1200/JCO.2004.06.075]
- 9 Stewart JS, Cohen EE, Licitra L, Van Herpen CM, Khorprasert C, Soulieres D, Vodvarka P, Rischin D, Garin AM, Hirsch FR, Varella-Garcia M, Ghiorghiu S, Hargreaves L, Armour A, Speake G, Swaisland A, Vokes EE. Phase III study of gefitinib compared with intravenous methotrexate for recurrent squamous cell carcinoma of the head and neck [corrected]. *J Clin Oncol* 2009; **27**: 1864-1871 [PMID: 19289630 DOI: 10.1200/JCO.2008.17.0530]
- 10 Machiels JP, Haddad RI, Fayette J, Licitra LF, Tahara M, Vermorken JB, Clement PM, Gauler T, Cupissol D, Grau JJ, Guigay J, Caponigro F, de Castro G, de Souza Viana L, Keilholz U, Del Campo JM, Cong XJ, Ehrnrooth E, Cohen EE; LUX-H&N 1 investigators. Afatinib versus methotrexate as second-line treatment in patients with recurrent or metastatic squamous-cell carcinoma of the head and neck progressing on or after platinum-based therapy (LUX-Head & Neck 1): an open-label, randomised phase 3 trial. *Lancet Oncol* 2015; **16**: 583-594 [PMID: 25892145 DOI: 10.1016/S1470-2045(15)70124-5]
- 11 de Souza JA, Davis DW, Zhang Y, Khattri A, Seiwert TY, Aktolga S, Wong SJ, Kozloff MF, Nattam S, Lingen MW, Kunnavakkam R, Stenson KM, Blair EA, Bozeman J, Dancay JE, Vokes EE, Cohen EE. A phase II study of lapatinib in recurrent/metastatic squamous cell carcinoma of the head and neck. *Clin Cancer Res* 2012; **18**: 2336-2343 [PMID: 22371453 DOI: 10.1158/1078-0432.CCR-11-2825]
- 12 Cohen EE, Halpern AB, Kasza K, Kocherginsky M, Williams R, Vokes EE. Factors associated with clinical benefit from epidermal growth factor receptor inhibitors in recurrent and metastatic squamous cell carcinoma of the head and neck. *Oral Oncol* 2009; **45**: e155-e160 [PMID: 19586795 DOI: 10.1016/j.oraloncology.2009.05.637]
- 13 Ruan Y, Jiang J, Guo L, Li Y, Huang H, Shen L, Luan M, Li M, Du H, Ma C, He L, Zhang X, Qin S. Genetic Association of Curative and Adverse Reactions to Tyrosine Kinase Inhibitors in Chinese advanced Non-Small Cell Lung Cancer patients. *Sci Rep* 2016; **6**: 23368 [PMID: 26988277 DOI: 10.1038/srep23368]
- 14 Liu S, Kurzrock R. Toxicity of targeted therapy: Implications for response and impact of genetic polymorphisms. *Cancer Treat Rev* 2014; **40**: 883-891 [PMID: 24867380 DOI: 10.1016/j.ctrv.2014.05.003]
- 15 Perisanidis C. Prevalence of EGFR Tyrosine Kinase Domain Mutations in Head and Neck Squamous Cell Carcinoma: Cohort Study and Systematic Review. *In Vivo* 2017; **31**: 23-34 [PMID: 28064216 DOI: 10.21873/invivo.11020]
- 16 Van Allen EM, Lui VW, Egloff AM, Goetz EM, Li H, Johnson JT, Duvvuri U, Bauman JE, Stransky N, Zeng Y, Gilbert BR, Pendleton KP, Wang L, Chiosea S, Sougnez C, Wagle N, Zhang F, Du Y, Close D,

- Johnston PA, McKenna A, Carter SL, Golub TR, Getz G, Mills GB, Garraway LA, Grandis JR. Genomic Correlate of Exceptional Erlotinib Response in Head and Neck Squamous Cell Carcinoma. *JAMA Oncol* 2015; 1: 238-244 [PMID: 26181029 DOI: 10.1001/jamaoncol.2015.34]
- 17 **Wen Y**, Li H, Zeng Y, Wen W, Pendleton KP, Lui VW, Egloff AM, Grandis JR. MAPK1E322K mutation increases head and neck squamous cell carcinoma sensitivity to erlotinib through enhanced secretion of amphiregulin. *Oncotarget* 2016; 7: 23300-23311 [PMID: 27004400 DOI: 10.18632/oncotarget.8188]
- 18 **Dumbrava EI**, Alfattal R, Miller VA, Tsimberidou AM. Complete Response to a Fibroblast Growth Factor Receptor Inhibitor in a Patient With Head and Neck Squamous Cell Carcinoma Harboring FGF Amplifications. *JCO Precision Oncology* 2018; 1-7 [DOI: 10.1200/PO.18.00100]

P- Reviewer: Bhalla AS, Dasgupta S

S- Editor: Ji FF **L- Editor:** A **E- Editor:** Bian YN





Published By Baishideng Publishing Group Inc
7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA
Telephone: +1-925-2238242
Fax: +1-925-2238243
E-mail: bpgoffice@wjgnet.com
Help Desk: <https://www.f6publishing.com/helpdesk>
<https://www.wjgnet.com>

