Takaokamycin

1. Discovery, producing organism and structure¹⁻⁵⁾

Takaokamycin was isolated from the culture broth of *Streptomyces* strain AC-1978 while screening for new antibiotics. It exhibited antibacterial activity against some Gram-positive bacteria. Though takaokamycin has been suggested to be a peptide containing three amino acids, its total structure was not elucidated¹). After a few years, N. Andres *et al.* isolated a compound from the culture broth of *Streptomyces griseoflavus* W-384 and named it hormaomycin. Its structure was elucidated and the physico-chemical properties of hormaomycin were quite similar to those of takaokamycin, except high-field ¹H-NMR signals^{2,3}). Recently, ¹H- and ¹³C-NMR spectra of takaokamycin were re-measured and data showed that takaokamycin was identical to hormaomycin⁴).



Takaokamycin (Hormaomycin)

2. Physical data

White powder. $C_{55}H_{69}N_{10}O_{14}Cl$; mol wt 1129.67. Sol. in DMSO, MeOH, acetone, CHCl₃. Insol. in H₂O, hexane.

3. Biological activity

1) Antibacterial activity¹⁾

Takaokamycin showed antibacterial activity against *Bacillus cereus* IFO3001 (MIC 12.5 μ g/ml) and *Micrococcus luteus* ATCC 9341 (MIC 1.56 μ g/ml).

2) Antimalarial activity against FCR3 and K1 strains of Plasmodium falciparum^{4,6)}

	IC ₅₀ (µM)
FCR3 strain (chloroquine sensitive)	1.21
K1 strain (chloroquine resistant)	0.59
Cytotoxicity (MRC-5 cells)	53.9

3) Other biological activity²⁾

Takaokamycin (hormaomycin) initiated the development of aerial mycelia and stimulated antibiotic production in some *Streptomyces* strains.

4. Biosynthesis^{7,8)}

The biosynthetic gene cluster for hormaomycin (takaokamycin) was identified from *Streptomyces griseoflavus* W-384 and it proposed to be synthesized by noribosomal peptide synthetase. The transcriptional regulators, hrmA and hrmB, are positive regulators in the biosynthesis.

5. References

- 1. [286] S. Ōmura et al., J. Antibiot. 37, 700-705 (1984)
- 2. N. Andres *et al.*, *Helv. Chim. Acta* **72**, 426-437 (1989)
- 3. E. Rössner *et al.*, *Angew. Chem. Int. Ed.* **29**, 64-65 (1990)
- 4. K. Otoguro, *Kagaku to Kyoiku* **50**, 355-359 (2002)
- 5. N. Andres *et al.*, *Z. Naturforsch.*, *C: Biosci.* **45**, 851-855 (1990)
- 6. [829] K. Otoguro *et al.*, J. Antibiot. **56**, 322-323(2003)
- 7. I. Höfer *et al.*, *Chem. Biol.* **18**, 381-391 (2011)
- 8. X. Cai et al., Chem. Biol. 20, 839-846 (2013)