

Drug-resistant trypanosome isolate populations in dogs in Enugu North Senatorial Zone, Southeastern Nigeria

^{1*}Obi, C.F., ¹Ezeh, I.O., ¹Okpala, M.I., ²Onyeabor, A., ¹Ezeokonkwo, R. C.



UNIVERSITY OF NIGERIA
To restore the dignity of man

¹University of Nigeria, Nsukka, Nigeria. ²Michael Okpara University of Agriculture, Umudike, Nigeria *Corresponding Author: Chukwunonso.obi@unn.edu.ng; +2348063707096.

Background

African animal trypanosomosis is an important endemic protozoan disease, causing morbidities and mortalities in animals in sub-Saharan Africa. Chemotherapy is the widely used method of African animal trypanosomosis control in dogs. Trypanocidal resistance is a serious, under-reported and rapidly emerging problem in Nigeria. A paucity of information exists on the prevalence of drug-resistant trypanosomes in Nigeria.

Objectives

To determine the sensitivity of *Trypanosoma sp.* isolated from dogs in Enugu North Senatorial Zone (ENSZ), Southeastern Nigeria to trypanocides {diminazene aceturate (DA) and isometamidium chloride (ISM)}.

Methodology Multidrug-resistant Trypanosoma brucei (n = 44) & T. Single dose tests in mice trypanosomes congolense (n = 4) from dogs (Obi et al., 2021) **Results Interpretation** based on WAAVP guidelines (Eisler et al., 2001) Multidose tests in mice Characterization of Cloning of Multidrug-resistant Multidrug-resistant Single & multidose tests trypanosomes via CD₅₀ trypanosomes of clones in mice

Results

Out of the 44 *Trypanosoma brucei* isolates screened, 20 isolates (45.5%) were resistant to trypanocides while all the *T. congolense* isolates were resistant to trypanocides (Table 1). Twelve (50%), four (16.7%) and eight (33.3%) of the drug-resistant trypanosome isolates were resistant to DA, ISM and both trypanocides (multidrug-resistant) respectively.

The clones of the randomly selected multidrug-resistant trypanosome isolate (VTH 22) expressed high levels of resistance to both diminazene aceturate and isometamidium chloride (Table 2) analogous to the original isolate. The CD_{50} of VTH 22 Clone 1 and 2 were about 1.3 times higher (P<0.05) than that of the original isolate.

The CD_{50} of the drug-resistant trypanosomes ranged between 11 and 34.69 mg/kg. Drug-resistant trypanosomes were characterized into highly resistant [$(CD_{50} = 11-24.99 \text{ mg/kg})$, (Table 3)] and very highly resistant ($CD_{50} = 25 \text{ mg/kg}$) trypanosome isolates (Table 4).

Table 1: Sensitivity of *Trypanosoma* Isolates Obtained from Naturally Infected Dogs in ENSZ to trypanocides

	T. brucei	T. congolense	Overall	
	Number (%)	Number (%)	Number (%)	95% C.I
Sensitive isolates	24 (54.5)	0 (0)	24 (50)	0.352 - 0.648
Resistant isolates	20 (44.5)	4 (100)	24 (50)	0.352 - 0.648
DAR isolates	9 (45)	0 (0)	12 (50)	0.314 - 0.686
ISMR isolates	3 (15)	1 (25)	4 (16.7)	0.061 - 0.365
MDR isolates	8 (40)	3 (75)	8 (33.3)	0.178 - 0.534

DAR – Diminazene Aceturate Resistant; ISMR – Isometamidium Resistant; MDR – Multidrug Resistant

Table 2: The diminazene aceturate resistance profile of *VTH 22* and some of its clones following multi-dose test

No of mice relapsed					CD ₅₀ (mg/kg)		
Isolates	1mg/kg	3mg/kg	10mg/kg	20mg/kg	40mg/kg	CD ₅₀	95% CI
VTH 22	6	6	5	4	3	27.92	19.87 – 69.13
VTH 22 Clone 1	6	6	4	5	3	35.19	22.10 - 161.94
VTH 22 Clone 2	6	6	6	4	4	37.19	22.34 - 704.22

Table 3: Some Highly Resistant *Trypanosoma* isolates in ENSZ

No of mice relapsed					CD ₅₀ (mg/kg)		
Isolates	1mg/kg	3mg/kg	10mg/kg	20mg/kg	40mg/kg	CD ₅₀	95% CI
Orba 10	6	6	5	3	1	20.52	14.7 - 29
Orba 19	6	6	5	2	1	19.06	13.25 – 26.46
Iba 8	6	6	4	3	2	21.72	14.8 - 45
Iba 4	6	6	4	3	2	21.72	14.8 - 36.78
VTH 5	6	5	4	3	2	21.04	12.89 - 45.30
Orba 4	6	6	5	3	2	22.85	16.36 – 36.02

Table 4: Very Highly Resistant *Trypanosoma* isolates in ENSZ

No of mice relapsed					CD ₅₀ (mg/kg)		
Isolates	1mg/kg	3mg/kg	10mg/kg	20mg/kg	40mg/kg	CD ₅₀	95% CI
Orba 9	6	6	6	5	2	34.19	25.04 – 54.22
VTH 11	6	5	3	4	3	34.69	26.31 – 60.1
VTH 22	6	6	5	4	3	27.92	19.87 - 69.13
lba 1	6	6	6	4	2	25.5	19.12 - 39.06

Conclusion

Drug-resistant trypanosomes are prevalent in dogs in ENSZ with CD_{50} ranging between 11 and 34.69 mg/kg. Multidrug-resistant trypanosomes were detected with clones recording high levels of resistance to both DA and ISM.







