

TOPICAL TREATMENT OF BASAL CELL CARCINOMA WITH NEOMYCIN

P. Cuevas¹, J. M. Arrazola²

¹Department of Research, ²Department of Dermatology, Hospital Ramón y Cajal. Madrid, Spain

Abstract: Basal cell carcinoma (BCC) is the most common skin cancer, occurring more frequently than malignancies of any other tissue or organ, either individually or in total. Medical treatment modalities of BCC offer cost reduction and clinical advantages in selected cases. Neomycin has been reported to have an important role on proliferation of endothelial cells and neoplastic cells. This finding may lead to new strategies for the therapeutic use of agents which block FGF activities in disease states associated with enhanced keratinocyte proliferation. We report here a case of BCC treated with neomycin 5% cream that induced a regression of BCC.

Key words: Basal cell carcinoma. Neomycin.

INTRODUCTION

Basal cell carcinoma (BCC), characterized by a non-aggressive behaviour, is the most common cancer among humans, and its incidence is increasing [1,2]. The most commonly used treatment modalities include simple excision, Mohs micrographic surgery, curettage and electrodesiccation, cryosurgery and irradiation therapy [2]. However, different topical medical options as chemotherapy, immunotherapy and photodynamic

therapy, have been proposed for BCC, according to number, sites of distribution and size of lesions, patient age, ease of treatment, costs and cosmetic results [3]. We previously reported that the aminoglycoside antibiotic, neomycin, exerts significant antiproliferative and antiangiogenic effects in gliomas [4-7] acting as a fibroblast growth factor (FGF) inhibitor [8]. FGF appeared to have a role in maintaining epidermal integrity, as well as in keratinocyte proliferation, wound healing [9,10] and skin carcinogenesis [11]. Furthermore, immunohistochemical studies revealed that endogenous FGF was localized within the cytoplasm of keratinocytes in BCC [12]. Since FGF family of proteins has an important function in proliferative- and angiogenesis-related diseases such as in malignant cutaneous neoplasms [13-15], we assessed the effect of topical neomycin in BCC. In the present study, the efficacy of neomycin application in BCC for two weeks is reported and discussed in a case report.

CASE REPORT

A 45-year-old man presented with a nodular BCC of more than 1 year duration in the left inferior eyelid. He was asymptomatic with no associated pain or bleeding. Patient medical history was unremarkable.

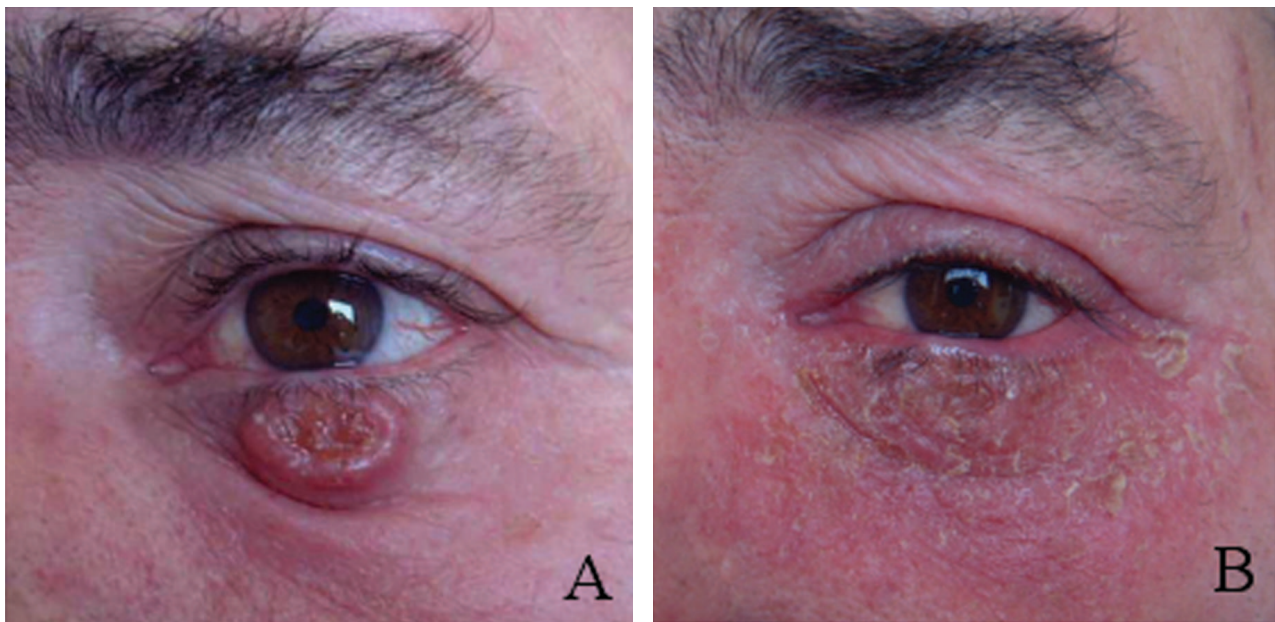


Fig. 1. Patient with BCC, (A) before treatment, (B) at two weeks of treatment with neomycin 5% cream twice a day.

After discussing the risks and benefits of the different treatment options, the patient declined surgical excision. A physical examination and photographic documentation were obtained at baseline and after treatment. He was started on neomycin sulphate (Sigma, St. Louis, USA) 5% cream twice daily for two weeks. As Figure 1 shows, treatment with neomycin induced a regression of the BCC. No recurrence was observed after 8 weeks of follow-up.

DISCUSSION

Medical treatment modalities for BCC may offer cost and clinical advantages in selected cases, such as tumours mainly located in low-risk areas, difficult sites on which to operate (nose, ears, eyelids), cases with a high number of neoplasms or otherwise inoperable patients [3]. We report a case of BCC regression treated with neomycin 5% cream. The mechanism of action of neomycin in the treatment of BCC is not known. Molecular effects similar to those observed in the case of gliomas [4-7], as antiproliferative and antiangiogenesis effects as well as proapoptotic activity, may mediate the therapeutic effect of neomycin reported here for BCC. Furthermore, this case report suggests that neomycin may be a potential treatment option for patients who are poor candidates to surgery or who face disfigurement and functional impairment from resection. Further randomized controlled trials are needed to evaluate the efficacy of neomycin for treatment of BCC.

REFERENCES

1. Diepgen TL, Mehler V (2002) The epidemiology of skin cancer. *Br J Dermatol* 14 (Suppl 61): 1-6
2. Wennberg AM (2000) Basal cell carcinoma: new aspects of diagnosis and treatment. *Acta Derm Venerol Suppl (Stockh)* 209: 5-25
3. Thissen MR, Newman MHA, Schouten LJ (1999) A systemic review of treatment modalities for primary basal cell carcinomas. *Arch Dermatol* 135: 1177-1183
4. Cuevas P, Diaz-Gonzalez D, Dujovny M (2002) Antiproliferative effect of neomycin in glioma cells. *Neurol Res* 24: 389-391
5. Cuevas P, Carceller F, Díaz-González D, Cuevas B, Fernández A, García-Gómez I, Dujovny M (2002) Inhibition of rat glioma growth by neomycin. Preliminary report. *Neurol Res* 24:522-524
6. Cuevas P, Diaz-Gonzalez D, Carceller F, Dujovny M (2003) Dual blockade of mitogen-activated protein kinases ERK-1 (p42) and ERK-2 (p44) and cyclic AMP response element binding protein (CREB) by neomycin inhibits glioma cell proliferation. *Neurol Res* 25: 13-16
7. Cuevas P, Díaz-González D, Dujovny M (2003) Antiproliferative action of neomycin is associated with inhibition of cyclin D1 activation in glioma cells. *Neurol Res* 25: 691-693
8. Hu G-F (2001) Neomycin inhibits the angiogenic activity of fibroblast and epidermal growth factors. *Biochem Biophys Res Commun* 287: 870-874
9. Halaban R, Moellmann G (1994) Fibroblast growth factors. In: *Epidermal Growth Factors and Cytokines* (Luger TA, Schwarz T. Eds) New York. Marcel Dekker 1994: 273-289
10. O'Keeffe EJ, Chin ML, Payne RE (1988) Stimulation of growth of keratinocytes by basic fibroblast growth factor. *J Invest Dermatol* 90: 677-679
11. Yamamoto N, Matsutani S, Yoshitake Y, Nishikawa K (1991) Immunohistochemical localization of basic fibroblast growth factor in A431 epidermoid carcinoma cells. *Histochemistry* 96: 479-485
12. Grimme Hu, Termeer CC, Bennett KI, Weiss JM, Schopf E, Aruffo A, Simon JC (1999) Colocalization of basic fibroblast growth factor and CD44 isoforms containing the variable spliced exon v3 (CD44v3) in normal skin and in epidermal skin cancers. *Br J Dermatol* 141:824-832
13. Halaban R (1996) Growth factors and melanomas. *Semin Oncol* 23: 673-681
14. Czubyko F, Liandet-Coopman ED, Aigner A, Tuveson AT, Berchem GJ, Wellstein A (1997) A secreted FGF-binding protein can serve as the angiogenic switch in human cancer. *Nat Med* 3: 1137-1140
15. Arbiser JL, Byers R, Cohen C, Arbeti J (2000) Altered basic fibroblast growth factor expression in common epidermal neoplasms: examination with in situ hybridization and immunohistochemistry. *J Am Acad Dermatol* 42: 973-977

Received: December 15, 2004 / Accepted: January 28, 2005

Address for correspondance:

Dr. Pedro Cuevas
 Servicio de Histología
 Departamento de Investigación
 Hospital Ramón y Cajal
 Ctra. de Colmenar, km. 9.100
 E-28034-Madrid - Spain
 Tel.: +3491-336 82 90
 Fax: +3491-336 82 90
 e-mail: pedro.cuevas@hrc.es