### A TOY MODEL THE EVOLUTION OF DNA POLYMERASE POLARITY

by

### Joshua Ballanco

### A DISSERTATION

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# DOCTOR OF PHILOSOPHY

Joshua Ballanco, Candidate	
ADVISORY COMMITTEE	
Marc Mansfield, Chairman	Date
A. V. Caramula	Date
A. K. Ganguly	Date
Knut Stamnes	Date
Nicholas Murgolo	Date

STEVENS INSTITUTE OF TECHNOLOGY Castle Point on Hudson Hoboken, NJ 07030 2009



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ABSTRACT

It is my thesis that life can be modeled as a thermodynamic system. Specifically, it

is my goal to use this approach to understanding life in order to investigate evolution

as an energy minimization problem. In this sense, ecological niches represent local

energy minima, selection criteria are nothing more than potential energy gradients,

and extinction is the rate of change of the energy landscape exceeding the kinetics

of adaptation. This idea is derived from the observation that life is, at its most

fundamental level, nothing more than an elaborate collection of chemical reactions.

Taken individually, each of these reactions is governed by the laws of thermodynamics,

and thus far there is no known upper limit to the scale to which thermodynamic

principles may be applied.

In this work I present a model systems which is inspired by this hypothesis.

In this model, I will look at the directionality of nucleotide polymerases, all of which

synthesize new nucleotide polymers in a 5' to 3' direction. This phenomenon could

be the consequence of a very early founder effect. On the other hand, it could be

that this directionality evolved due to an inherent advantage. I would propose that

the process of nucleic acid polymerization provides a clue as to what that advantage

might have been. The toy model I have constructed demonstrates this, and the results

provide an insight into the forces that might have driven early biological evolution.

Author: Joshua Ballanco

Advisor: Marc Mansfield

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### Chapter 1

#### Introduction

Evolution has been, since the time of Darwin, a science mostly concerned with explaining observations and predicting what new evidence from the past should eventually turn up, such as the existence of transitional forms. In this regard, evolution has been wildly successful. Where the study of Evolution has been lacking is in its ability to predict the future. That is, looking at collected fossil and current natural evidence, the Theory of Evolution gives us the ability to identify which selective pressures acted on past populations. What the Theory of Evolution cannot do, at present, is predict which environmental or other influences will act as selective pressures going forward. At best, we can make educated guesses based on past evidence, but we lack even the basic ability to assign a concrete measure of confidence in such predictions.

### Chapter 2

### Design of the Polymerase Evolution Model

In this chapter, I will provide a detailed description of the design of the model system used to investigate the question of the evolution of polymerase polarity. The goal of this model is to have organisms containing either  $5' \rightarrow 3'$  or  $3' \rightarrow 5'$  compete with each other in order to see which strategies are either evolutionarily dominant or evolutionarily stable.

### Overview

### Vita

### Joshua Ballanco

Place of birth Hazel Crest, IL

Date of birth October 5, 1980

Education Stevens Institute of Technology, Hoboken, NJ

Doctoral Candidate in Chemical Biology expected date of graduation, September 2009

Stevens Institute of Technology, Hoboken, NJ

Masters of Science in Chemistry

May 2008

Stevens Institute of Technology, Hoboken, NJ Bachelors of Science in Chemistry with Honors

May 2002