## GEORGIA INSTITUTE OF TECHNOLOGY OFFICE OF CONTRACT ADMINISTRATION SPONSORED PROJECT INITIATION

	Date:June_18, 1979				
Project Title: Developmental - Genetic Reg	ulation of Brain Tryptophan Transport				
which we are a comparable to the contract of t					
Project No: G-32-658 GRaew	carel				
Project Director: Dr. James A. Diez					
Sponsor: National Science Foundation					
Agreement Period: From 6/15/79	Until 11/30/82 (Grant Period)				
Type Agreement: Grant No. BNS-7905601,	dated 6/8/79				
Amount: \$85,000 NSF	÷0.				
. 4,474 GIT (G-32-327) \$89,474 TOTAL					
\$69,474 TOTAL					
Reports Required: Annual Progress Reports;	Final Project Report				
A STATE OF THE STA					
Sponsor Contact Person (s):					
Sponsor Contact Leison (s).					
Technical Matters	Contractual Matters				
	(thru OCA)				
NSF Program Official					
Nathaniel G. Pitts Assistant Program Director, Neurobiology	NSF Grants Official qu Program Mr. Thomas F. Griffin				
Neurosciences	DBS/SE Branch				
Div. of Behavioral and Neural Sciences	Division of Grants and Contract.				
Directorate for Diological, Behavioral					
Social Sciences	National Science Foundation				
National Science Foundation	Washington, D. C. 20550				
Washington, D. C. 20550	202/632-7496				
202/634-4036					
Defense Priority Rating: n/a	¥ ·				
Assigned to: Biology	(School/Laboratory)				
COPIES TO:					
COFFES TO:					
Project Director	Library, Technical Reports Section				
Division Chief (EES)	EES Information Office				
School/Laboratory Director	EES Reports & Procedures				
Dean/Director—EES Accounting Office	Project File (OCA)				
Procurement Office	Project Code (GTRI) Other				

Security Coordinator (OCA)
Reports Coordinator (OCA)

## SPONSORED PROJECT TERMINATION SHEET

4		Dat	e <u>July 12, 1983</u>	
Project Title:	Developmental - G	enetic Regulation	of Brain Tryptophan	Transport
Project No:	G-32-658		and the second s	
Project Direct	tor: Dr. James A. Di	ez	96.446 NA N 2 QUIN	
Sponsor: N	ational Science Found	dation		
Effective Teri	mination Date: 11/3	0/82		
Clearance of	Accounting Charges:	11/30/82		
Grant/Contrac	t Closeout Actions Rema	aining:		9
e take a je			T.	<b>↓</b> 4.4
			***	2
		osing Documents	** 1	
_	x Final Fiscal Report 5	FCTR		
1	Final Report of Inver	ntions		
(1	Govt. Property Inven	tory & Related Certif	icate	
	Classified Material Ce	rtificate		
	Other			
		<del> </del>		
A.C.				
	- T			
Assigned to:	Applied Biology		(School/Laborator	w)
				<del>777</del>
COPIES TO:	Marchine of the reducement of the second	وريا المتوجه المتحافظ	and the second s	
Accounting	perty Management SL	esearch Security Services (OCA) ibrary		put

## NATIONAL SCIENCE FOUNDATION FINAL PROJECT REPORT Washington, D.C. 20550 PLEASE READ INSTRUCTIONS ON REVERSE BEFORE COMPLETING PART I-PROJECT IDENTIFICATION INFORMATION 2. NSF Program 3. NSF Award Number 1. Institution and Address Georgia Institute of Technology BNS-7905601 Neurobiology 5. Cumulative Award Amount 4. Award Period Atlanta, Georgia 30332

From 6/15/79 To 11/30/82

\$91,312

6. Project Title

Developmental-genetic regulation of brain tryptophan transport.

## PART II-SUMMARY OF COMPLETED PROJECT (FOR PUBLIC USE)

The major aim of this project was to determine whether the system which transports tryptophan (TRP) across the neuronal cell membrane shows significant physiological variation due to genotype of developmental age. TRP is the precursor of the neurotransmitter serotonin; variations in TRP availability can alter serotonin synthesis, and thereby affect the many behaviors modulated by serotonin.

The membrane transport system for TRP was studied in synaptosomes (nerve endings) prepared from whole mouse brain; the accumulation of radioactive TRP was used to characterize the maximum transport rate (Vmax) and the affinity of the carrier for TRP (Km). The transport constants were measured in preparations from several strains of mice which show differing behavioral traits; developmental changes in the constants were studied from birth to sexual maturity (approx. 8 wks.).

The majot hypotheses of this study were confirmed: significant genetic differences in the Vmax and developmental changes in both the Km and Vmax for TRP transport were identified. Attempts to find a hormonal basis for the differences were not successful.

Experiments on the mechanism by which TRP is accumulated in synaptosomes have helped to resolve a controversy about how many carrier systems move TRP across the membrane. Our results indicate the existence of one system with relatively high affinity; the "low affinity" system which also appears in most TRP uptake studies seems to result from intra-cellular binding rather than movement across the membrane. These experiments also led to the discovery of a TRP-binding "phenomenon" in brain cell membrane fragments. This binding appears to behave much like a receptor for TRP, except that its dissociation constant (Kd) is relatively high (1  $\mu$ m). The binding we measure could be to the carrier which transports TRP, except that it does seem to be unique to brain (we have not been able to measure it in liver, kidney, heart, erythrocytes, or platelets).

PART III-TECHNICAL INFO	RMATION (FOR P.	ROGRAM MAN	AGEMENT USES		FIIDNICHED
1. ITEM (Check appropriate blocks)	NONE	ATTACHED	PREVIOUSLY FURNISHED	TO BE FURNISHED SEPARATELY TO PROGRAM	
	None			Check (√)	Approx. Date
a. Abstracts of Theses					7/15/83
b. Publication Citations			<u> </u>	V	7/15/83
c. Data on Scientific Collaborators	V			<u> </u>	
d. Information on Inventions	V				- 1-
e. Technical Description of Project and Results					7/15/83
f. Other (specify)					
2. Principal Investigator/Project Director Name (Typed)  James A. Diez	3. Principal Inve	estigator/Project	Director Signatur	e	4. Date 5/18/83
SF Form 98A (5-78) Supersades All Previous Editions		<u> </u>	<del></del>	Form A	pproved OMB No. 99R0

NSF Form 98A (5-78) Supersedes All Previous Editions