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ponsor:DHEW/PHS/NIH - National	l Institute of Al	lergy and Infe	ectious Disease	es
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ype Agreement: Grant No. 5 K04	AI00332-03 (year	03)		
ward Period: From 7/1/81 To	6/30/82 (1	Performance)	9/30/82	(Reports)
Sponsor Amount: \$38,718			Contracte	d through:
Cost Sharing: \$ 2,863			GTRI/	COOX
Title: Interaction of RNA Polymera	se with DNA Sites	3		-
ADMINISTRATIVE DATA	OCA CONT	Willia:	m F. Brown x4	320
L) Sponsor Technical Contact: Pro Molecular Microbiology and				
Officer; Grants Management Reports: See Deliverable Schedule			20014 (301) None	496-7075
Defense Priority Rating: N/A				
RESTRICTIONS				
See Attached NIH	Supplemental Inf	formation Sheet	t for Additiona	al Requireme
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OFFICE OF CONTRACT ADMINISTRATION

SPONSORED PROJECT TERMINATION/CLOSEOUT SHEET

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Project No. G-41-D03	School/Kask	Physics
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Project Director(s) Dr. R. M. Wartell		发环和 / GIT
Sponsor DHEW/PHS/NIH - National Institute of Allergy	v and Infectious Di	sease
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Title Interaction of RNA Polymerase with DNA Sites	· · · · · · · · · · · · · · · · · · ·	
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	GHANT NUMBER	-41-DO3/Wartell	
PROGRESS REPORT	CHANT NUMBER G-41-DO3/Wartell 5-K04-AI-00332-04		
NAME OF HEAD OF DEPARTMENT OR DEPARTMENTAL SUBDIVISION	ACCOMPLISHMENTS COVERING PERIOD		
"Professor Edward W. Thomas	FROM	THROUGH	
Professor Roger M. Wartell	7/1/81	6/30/82	
Georgia Institute of Technology			

G-41-D03

Interaction of RNA Polymerase with DNA Sites

STATEMENT OF ACCOMPLISHMENT (IF SPACE IS INADEQUATE, USE CONTINUATION PAGE)

RESEARCH

During the past year progress was made in developing methods of examining DNA conformation in solution. New results on junctions between different DNA conformations were obtained. Raman spectroscopy was employed to determine the vibrational spectra of short DNA molecules. Also, the temperature induced duplex to single strands transition of DNA promoter fragments was examined both experimentally and theoretically.

One study examined the Raman spectra of the DNA polymer $(dG-dC)_n \cdot (dG-dC)_n$, and two DNA restriction fragments, 95 base pairs (bp.) long and 157 bp. long (1). The 95 bp. DNA contains the RNA polymerase binding site of the E. coli lactose operon transcription start region. The 157 bp. DNA has 26 and 32 bps. of (dG-dC) sequences at each end of the 95 bp. DNA. The Raman spectra show that in 10mM NaCl, the two (dG-dC) regions of the 157 bp. DNA form a left-handed "Z" conformation which alters vibrations of the middle 95 bp. section. By itself the 95 bp. DNA has a B conformation in 4.5 M NaCl whereas $(dG-dC)_n$. (dG-dC) is in a Z conformation. The results show that left handed helical segments of DNA can form junctions with, and distort DNA normally in a right handed helix. Raman data has been obtained which follows the intensities of 20 different vibrational bands of the 157 bp. DNA during the NaCl induced transition of the (dC-dG) regions from right to left handed helices. Computer methods were developed and are being applied to quantifying peak intensities and widths of overlapping Raman bands. These methods were successfully applied to examine the B to A transition of a heterogenous sample of calf thymus DNA (2).

Theoretical analysis was carried out on the duplex of coils transition of three DNA fragments containing different promoter sequences. The influence of 20 known base pair mutations on the DNA melting curves were predicted. (3) Some of the predictions are currently being tested.

PUBLICATIONS

- "The Junction Between 'Z' and 'B' Conformation in a DNA Restriction Fragment: Evaluation by RAMAN Spectroscopy" R. M. Wartell, J. Klysik, W. Hillen, and R. D. Wells, (1982) Proc. Natl. Acad. Sci., (USA) in press.
- 2. "Changes in Raman Vibrational Bands of Calf Thymus DNA During the B to A Transition" J. C. Martin and R. M. Wartell (1982) <u>Biopolymers 21</u> 499.
- 3. "Fluctuational Base Pair Opening in DNA at Temperatures Below the Transition Region" by R. M. Wartell and A. S. Benight (1982) <u>Biopolymers</u>, in press.
- 4. "Influence of Base Pair Changes and Cooperativity Parameters on the Melting Curves of Short DNAs", A. S. Benight and R. M. Wartell (in preparation).

Contractor (Contractor)

Professor Wartell taught two courses in the School of Biology this winter quarter. One course was Biophysical Genetics, Biol. 4470. The second course, Biol 3308, was Genetic Engineering and was introduced into the curriculum for the first time by Professor Wartell and Professor Hall. Professor Wartell has also taught Biophysics II, Phys. 4252 and the undergraduate biophysics laboratory course Phys. 4254 in the spring quarter. Both of these courses are essential components of the option in biophysics. Two graduate students are working on their doctoral thesis research in Professor Wartell's laboratory.

Professor Wartell presented invited lectures at the University of Alabama -Birmingham (June 1981) and at Brookhaven National Laboratories (February 1982).

The biophysics courses and research have become a major feature of our programs under his strong leadership. They attract students at all levels and we see an increasing proportion of entering graduate students declaring this to be their area of proposed work. In recognition of these developments Professor Wartell serves on the committee to recruit and select graduate students. He is relied on as a source of well balanced judgements in all aspects of the school's activities. The faculty has elected him to serve on the school's advisory committee and the Director has appointed him to chair the committee to search for new faculty. In recognition of the contribution made by Prof. Wartell's research and instructional programs, we are searching for a further faculty person in the area of Biophysics.

In all respects Professor Wartell's contributions and career development are at a superior level.