# ACTIVE IMMUNIZATION OF MICE WITH THE AVIRULENT Y STRAIN OF TRYPANOSOMA CRUZI AGAINST HETEROLOGOUS VIRULENT STRAINS OF THE SAME PARASITE

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### SUMMARY

Mice vaccinated with the avirulent Y strain of *Trypanosoma cruzi* became protected against infection with other virulent strains of the same parasite. This protection could be detected until 11 weeks after vaccination.

# INTRODUCTION

The cross immunization of laboratory animals with different strains of *Trypanosoma* cruzi is a well known fact <sup>1, 4, 5, 7, 8</sup> with few exceptions <sup>2</sup>. The real aim of this work is to verify whether the Y strain maintained in culture for several years and shown to be avirulent to mice <sup>6</sup> is able to protect these animals against challenge with other virulent strains of *T. cruzi*, besides the virulent Y strain.

# MATERIAL AND METHODS

Forty white mice with a mean body weight of 10 g were vaccinated with about 3  $\times$  10  $^6$  trypanosomes by subcutaneous route.

The flagellates were obtained from a culture 35 days old, in Packchanian medium, of the Y strain maintained in this medium for 16 successive years, in the Department of Parasitology of our Medical Faculty.

The trypanosomes were washed and suspended in saline solution and the suspension used immediately after preparation.

Forty other mice of the same provenience and with the same body weight were kept as controls. Ten mice, 4 weeks after vaccination, and 10 controls were inoculated by intraperitoneal route with 5,000 parasites/g body weight of the strain RC1729 isolated from a wild dog (Cerdocyon thous azarae) and maintained in mice.

The second similar group was infected with the same amount of parasites belonging to the strain M 1418 isolated from bat (Desmodus rotundus rotundus).

The third group with 10 mice controls and 10 vaccinated was inoculated, 11 weeks after the vaccination, with parasites of the strain "Berenice" isolated from a human chronic case of Chagas' disease.

Finally a fourth group, 11 weeks after the immunized animals had received the vaccine, was challenged with trypanosomes of the "ABC" strain isolated from a chronic human chagasic patient.

The number of inoculated parasites and the route of inoculation was the same for all the animals of the four groups and the infective parasites came from blood of mice at the 8th day of infection.

Parasitemia by thick drop was verified on the 8th, 15th and 30th day after challenge,

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except for the group of the RC 1729 strain of which the animals were killed after 3 weeks.

RESULTS

The results of all these experiments are summarized in Tables I to IV and in Graphs I to IV.

TABLE I

RC 1729 (wild dog Cerdocyon thous)

	Mice	Number of parasites/5 mm³ of blood				
		Days after infection				
		8	15	21		
CONTROLS	1 2 3 4 5 6 7 8 9	17.500 14.000 35.000 7.875 28.000 	4.375			
	Mean	20.903	3.552	525		
	Median	17.500	3.552	525		
VACCINATED	1 2 3 4 5 6 7 8 9	0 105 70 105 35 175 70 175 0	0 0 0 0 70 35 35 0	0 0 0 0 0 0 0 0		
	Mean	73	14	0		
	Median	70	0	0		

TABLE II

M 1418 (bat, Desmodus rotundus rotundus)

,, , , , , , , , , , , , , , , , , , , ,	Mice	Number of parasites/5 mm³ of blood  Days after infection				
CONTROLS						
		8	15	30		
	1 2 3 4 5 6 7 8 9	10.500 10.500 3.920 10.500 4.495 17.500 9.450 35.000 14.000 5.285	4.025			
	Mean	12.115	4.025	0		
	Median	10.500	4.025	0		
VACCINATED	1 2 3 4 5 6 7 8 9	175 280 140 70 140 70 140 175 35 70	0 0 0 0 0 0 0 0	0 0 0 0 0 0 0 0		
	Mean	129	0	0		
	Median	140	0	0		

From the observation of all of them it can be seen that the avirulent Y strain of *Trypanosoma cruzi* gave a very good protection against challenge with virulent strains of the same trypanosome.

The low mortality with the "ABC" strain has been already mentioned by Tafuri & Brener 9 as a peculiarity of this strain.

One vaccinated animal of this same group "ABC" was killed, by accidente, during the determination of the parasitemia on the 30th day.

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TABLE III

"ABC" (human)

TABLE IV

"BERENICE" (human)

	Mice	Number of parasites/5 mm³ of blood			Mice	Number of parasites/5 mm³ of blood  Days after infection			
		Days after infection							
		8	15	30		·	8	15	30
CONTROLS	1 2 3 4 5 6 7 8 9 10	10.500 17.500 5.075 5.775 2.730 7.280 2.870 4.025 10.500 5.355	4.025 10.500 2.870 2.835 4.375 3.570 3.780 7.085	280 ————————————————————————————————————	CONTROLS	1 2 3 4 5 6 7 8 9	2.520 4.515 2.555 	770   1.155 2.170 1.645 735	35 
	Mean Median	7.161 	4.880 3.902	365		Mean Median	2.555	1.295	17,5
VACCINATED	1 2 3 4 5 6 7 8 9	0 0 0 70 0 140 175 0 105 70	70 0 0 35 0 0 0 35 35 35	0 0 0 0 70 	VACCINATED	1 2 3 4 5 6 7 8 9	70 70 280 280 140 140 105 175 105 70	0 70 0 35 70 0 0 0	0 35 0 35 0 0 0 0
	Mean	56	19	1,1		Mean	143	17,5	7,0
	Median	35	0	0		Median	122	0	0

killed by accident

# COMMENTS

GRAY<sup>3</sup> in his work on immunology of trypanosomiasis says that recovery from an infection with a high infective strain gives no immunity or a very low one against a latter infection with other strains.

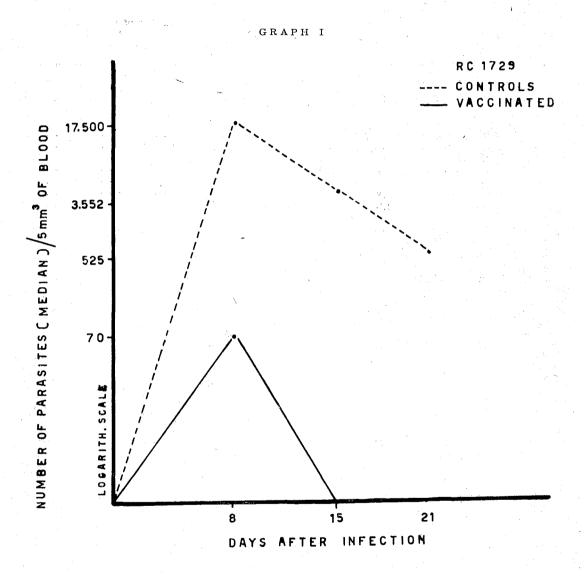
This statement (although not explicit) seems to apply only to the African trypanosomiasis because in the South American try-

panosomiasis this is perhaps the exception and not the rule.

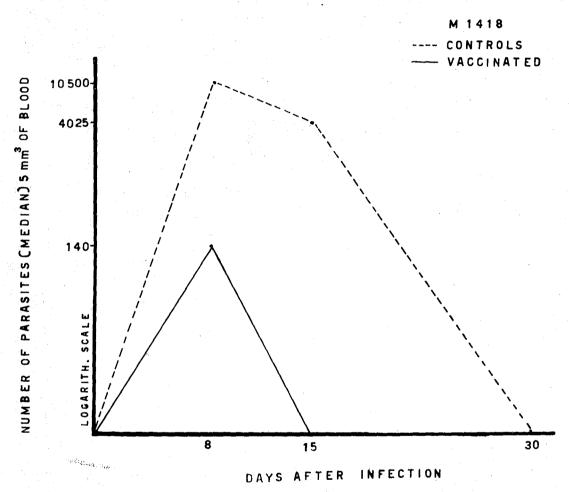
This is a crucial point for the success in any attempt to obtain a vaccine against the disease.

The specificity of the vaccine being of strain and not of species it would be necessary to have a great mixture of antigens and would make the vaccination very difficult or even impossible.

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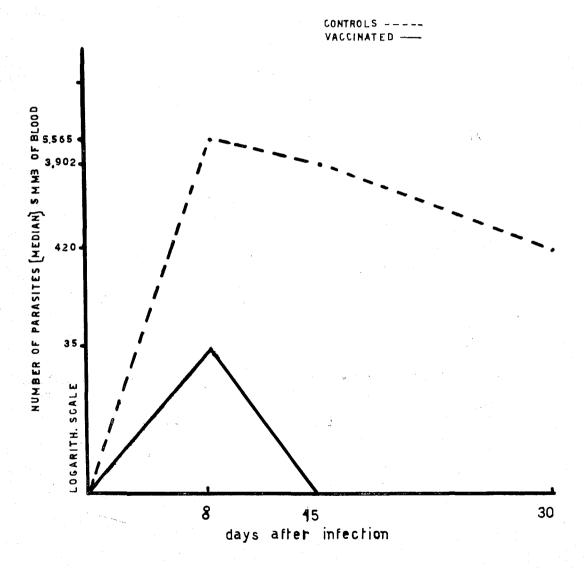


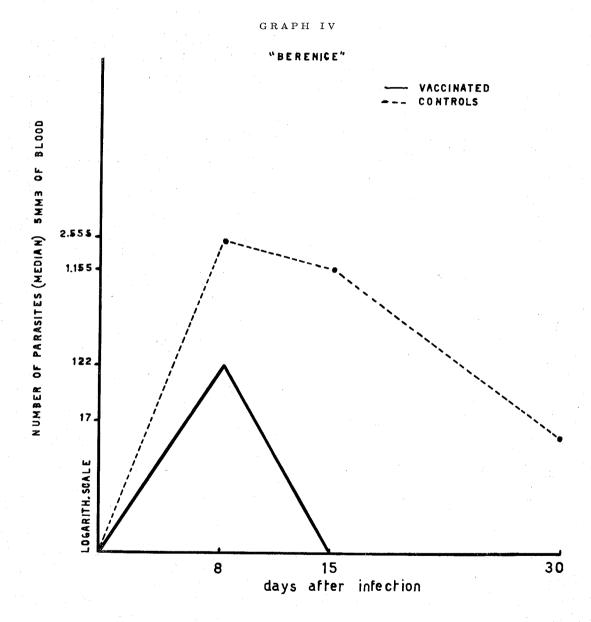




GRAPH III

# "ABC"





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Our experiments show that the avirulent Y strain keeps the ability to give cross protection against different virulent strains of T. cruzi.

## RESUMO

Imunidade cruzada de camundongos vacinados com cepa avirulenta de T. cruzi contra outras cepas, virulentas, do mesmo parasita

A cepa Y, avirulenta, confere a camundongos, imunidade cruzada contra infecção por várias cepas virulentas do *Trypanosoma cruzi*.

Essa imunidade foi determinada até 11 semanas após a vacinação.

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