TRANSACTIONS OF THE ROYAL SOCIETY OF TROPICAL MEDICINE AND HYGIENE, VOL. 74, NO. 4, 1980

# **Brief Communications**

# Activity of precocenes on the Chagas's disease vector, Panstrongylus megistus

### A. M. DE OLIVEIRA FILHO, R. PINCHIN, C. E. SANTOS

Núcleo de Pesquisas de Produtos Naturais, Centro de Ciências da Saúde, Universidade Federal do Rio de Janeiro, Rio de Janeiro, Brasil

AND

## W. S. BOWERS

## Dept. of Entomology, Entomology-Plant Pathology Laboratory, Cornell University, Geneva, N.Y. 14456, USA

The continued use of the organochlorine insecticides, HCH and dieldrin, for triatomine control is at present under review. Several organophosphorus, carbamate and pyrethroid insecticides are currently being tested and one, malathion, has already been introduced in the national vector control campaign in Argentina. It also has been suggested that juvenile hormone analogues (JHAs) may be used as control agents (GILBERT, 1976). However, as triatomine bugs are most susceptible to JHAs only during a few weeks of the fifth instar, their practical use is somewhat limited (PINCHIN et al., 1978a). This may not be so for another class of compounds with insect growth regulatory activity-the plant-derived precocenes. In 1976, Bowers announced the discovery of such compounds with anti-juvenile hormone activity. Their principal effects on Oncopeltus fasciatus (Dallas) were the induction of precocious metamorphosis, and anti-gonadotropic and ovicidal activity (Bowers 1976, 1977; Bowers et al., 1976). As we have noted some similarity in the structure-activity relationships for JHAs for O. fasciatus and Panstrongylus megistus (Burmeister), it was considered feasible that the precocenes might also exhibit anti-juvenile hormone activity on P. megistus and other triatomine bugs of importance in the transmission of Chagas's disease. We describe here the results of initial experiments with the precocenes and discuss their possibilities as future vector control agents.

### Materials and Methods

The specimens of *P. megistus* used in these experiments were from a laboratory colony maintained at  $27 \pm 2^{\circ}$ C and  $65 \pm 15\%$  R.H. and given a weekly opportunity to feed on chickens. Acetone solutions of precocenes I and II (7-methoxy- and 6,7-dimethoxy-2,2-dimethylchromene, respectively) were applied topically, using a microsyringe, to the dorsal abdomen of each of the five nymphal instars, so that each insect received 100  $\mu$ g of the compound. Insects in the control groups were treated with acetone (1  $\mu$ l) only. The insects were then maintained in clean glass jars (700 ml capacity) fitted with filter paper roosts and closed with nylon netting. They were given a weekly opportunity to feed on chickens and their development was observed daily.

Similarly, third and fourth instars, one to seven days after ecdysis, were treated with precocene I, precocene II and precocene II plus N-(2, 5dichlorophenyl)-3, 7-dimethyl-2, 6-octadienylamine a JHA effective in inducing juvenilization in *P. megistus* at  $1 \cdot 0 \mu g/fifth$  instar (PINCHIN *et al.*, 1978b) and  $10 \mu g/fourth$  instar (Oliveira F., unpublished results), at the dosage given in Table II, and observed as before. The mortality, time taken to ecdysis and morphological characteristics of the subsequent instars were recorded.

#### **Results and Discussion**

Table I shows the results of topical application of precocene I and II to all the nymphal instars of P. megistus. Treatment of the third and fourth instars with precocene II resulted in high percentages of precocious adultoids at the next ecdysis. Precocene I, however, showed only weak activity on the fourth instar. The percentages, corrected (Abbott's formula) with respect to control group mortality, of apparently normal insects emerging upon the first moult after treatment, demonstrated the toxic effect of precocene II on the first and second instars and, to a lesser degree, of precocene I on the first instar. Furthermore, of all the insects which underwent ecdysis after treatment with precocene II, only one, a third instar which moulted to give an apparently normal fourth instar, moulted a second time. The resulting fifth instar died the day after ecdysis. Thus precocene II was 100% effective in blocking normal development when applied to the first to fourth instars. For precocene-treated P. megistus, a marked delay in moulting was observed in those insects which underwent precocious metamorphosis.

The precocious adultoids obtained had the appearance of adults in miniature, with the characteristic red and black coloration on the conexivium. The wings, though membranous and veined, were only partially developed, being small and curled up. The tarsi were, in general, two-segmented although some of the adultoids resulting from treated fourth instars had three-segmented tarsi. It had been

Puinomo	Treated	No. insects		After 1st moult		After 2nd moult
Componing	1910111	וו כמוכת	No. precocious adultoids	% precocious adultoids % apparently normal /insects moulted /insects treated	% apparently normal /insects treated	% apparently normal /insects treated
	Ist	06	0	0.0	22.2 (39.3) <sup>1</sup>	20.0 (43.6)1
	2nd	65	0	0.0	70-8 (92-7)	60.0 (94.6)
Precocene I	3rd	50	0	0.0	82.0 (92.9)	66.0 (82.5)
	4th	50	4	9.3	$78 \cdot 0(100 \cdot 0)$	68.0/100.0)
	5th	50	0	0.0	70 • 0(100 • 0)	
	lst	30	0	0.0	10.0 (17.7)	
	2nd	25	0	0.0	8.0 (10.5)	
Precocene II	3rd	20	10	83.3	10-0 (11-3)	5.0 (6.2)
	4th	15	11	84.6	$13 \cdot 3 (18 \cdot 9)$	
	5th	10	0	0.0	$50 \cdot 0 (83 \cdot 3)$	

ar $P$ . megistus (1 to 7 days old) with precocene I, II or II plus the JHA $N$ -(2, 5-dichloro-	
days	
to 7	
ıs (1	
egistı	
т. Г.	
ar ]	
insi	
and 4th instar amine	
and	
iyl	
of	I
fment o 6-octadi	,
treatmer yl-2, 6-oc	ļ
of tr hyl-	1
yl)-3, 7-dimethyl-2, 6-octadier	
Effe 7-di	
-   3,	
able l tenyl	
Table II—Effects of phenyl)-3, 7-dimethy	

Treated	ed r Compound	Dose No.	No. insects treated		After 1st moult		A	After 2nd moult	lt
01011		heb/art tur		No. precocious adultoids	% precocious adultoids/insects moulted	% apparently normal/insects treated	No. precocious adultoids in	% precocious adultoids/ n insects moulted	% precocious % apparently adultoids/ normal/insects isects moulted treated
	Precocene I	100	30	0	0.0	$73 \cdot 3 (84 \cdot 4)^1$	0	0.0	70.0/86.6)1
	Precocene II	100	30	19	100.0	(0.0) 0.0	' {	, I	
3rd	Precocene II	10	30	6	47.4	33 · 3 (38 · 2)	2	22.2	13.3(78.8)
	Precocene II		31	0	0.0	71.0 (81.4)	0		58.1(71.0)
	Prec. II + JHA	100 + 10	30	0	0.0	46.7 (53.5)	12	100.0	
	Precocene I	100	20	0	0.0	$90 \cdot 0(100 \cdot 0)$	0	0.0	55.0(64.0)
	Precocene II	100	20	19	100.0	$(0 \cdot 0)$ (0 · 0)	1	, I	
4th	Precocene II	10	20	0	0.0	85.0 (94.0)	0	0.0	75.0(88.5)
	Prec. II + JHA	100 + 10	20	0	0.0	70.0 (77.8)	9	$42.8^{\circ}$	40.0(47.2)
<sup>1</sup> Nu <sup>2</sup>	<sup>1</sup> Numbers in parentheses correspond to the percentages corrected (using Abbott's formula) with respect to control group mortality. <sup>2</sup> Adultoids not precocious because they moulted from the 5th instar.	es correspond us because th	l to the perce sey moulted f	ntages corrected ( from the 5th insta	using Abbott's for tr.	mula) with respec	t to control group	mortality.	

546

noted previously that if precocious metamorphosis occurs from the fourth instar, either with precocene application to third-instar *O. fasciatus* (BOWERS *et al.*, 1976) or to fourth instar *Rhodnius prolixus* (Stal), as described by TARRANT & CUPP (1978), the tarsi in the resultant fifth instar adultoid are threesegmented, as in a normal adult.

The adultoids were unable to copulate with each other or with normal adults and oviposition did not occur. They were also apparently unable to feed. Life expectancy was short, averaging  $20.9 \pm 4.2$  days for fourth-instar adultoids that moulted from treated third instars; adult female *P. megistus* can live under laboratory conditions for five months and males for even longer (SZUMLEWICZ, 1976). Consequently these fourth and fifth instar adultoids can be considered to have very low vector potential.

Table II shows the results of the treatment of third and fourth instars with the precocenes at various dosages, and with precocene II plus a potent JHA for this insect. The slight difference in response to the precocenes for the insects aged up to one week (Table II) and those of undefined age (Table I) can be attributed to the fact that some individuals of the latter group had several opportunities to feed before treatment, and hence the physiological processes involved in development and ecdysis would already have been initiated in some insects.

Precocene II, applied to the third instar at  $10 \mu g/\text{insect}$ , caused the emergence of precocious adultoids upon both the first and second moults after treatment.

As expected, concurrent treatment with a JHA counteracted the effects caused by precocene. The third and fourth instars, treated with precocene II and a JHA (100 and 10  $\mu$ g/insect, respectively), after passing through an intercalary instar, gave rise to fifth and sixth instar adultoids respectively All the fifth and 50% of the sixth instar adultoids were similar in appearance to the precocious adultoids described above, although the sixth instar adultoids cannot strictly be termed precocious. However, the other sixth instars were characteristic of the partially juvenilized adultoids (PINCHIN et al., 1978b) which result from the application of JHA alone. Thus precocene apparently does not block the JH-receptor centres and probably inhibits JH production. This agrees with the results obtained for O. fasciatus (BOWERS, 1977), in which females sterilized by precocene, upon receiving a dose of JH III, resumed oocyte development.

Evidently compounds with anti-juvenile hormone activity may have a future as control agents. Contrary to JHAs, they have the advantages of shortening the life-cycle of triatomine bugs and inducing the appearance of sterile adultoids with negligible vector potential. At present it seems unlikely, however, that either JHAs or precocene analogues will completely replace insecticides in control campaigns, although they could well be useful as complementary pesticides in integrated control programmes.

#### Acknowledgements

We are indebted to the Ministry of Planning (FINEP), the National Research Council of Brazil (CNPq), and the Research Council of this University (CEPG/UFRJ) for financial assistance.

#### References

- Bowers, W. S. (1976). Discovery of insect antiallatotropins. In: The Juvenile Hormones. Gilbert, L. I. (Editor). New York: Plenum Press, pp. 394-408.
  Bowers, W. S., Ohta, T., Cleere, J. S. & Marsella,
- Bowers, W. S., Ohta, T., Cleere, J. S. & Marsella, P. A. (1976). Discovery of insect anti-juvenile hormones in plants. *Science*, **193**, 542-547.
- Bowers, W. S. (1977). Anti-juvenile hormones from plants: chemistry and biological activity. In: *Produits Naturels et la Protection des Plantes*. Citta del Vaticano: Pontifica Academia Scientiarum, pp. 129-156.
- Gilbert, B. (1976). Possible use of insect hormone mimics in vector control. In: New approaches in American trypanosomiasis research. Proc. Int. Symp., Belo Horizonte, 1975. Pan American Health Organization, Washington, D.C., pp. 252-288.
- Pinchin, R., Oliveira Filho, A. M. de, Figueiredo, M. J., Muller, C. A. & Gilbert, B. (1978a). Slow-release juvenile hormone formulations for triatomine control. *Transactions of the Royal Society of Tropical Medicine and Hygiene*, 72, 322-323.
- Pinchin, R., Oliveira Filho, A. M. de, Figueiredo, M. J., Muller, C. A., Gilbert, B., Szumlewicz, A. P. & Benson, W. W. (1978b). Screening and structure-activity relationships for juvenile hormone analogues for *Panstrongylus megistus*, a primary vector of Chagas' disease in Brazil. *Journal of Economic Entomology*, **71**, 950-955.
- Szumlewicz, A. P. (1976). Laboratory colonies of Triatominae, biology and population dynamics. In: New approaches in American trypanosomiasis research. Proc. Int. Symp., Belo Horizonte, 1975. Pan American Health Organization, Washington, D.C., pp. 63-82.
- Tarrant, Ĉ. A. & Cupp, E. W. (1978). Morphogenetic effects of precocene II on the immature stages of *Rhodnius prolixus*. Transactions of the Royal Society of Tropical Medicine and Hygiene, 72, 666-668.

Accepted for publication 31st August, 1979.