

HMRF Research Fellowship Scheme – Experience Sharing

Dr. Eric WAN Assistant Professor Department of Family Medicine and Primary Care & Department of Pharmacology and Pharmacy The University of Hong Kong



Outline

- Introduction of my Research Fellowship project
- Proposal writing and interview
- Challenges during the implementation phase
- Take-home messages



"When should statin therapy be initiated for patients with diabetes mellitus"

Mentor: Professor Cindy Lam (HKU Family Medicine) Overseas supervisor: Professor Goodarz Danaei (Harvard Cardiovascular Health)





Current evidence

LDL-C level

Existing evidence or guideline

— 4.0 mmol/L	RCT in Japan : a lower risk of coronary heart disease in the treatment arm using pravastatin in the cohort (21% were diabetic) with a mean baseline LDL-C of 4 mmol/L
— 3.0 mmol/L	RCT in UK, Multi-center RCT in 21 countries, Cohort study in Spain: demonstrated the benefits of statin for primary prevention in participants with a mean baseline LDL-C of at least 3 mmol/L
— 2.6 mmol/L	Guideline in mainland China and HK: LDL-C ≥ 2.6 mmol/L in patients with DM
— 1.8 mmol/L	Guideline by ACC/AHA : suggesting the use of statin in diabetic patients who aged 40 to 75 years with LDL-C \geq 1.8 mmol/L



Objective

Comparing effectiveness and safety of initiating statin therapy at different LDL-C thresholds (1.8mmol/L vs. 2.6mmol/L) for primary prevention



Overview of Method: target trial emulation





Results: Estimated effects of statin initiation

Intention-to-treat analysis

Per-protocol analysis

7

	Hazard ra	tio (95% CI)	Hazard	ratio (95% Cl)	
Baseline LDL-C: 1.8-2.5 mmol/L					
Overall CVD	0.78 (0.72, 0.84)	⊢ •−1	0.59 (0.51, 0.68)	 -1	
Myocardial infarction	0.84 (0.74, 0.96)	·	0.61 (0.39, 0.95)	·	
Heart failure	0.87 (0.77, 0.98)	،	0.39 (0.23, 0.65)	• •	
Stroke	0.72 (0.65, 0.80)		0.41 (0.31, 0.55)	• 1	
Ischemic stroke	0.65 (0.55, 0.76)		0.41 (0.22, 0.76)	·	
Haemorrhage	0.73 (0.55, 0.98)	•	0.24 (0.05, 1.12)	· · · · · · · · · · · · · · · · · · ·	
Myopathies	1.03 (0.74, 1.42)		0.45 (0.13, 1.50)	•	
Liver dysfunction	0.86 (0.79, 0.93)		0.96 (0.71, 1.30)	·•	
Mortality	0.80 (0.74, 0.86)		0.70 (0.52, 0.94)	·	
Baseline LDL-C≥2.6 mmol/L					
Overall CVD	0.90 (0.88, 0.92)		0.77 (0.74, 0.81)	H a ri	
Myocardial infarction	0.86 (0.82, 0.89)	⊢⊷ ∣	0.68 (0.62, 0.75)	⊢← -i	
Heart failure	0.91 (0.88, 0.95)		0.69 (0.63, 0.75)		
Stroke	0.89 (0.87, 0.92)	He-I	0.78 (0.73, 0.83)		
Ischemic stroke	0.82 (0.78, 0.86)	⊢ •−1	0.65 (0.58, 0.72)		
Haemorrhage	0.92 (0.85, 1.01)	·	0.85 (0.69, 1.04)	·+	
Myopathies	1.02 (0.92, 1.13)	·•_	→ 0.87 (0.67, 1.14)		<u> </u>
Liver dysfunction	0.92 (0.89, 0.94)	-+-	0.85 (0.80, 0.90)	⊢ •-1	
Mortality	0.92 (0.90, 0.94)		0.81 (0.76, 0.86)	•••	
		0.5 1.0	1.5	0.0 0.5 1.0	1.5



Conclusion

Initiating Statin in diabetic patients with **baseline LDL-C \geq1.8mmol/L** was associated with reduced risks of incident CVD and all-cause mortality among the Chinese population without a significant increase in the risk of myopathy, liver function derangement, or cancer.

Published in Diabetes, Obesity & Metabolism

• Wan EYF et al. Evaluating different low-density lipoprotein cholesterol thresholds to initiate statin for prevention of cardiovascular diseases in patients with type 2 diabetes mellitus: A target trial emulation study. Diabetes, Obesity and Metabolism. 2024

Extended outcomes

- Xu W et al. Long-term statin use and risk of cancers: a target trial emulation study. Journal of Clinical Epidemiology. 2024
- Xu W et al. Benefits and Risks Associated With Statin Therapy for Primary Prevention in Old and Very Old Adults: Real-World Evidence From a Target Trial Emulation Study. Annals of Internal Medicine. 2024
- HMRF 2023 (Recommend for Support)



Proposal writing and interview



Proposal writing - Research project

Components	Advices
Clinical important (e.g. what is the gap, what is the impact)	Seek the advice of Health Professions (e.g. clinicians, nurses, pharmacists, OT, PT, dietitian etc.)
Feasibility (e.g. whether the study - can be done)	Include the results from pilot study Include preliminary data (especially in sample size calculation section)
Methodology (e.g. whether use the correct data analysis method)	seek the advice from statisticians
Approval (e.g. whether the study is approved by IRB, any data sources approval?)	Apply the approvals early



Proposal writing – Mentor and Overseas training program

Components		Advices
Agreement from mentor and	- A	Approach potential them early, and work
overseas supervisors (if any)	t	he research project with them early
Additional to overseas training progra	m	
Set clear learning objectives	- N	Must be relevant to the projects
Include deliverables and time frame	- v t	work with overseas supervisors on the training schedule
Budget	- A	Avoid over- or under-budget



Interview

Components	Advices
Prepare Your Research Narrative	 Clearly articulate your research background, key plans. Practice summarizing your work in a concise way that highlights its significance and potential impact.
Anticipate Questions	 Prepare for common interview questions, such as, challenges you've faced in research, and your long- term career goals. Formulate thoughtful questions to ask the interviewers as well.
Mock Interviews	 Conduct mock interviews with a friend, mentor, or advisor. This helps you practice answering questions and receive constructive feedback on your responses and presentation style



Challenges during the implementation phase



Challenges during the implementation phase

Challenges	Advices
COVID-19 (e.g. changing the training	- Seek the advice of oversea supervisor
period; delaying the data extraction;	and HMRF research office
project extension etc.)	



Take-home Messages



Take-home Messages

For junior research staff, the HMRF Research Fellowship Scheme presents a golden opportunity to develop your own research projects and acquire new skills and methodologies for future research

Prepare Your Application Early: Identify potential mentors and overseas supervisors at an early stage to strengthen your application.
Emphasize Your Training/Mentorship Plan: A well-structured Training and Mentorship Plan is just as important as your Research Plan. Ensure both are

clearly articulated.

Prepare Thoroughly for the Interview: Take the time to practice and prepare for the interview to present your ideas confidently and effectively.

Maintain Timely Communication: Keep in regular contact with the HMRF research office regarding any amendments to your application or budget virement to ensure a smooth process.



Acknowledgement

- HMRF research fellowship scheme (Project No. FHB/H/41/69 (Application No: 05190107))
- HMRF research office
- Co-investigators in this project
 - Professor Cindy Lam (HKU)
 - Professor Ian Wong (HKU)
 - Professor Goodarz Danaei (Harvard University)
 - Dr Esther Yu (HKU)
 - Professor Esther Chan (HKU)
 - Dr Weng Chin (HKU)
 - Dr Celine Chui (HKU)
 - Ms Wanchun Xu (HKU)
- Hospital Authority for helping data extraction
- HKU ITS High Performance Computing service