Virustomycin A ©

1. Discovery, producing organism¹⁾ and structure²⁾

Virustomycin A was isolated from the culture broth of the actinomycete strain AM-2604 and recognized by the plaque reduction method as an antiviral substance.



Streptomyces sp. AM-2604

2. Physical data^{1,3)}

Lipophilic pale-yellow needles. $C_{48}H_{71}NO_{14}$; mol wt 885.49. Sol. in CHCl₃, EtOAc, acetone. Slightly sol. in MeOH, EtOH. Insol. in H_2O , hexane.

3. Biological activity $^{1,2,4)}$

1) Virustomycin A diminished 50% of plaque formed by both RNA and DNA viruses at a very low concentration (3 ng/ml). The MIC of virustomycin A against *Trichomonas foetus* is 1.6 μ g/ml.^{1,2)}

2) Anti-trypanosomal activity⁴⁾

Virustmycin showed anti-trypanosomal activity against *Trypanosoma b. brusei* GUTat 3.1 strain with IC_{50} of 0.45 ng/ml.

4. Mode of action³⁾

Virustomycin A is believed to interfere with the formation of a phosphate donor(s), which is required for the organism nucleotide (UMP, UDP and UTP) formation.

Virustomycin A Precursor			Nucleotide formed (% of control)				
added (µg/ml		uracil	Uridine	UMP	UDP	UTP*	Total nucleotide
[³ H]Uridine	0	100 (704cpm)	100 (1,175)	100 (3,365)	100 (5,789)	100 (6,589)	100 (15,743)
	0.1	103	90	30	48	35	38
[³ H]Uracil	1.0 0 0.1 1.0	57 100 (3,050) 83 91	64 100 (13) 72 79	23 100 (2,625) 63 48	19 100 (5,168) 53 42	27 100 (8,505) 60 53	23 100 (16,298) 58 48

^{*}Contains UDP-sugars

5. References

- 1. [255] S. Ōmura et al., J. Antibiot. 35, 1632-1637 (1982)
- 2. [278] S. Ōmura et al., J. Antibiot. 36, 1783-1786 (1983)
- 3. [276] S. Ōmura et al., J. Antibiot. **36**, 1755-1761 (1983)
- 4. [993] K. Otoguro et al., J. Antibiot. 61, 372-378 (2008)