

Compound	Tracheal chain ED ₅₀ in mcg.	Rat uterus ED ₅₀ in mcg
A - 1	2.19	2.45
A - 2	4.36	4.37
Isoproterenol	2.09	2.75
A - 3	8.91	9.33
A - 4	4.00	7.24
Isoproterenol	2.13	1.51

In conclusion, all compounds exhibited beta adrenergic receptor stimulant activity and compound A-1 was as potent as Isoproterenol. All the compounds showed a high degree of selectivity for the beta₂ receptor sub-type, however, they also exhibited

a small degree of cardiac beta receptor blocking activity at the doses studied.

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Synthesis and Antimicrobial activity of some new Imidazolones having Thymolmoiety

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Received 4th October 1994

Some new-1-(2'-Isopropyl-4'-nitroso-5' methyl phenoxy-acetamido)-2-methyl/phenyl-4-arylidene-5-imidazolones were prepared by reacting 4-nitroso hydrazinocarbonyl methyl thymol with preformed azalactone. The structure of the compounds have been confirmed by IR, PMR and elemental analysis. The products were screened for their antimicrobial activity. Some of the products exhibited comparable antimicrobial activity with standard drugs at same concentration.

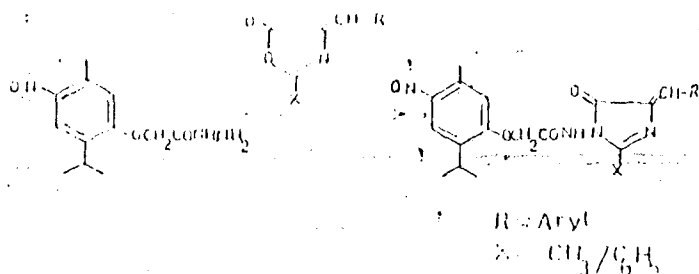
IMIDAZOLONE¹⁻⁴ derivatives have wide range of biological activities as thymol⁵⁻⁷ derivatives have been found to possess a broad pharmacological spectrum.

Reaction to acetic anhydride, aromatic aldehydes with hippuric acid or acetyl glycine in the presence of sodium acetate has been known to pro-

duce azalactone⁸ (I). I on refluxing with 4-nitroso hydrazinocarbonyl methyl thymol⁹ in 1:1 ratio in the presence of pyridine yielded the title compound (II) (Table-I). The latter was synthesised by condensation of 4-nitrosothymol and ethyl-chloro acetate followed by the reaction with hydrazine-hydrate.

The melting points were uncorrected. The IR(KBr) spectra were recorded on a Shimadzu-435

Scheme



infrared spectro-photometer. The PMR spectra were recorded in CDCl₃ on Hitachi R-1200 (60 MHz) TMS as internal reference. Chemical Shifts are expressed in τ (ppm). Elemental analyses are quite comparable with their structures.

1-(2'-Isopropyl-4'-nitroso-5' -methyl phenoxy acetamido) - 2- phenyl-4-(p-methoxybenzylidene)-5- imidazolinone (II) was prepared from a mixture of (I) 0.01 M) and 2-phenyl-4-p)-methoxy- benzylidene- 5-oxolone (0.01 M), pyridine (10 ml) which was refluxed for 6 hrs. The product was isolated and crystallised from ethanol-water, m.p. 124°C, yield 68% IR (KBr) $\nu_{cm^{-1}}$: 1250 (C-O-C str.), 1640 (C=O str. amide), 1620 (N-H def + C-N str.), 1320 (C-N str.), 3170 (N-H str.), 1510 (C-NO str.), 1760, 1630 (C=O + C=N imidazolinone). PMR (CDCl₃) δ : 1.2-1.3 (d, 6H, (CH₃)₂ CH), 2.27 (6, 3H + 2H, CH₃ + OCH₂) 3-3.5 (m, 1H, CH- CH₃)₂, 4-005 (5, 3H, OCH₃), 6-907-7.197 (m, 6H, Ar-H) 8.2 (5, 1H, -C=CH-R), 9.0 (S (br), 1H, NH).

Table I: Physical data of 1-(2'-Isopropyl-4'-nitroso-5'-methyl-phenoxyacetamido)-2 methyl/phenyl-4- arylidene-5-imidazolines

Compound	R	X=CH ₃	X=C ₆ H ₅
		M.P.°C II a-n	M.P.°C II a'-n'
IIa-a'	C ₆ H ₅	138	127
IIb-b'	3-NH ₂ C ₆ H ₄	135	—
IIc-c'	2-ClC ₆ H ₅	121	129
IId-d'	4-ClC ₆ H ₄	142	135
IIE-e'	C ₄ H ₃ O	148	151
IIf-f'	2-OHC ₆ H ₄	118	141
IIg-g'	4-OHC ₆ H ₄	210	138
IIh-h'	4-OCH ₃ C ₆ H ₄	131	124
IIi-i'	4-OH-3-OCH ₃ -C ₆ H ₃	129	144
IIj-j'	4-N, N(CH ₃) ₂ C ₆ H ₄	129	—
IIk-k'	2-NO ₂ C ₆ H ₄	195	132
IIl-l'	3-NO ₂ C ₆ H ₄	185	117
IIm-m'	4-NO ₂ C ₆ H ₄	—	126
II n-n'	C ₆ H ₅ CH=CH	140	181

Table II
Antimicrobial activity of 1-(2'-Isopropyl-4'-nitroso-5'-methylphenoxy-acetamido)-2 methyl/phenyl-4-arylidene-5-imidazolinones

No.	R	X	Antibacterial activity				Antifungal activity
			Zone of inhibition in mm.				Zone of inhibition in mm.
			S. citrus	B. mega	E.coli	S.typhosa	A. niger
II _d	4-ClC ₆ H ₄	CH ₃	22	19	21	14	14
II _i	4-OH-3-OCH ₃ -	CH ₃	22	12	27	13	13
II	C ₆ H ₃						
II _a '	C ₆ H ₅	C ₆ H ₅	22	14	21	13	14
II _d '	4-ClC ₆ H ₄	C ₆ H ₅	22	13	23	11	13
II _i '	4-OH-3-OCH ₃ C ₆ H ₃	C ₆ H ₅	22	16	22	12	14
II _k '	2-NO ₂ C ₆ H ₄	C ₆ H ₅	20	16	19	14	13
	Ampicillin (50 µg)		22	17	24	17	—
	Chloramphenicol (50 µg)		24	26	24	25	—
	Norfloxacin (50 µg)		33	23	26	28	—
	Grecioflavin (50 µg)		—	—	—	—	23

Similarly, other imidazolinones were prepared. The physical constant are recorded in Table-1.

All the compounds (II_{a-y}) were screened for their antimicrobial activity against gram positive **Staphylococcus citrus**, **Bacillus megaterium**, gram negative **Escherichia coli**, **Salmonella tyohosa** and antifungal activity against **Asperguillus niger** at a concentration of 50 µg using DMF as solvent. The zone of inhibition was measured in mm. and are presented in Table-1. The activity was compare with standard drugs such as ampicillin, chloroamphenical, norfloxacin, griseofulvin at same concentration.

From the test results, it is observed that Imidazolinones of type (II_{a-x}) were found to be less active against various strains of bacteria. However, compounds II_d (22 mm), II_i (92 mm), II_n (22 mm), II_p (22 mm), II_v (22 mm), II_u (22 mm), exhibited comparable activity with the standard drug ampicillin (22 mm) at the same concentration against **Staphylococcus citrus**. In case of antifungal activity, com-

pounds (II_{a-y}) showed less activity again **Aspergillus niger** in comparision to standard drug griseofulvin.

The authors are thankful to Dr. A.R. Parikh, Professor and Head of the Chemistry Department, Saurashtra University, Rajkot for providing research facilities.

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In Vivo Antisnake Venom Activity of A Furanoid Diterpene from *Aristolochia albida* Duch (Aristolochiaceae)

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Received 5 January 1995

The antisnake venom activity of a furanoid diterpene lactone isolated from the rhizome of *Aristolochia albida* (Family : Aristolochiaceae) was evaluated. The whole animal (*in vivo*) studies were conducted using the mortality of male Swiss albino mice after Intra-peritoneal (i.p.) injection of lethal doses (LD₁₀₀), 8.75 mg/kg and 4.20 mg/kg of venoms of *Naja nigricollis* (spitting cobra) and *Bitis arietans* (puff-adder) respectively. The diterpene was found to significantly reduce the toxic symptoms and protect the mice against the lethal doses of these two snake species commonly found in Northern Nigeria. However, the compound is more effective against the venom of *N.nigricollis* (ED₅₀=45 mg/kg) than that of *B.arietans* (ED₅₀=74 mg/kg).

ARISTOLOCHIA ALBIDA is a climbing shrub commonly found in the tropical West Africa¹ and is used in several gastro- intestinal disorders². There are reports that the rhizomes are used in skin diseases and against snake bites^{3,4}. The isolation of fargesin (neolignan) was reported earlier⁵. Recently, the isolation of phytosterols and glucose has been reported⁶. No pharmacological work on this medicinal plant has been reported so far to validate the claims of the folk-loric uses except for the molluscidal activity of the plant.⁷ Ethnopharmacological reports from northern Nigeria about the use of the rhizomes of this plant against snake bites⁸ have

prompted us to investigate the plant phytochemically and pharmacologically. This communication reports the antisnake venom activity of a diterpene isolated from *A.albida*.

The plant was collected in mid-July and authenticated by the Ahmadu Bello University herbarium, Zaria. The air dried, powder of the rhizomes was defatted with petroleum ether (b.p.60- 80°) and then extracted with methanol in a Soxhelt apparatus. The column chromatography (silica gel) of the methanolic extract furnished the diterpene which upon crystallisation from chloroform/methanol gave fine colorless needlelike colourless needles, m.p. 182°.

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