PERPUSTAKAAN HAMDAN TAHIR UNIVERSITI SAINS MALAYSIA



# UNIVERSITI SAINS MALAYSIA GERAN PENYELIDIKAN UNIVERSITI PENYELIDIKAN LAPORAN AKHIR

# MOLECULAR MECHANISM OF THE ACTIN-ASSOCIATED PROTEIN HIP1R ON MALIGNANT B-CELL MIGRATION

**PENYELIDIK** 

DR. WONG KAH KENG

PENYELIDIK BERSAMA

PROF. MUSTAFFA MUSA PROF. GAN SIEW HUA MS. LOO SUET KEE

2015



## FINAL REPORT FUNDAMENTAL RESEARCH GRANT SCHEME (FRGS)

Laporan Akhir Skim Geran Penyelidikan Fundamental (FRGS) Pindaan 1/2015

A RESEARCH TITLE: Molecular Mechanism of the Actin-Associated Protein HIP1R on Malignant B-cell Migration.

PHASE & YEAR: 2, Year 2015

START DATE: 1st June 2012 END DATE: 31st May 2014

EXTENSION PERIOD (DATE): RMC LEVEL: 30th November 2014

**KPM LEVEL: -**

PROJECT LEADER: Wong Kah Keng

I/C / PASSPORT NUMBER: 830808-14-5513

PROJECT MEMBERS: 1. Prof. Mustaffa Musa (including GRA) 2. Prof. Gan Siew Hua

3. Ms. Loo Suet Kee (student)

PROJECT ACHIEVENESH (Present Projek)

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	ACHIEVEMENTYPER	CENTAGE		
Project progress according to milestones achieved up to this period	0 - 50%	51 - 75%	76 - 100%	
Percentage (please state #%)	100%	100%	95%	
	RESEARCH OU	TPUT		
Number of articles/ manuscripts/ books	Indexed Journal		Non-Indexed Journal	
(Please attach the First Page of Publication)	1 (Impact factor: 9.379)		-	
Conference Proceeding	International -		National	
(Please attach the First Page of Publication)			-	
Intellectual Property (Please specify)				

HUMAN CAPITAL DEVELOPMENT							
Human Capital	Number				Others	_	
	On-going		Graduated		(please specify)		
Citizen	Malaysian	Non Malaysian	Malaysian	Non Malaysian	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	_	
No. PHD STUDENT	1	-		-	]		
Student Fullname: IC / Passport No: Student ID:	Loo Suet Kee 871002-35-5344 P-UD0006/13(R)	-	-	-			
No. MASTER STUDENT	-	-	-	-			
Student Fullname: IC / Passport No: Student ID:	-	-	-	-			
No. UNDERGRADUATE STUDENT	+	-	-	_			
Student Fullname: IC / Passport No: Student ID:	-	_	**	_			
Total	1 (PhD)	-	-	-			

#### EXPENDITURE (Parishington) as Borang X1(RINC)

Budget Approved (Peruntukan diluluskan) : RM 125,000 Amount Spent (Jumlah Perbelanjaan) : RM 125,000

Balance (Baki) : RM 0 Percentage of Amount Spent : 100%

(Peratusan Belanja)

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ADDITIONAL RESEARCH ACTIMITIES THAT CONTRIBUTE TOWARDS DEVELOPING SOFT AND HARD SKILLS (AKIMI) Penyalidikan Sampingan (eng menyambang kepada pembangunan kemahiran insaniah)

Activity	Date (Month, Year)	Organizer		
(e.g : Course/ Seminar/ Symposium/ Conference/ Workshop/ Site Visit)	-	-		
National	<u> </u>			
Activity	Date (Month, Year)	Organizer		
Next Generation Researchers Cancer Symposium (my role: presenter and organiser) (e.g: Course/ Seminar/ Symposium/ Conference/ Workshop/ Site Visit)	19th November 2014	Centre for Research Initiative – Clinical and Health Sciences (CRI- CHS), Health Campus, Universiti Sains Malaysia.		

### PROBLEMS/MECONSTRAINTS IF ANY (Masalan/Kelengen

The first phase of the project (activity no. 1 and no. 2, please refer to the Gantt chart) had been completed, resulting in a paper published in Leukemia (impact factor: 9.379) in early 2014 and acknowledged FRGS, please refer to the evidences submitted together with this report. In the publication, we reported that HIP1R (a putative tumour suppressor) expression was repressed by the transcription factor FOXP1 (an oncoprotein), and their inverse expression pattern conferred worse survival in a series of B-cell lymphoma cases.

The next phase of the project (activity no. 3 and no. 4) had been completed mid of last year and drafting of the manuscript is still in progress. The manuscript entitled "Prognostic and subtyping utility of the actin-binding protein HIP1R in diffuse large B-cell lymphoma" will be submitted into the journal British Journal of Haematology (impact factor 4.959), please refer to the first page of the manuscript draft. It will be submitted in late February or early March 2015 and the evidence for the submitted manuscript will be sent to CRI as soon as possible.

#### RECOMMENDATION (Crotengen Penemberhoelkein)

To submit the next manuscript entitled "Prognostic and subtyping utility of the actin-binding protein HIP1R in diffuse large B-cell lymphoma" into British Journal of Haematology as soon as possible (by end of Feb or early March 2015).

#### RESEARCH/ARSIRACT — Not Nove Them 200 Words (Abstrak Penyaltilken — Titak Meledihi 200 patah perkataan)

In this FRGS project, we aimed to uncover why HIP1R is expressed and how its expression is regulated in diffuse large B-cell lymphoma (DLBCL), an aggressive malignancy and the most common type of non-Hodgkin's lymphoma. We found that HIP1R was expressed at high levels in non-malignant mature B cells, and its expression was maintained in a subtype of DLBCL termed as GCB-DLBCL, the subtype with a better outcome compared to the more aggressive subtype, the ABC-DLBCL. In a patient cohort (n=157), HIP1R expression alone was predictive of outcome in the disease. We demonstrated that HIP1R expression was downregulated by FOXP1, a transcription factor involved as an oncoprotein in the disease, and that their inverse expression pattern conferred worse survival in the patient cohort. HIP1R was the first direct target of FOXP1 uncovered and our findings were published in the journal Leukemia (impact factor 9.379). Finally, we tested the prognostic and subtyping utility of HIP1R in a series of HUSM DLBCL patient cohort (n=73). In this independent series, HIP1R was predictive of survival and its expression was correlated with the GCB-DLBCL subtype, indicating that the molecule is a strong predictive marker in the disease as demonstrated in two independent patient cohort. (word count: 200)

Date Tarikh : 30th Jan 2015

Project Leader's Signature: Tandatangan Ketua Projek

COMPARAMENTS, IF ANYMENDORSEMENT BY RESEARCH MANAGEMENT GENTER (RMG) (Koman, sekirinya ada/ Pangasahan elah Pusat Pangurusan Panyaldikan)

Name:

PROF. DR LEE KEAT TEONG

Signature:

Nama:

Pengarah Pejabat Pengurusan & Kreativiti Penyelidikan Tandatangan:

Date: Tarikh: Universiti Salns Malaysia

11800 USM, Pulau Pinang.