SPECIFIC INTERACTIONS BETWEEN NIPAH VIRUS NUCLEOCAPSID (N) PROTEIN AND PHOSPHO-(P) PROTEIN USING THE YEAST TWO-HYBRID SYSTEM

By

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Thesis Submitted to the School of Graduate Studies, Universiti Putra Malaysia, in Fulfilment of the Requirement for the Degree of Master of Science

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Dedicated to my parents, sister and brother

Abstract of thesis presented to the Senate of Universiti Putra Malaysia in fulfilment of the requirement for the degree of Master of Science

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Chairman: Professor Datin Khatijah Mohd Yusoff, PhD

Faculty : Biotechnology and Biomolecular Sciences

Nipah virus (NiV) which is a member of a new genus, *Henipavirus*, in the family *Paramyxoviridae*, encodes an unusually large phospho- (P) protein compared to other known paramyxoviruses. In this study, the region(s) involved in the interaction between this exceptionally large P protein with its nucleocapsid (N) protein was investigated *in vivo* using the yeast two-hybrid system. Deletion analysis was used to map the domain(s) of both the N and P proteins involved in N-P and P-N interactions. Mapping of the domains of N protein involved in its interaction with the P protein revealed that the C-terminal 30 amino acids (423-452 residues) are crucial for N-P interaction. However, mapping of the domains of P protein involved in the P-N association demonstrated that both the C-terminal 63 amino acids (470-532 residues) and the immediate N-terminal 62 amino acids (1-62 residues) simultaneously play a major role. Comparison of these findings with other studies indicates that paramyxoviruses are different in terms of

interaction domains(s) between these two essential viral proteins involved in genome replication.

Abstrak tesis yang dikemukakan kepada Senat Universiti Putra Malaysia sebagai memenuhi keperluan untuk ijazah Master Sains

TINDAK BALAS SPESIFIK DI ANTARA PROTEIN NUKLEOKAPSID (N) DAN PROTEIN FOSFO (P) NIPAH VIRUS MENGGUNAKAN SISTEM DWI-HIBRID YIS

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Nipah virus (NiV) yang merupakan ahli *Henipavirus* iaitu satu genus baru dalam famili *Paramyxoviridae*, mengkodkan suatu protein fosfo (P) yang besar jika dibandingkan dengan paramyxovirus lain yang diketahui. Dalam kajian ini, bahagian yang terlibat dalam tindak balas di antara protein P ini dengan protein nukleokapsid (N) telah dikaji secara *in vivo* dengan menggunakan sistem dwi-hibrid yis. Analisis pemotongan telah digunakan untuk memetakan domain yang terdapat pada kedua-dua protein N dan P yang terlibat dalam tindak balas N-P dan P-N. Pemetaan domain protein N yang terlibat dalam tindak balas dengan protein P menunjukkan bahawa 30 asid amino pada terminal-C (residu 423-452) adalah penting untuk tindak balas N-P. Walau bagaimanapun, pemetaan domain protein P yang terlibat dalam penggabungan P-N menunjukkan bahawa kedua-dua 63 asid amino pada terminal-C (residu 470-532) dan 62 asid amino pada terminal-N (residu 1-62) memainkan peranan yang sama penting. Perbandingan antara penemuan kajian ini dengan kajian-kajian yang lepas menunjukkan bahawa

paramyxovirus adalah berbeza dari segi bahagian-bahagian protein N dan protein P yang bertindak balas di antara satu sama lain dalam proses replikasi genom.

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I certify that an Examination Committee has met on 7th February 2006 to conduct the final examination of Taznim Begam binti Mohd Mohidin on her Master of Science thesis entitled "Specific Interactions Between Nipah Virus Nucleocapsid (N) Protein and Phospho-(P) Protein Using the Yeast Two-hybrid System" in accordance with Universiti Pertanian Malaysia (Higher Degree) Act 1980 and Universiti Pertanian Malaysia (Higher Degree) Act 1980 and Universiti Pertanian Malaysia (Higher Degree) Regulations 1981. The committee recommends that the candidate be awarded the relevant degree. Members of the Examination Committee are as follows:

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DECLARATION

I hereby declare that the thesis in based on my original work except for quotation and citations, which have been duly acknowledged. I also declare that it has not been previously or concurrently submitted for any other degree at UPM or other institutions.

TAZNIM BEGAM BINTI MOHD MOHIDIN

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TABLE OF CONTENTS

Page

DEDICATION	ii
ABSTRACT	iii
ABSTRAK	V
ACKNOWLEDGEMENTS	vii
APPROVAL	ix
DECLARATION	xi
LIST OF TABLES	xiv
LIST OF FIGURES	XV
LIST OF ABBREVIATIONS	xvi

CHAPTER

1	INTRODUCTION	1
2	LITERATURE REVIEW	3
	2.1 Introduction	3
	2.2 Nipah virus	6
	2.2.1 Phylogenetic analysis	6
	2.2.2 Morphology of NiV	9
	2.2.3 Nucleoprotein of NiV	12
	2.2.4 Phosphoprotein of NiV	13
	2.3 Protein-protein interaction	15
	2.3.1 Interactions between N and P	15
2.3.2	Advantages of interaction mapping 16	
	2.4 Two-hybrid system	18
	2.4.1 Applications of the two-hybrid system	19
	2.4.2 The yeast two-hybrid system	20
	2.4.3 Advantages of the yeast two-hybrid system	22
3	MATERIALS AND METHODS	24
	3.1 Source of genes of NiV	24
	3.2 Source of chemicals and biochemicals	24
	3.3 Construction of bait and prey plasmids	24
	3.3.1 Amplification of N and P full length genes	24
	3.3.2 Propagation of bait and prey vector plasmids	26
	3.3.3 Preparation of competent <i>Escherichia coli</i> TOP10 cells	27
	3.3.4 Restriction enzyme digestion	28
	3.3.5 Directional ligation	29
	3.3.6 E. coli transformation	29