





# Age and Sex Differences in LDL Cholesterol Distribution in Adults in Malaysia: A Cross-Sectional Study (2010–2021)

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## Introduction

Cardiovascular disease (CVD) is the leading cause of death worldwide. Hypercholesterolaemia, especially higher low-density lipoprotein cholesterol (LDL-c), has been recognised as one of the major key risk factors for CVD.<sup>1</sup> Lipid-lowering treatments, such as statins, have been shown to impact both primary and secondary cardiovascular prevention.<sup>2</sup>

In Europe and America, the prevalence of hypercholesterolaemia in adults was 53.7% and 47.7%, respectively, with a lower prevalence in Southeast Asia (30.3%) and Africa (23.1%).<sup>3</sup> In Malaysia, there is an increasing trend in the prevalence of hypercholesterolaemia, with a recent prospective study of 11,288 adults reporting a prevalence of 64.0% (95% confidence interval (CI) 63.0–65.0%) and 56.7% (95% CI 55.7–57.7%) for elevated total cholesterol (ie, > 5.2 mmol/L) and elevated LDL-c (ie, > 3.4 mmol/L), respectively.<sup>4</sup>

Establishing the mean LDL-c for each age range in the population is vital for preventive and management strategies for hypercholesterolaemia. In Europe and North America, the mean LDL-c is well established,<sup>5,6</sup> however; the mean LDL-c level among the primary care population in Malaysia is not well established.

This study, therefore, aimed to describe the mean LDL-c levels in the Malaysian primary care adult population and additionally describe the age and sex differences in the mean LDL-c levels.

## Materials and Methods

A cross-sectional study was conducted in 11 out of the 21 Ministry of Health primary care clinics in Selangor, Putrajaya and Kuala Lumpur in Malaysia. Of the 11 primary care clinics involved in this study, 6 were in Selangor, and the remaining 5 were in Kuala Lumpur and Putrajaya. Demographic data (age and gender) and LDL-c readings from 2010 to 2021 were electronically extracted from the electronic medical record (EMR). Adult patients aged ≥18 years old with LDL-c results recorded in the EMR system were included in this study. Patients with missing information on age and/or sex were excluded. The study setting and population have been previously described in detail.<sup>7</sup>

Descriptive analyses were conducted to describe the mean LDL-c levels overall and by sex stratified by 10-year age band. Baseline characteristics were presented as frequencies with percentages, means with standard deviation (SD) or medians with interquartile ranges. Analyses were performed using R statistical software (version 4.0.5).

**Table 1** Baseline Characteristics of the Study Population

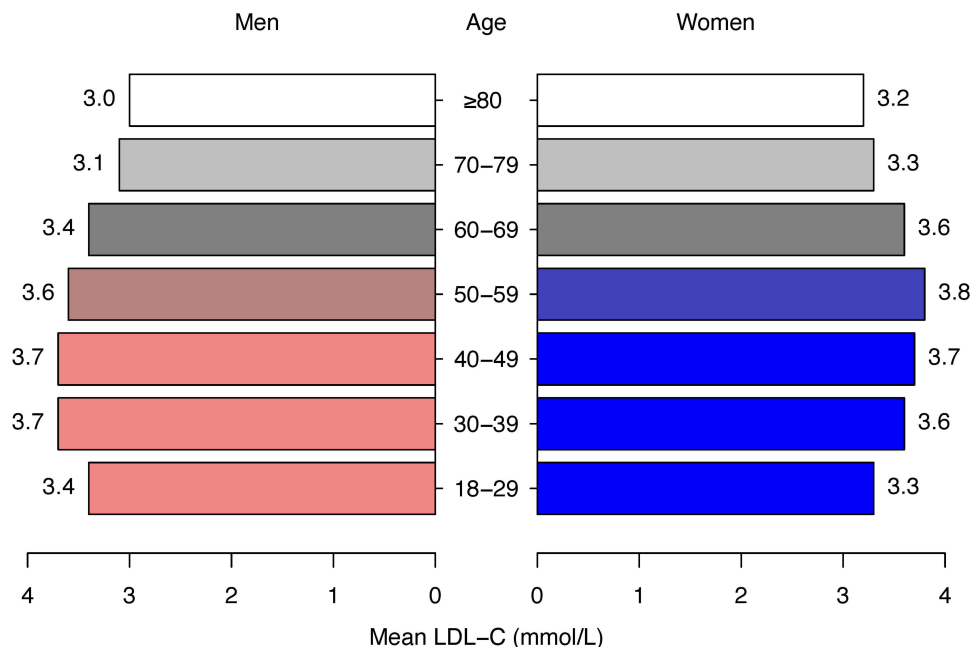
	Total n=139,702	Men n=63,674 (45.6%)	Women n=76,028 (54.4%)
Age at the time of LDL-c assessment (years)			
Mean (SD)	58.4 (14.2)	57.4 (14.5)	59.2 (13.9)
Median (IQR)	60 (49–69)	59 (47–68)	60 (50–69)
Age categories, n (%)			
18–29 years	3933 (2.8)	2123 (3.3)	1810 (2.4)
30–39 years	12,391 (8.9)	6668 (10.5)	5722 (7.5)
40–49 years	19,884 (14.2)	9657 (15.2)	10,228 (13.5)
50–59 years	32,397 (23.2)	14,285 (22.4)	18,111 (23.8)
60–69 years	39,756 (28.5)	17,502 (27.5)	22,255 (29.3)
70–79 years	23,415 (16.8)	10,275 (16.1)	13,140 (17.3)
≥80 years	7926 (5.7)	3164 (5)	4762 (6.3)
Primary care clinic area, n (%)			
Urban	88,641 (63.4)	40,973 (64.3)	47,669 (62.1)
Sub-urban	51,061 (36.6)	22,701 (35.7)	28,359 (37.3)

**Notes:** All figures as presented as absolute numbers/frequency (percentages), unless otherwise specified.

**Abbreviations:** IQR, interquartile range; LDL-c, low-density lipoprotein cholesterol; n, frequency; SD, Standard Deviation; %, percentage.

## Results

A total of 139,702 participants were included in the analysis. The number of study participants from each primary care clinic varied, ranging from 6189 (4.4%) to 19,781 (14.2%). The mean age was  $58.4 \pm 14.2$  (range, 18 to 108) years old, with the majority of participants being women (54.4%) (Table 1). The mean (SD) LDL-c level of the overall population was 3.5 (1.10) mmol/l. The mean LDL-c levels ranged from 3.0 to 3.7 mmol/L and 3.2 to 3.8 mmol/L in men and women, respectively (Figure 1). Women aged 50 to 59 years had the highest overall mean LDL-c level (3.8 mmol/L), while the highest mean LDL-c in men (3.7 mmol/L) was in those aged 30 to 49 years. Men had higher LDL-c levels compared to women between 18 to 39 years old; however, this was reversed after the age of 50 years old.



**Figure 1** Mean LDL-c (mmol/L) stratified by 10-year age bands and sex.

## Discussion

This study using electronic medical records from 11 primary care clinics in Malaysia found that the highest mean LDL-c levels were in individuals aged 30 to 59 years old. Additionally, we observed sex and age-related variations in LDL-c levels. Men had higher LDL-c levels compared to women between the ages of 18 to 39 years old. However, this pattern was reversed after the age of 50 years, with women having higher LDL-c levels. Furthermore, we found a consistent decline in LDL-c measurements after the age of 60 years in our study population.

## Relationship with Other Literature

Our findings align with previous research, particularly a prospective study of 11,288 Malaysian adults, which reported similar mean LDL-c levels across the various age groups. The highest mean LDL-c level reported in this Malaysian study was 3.79 ( $\pm$  1.14) mmol/L among individuals aged 50 to 60 years old.<sup>4</sup> Furthermore, our finding of higher LDL-c in women aged 51 to 60 years old is consistent with previous literature, and this phenomenon may be partially explained by hormonal changes associated with menopause.<sup>8</sup> The population-based Malaysian Cohort Study of 106,527 recruited individuals reported a 44.9% prevalence of hypercholesterolaemia with similar mean LDL-c levels across the various age-sex groups.<sup>9</sup>

## Strengths and Limitations

This study has several strengths. It is the first study in Malaysia reporting on mean LDL-c levels using primary care EMR data. The primary strength of this study is the inclusion of a large sample size.

However, some limitations should be mentioned. The LDL-c testing was not part of a general screening programme but following consultation in primary care clinics. In younger individuals, cholesterol testing/assessment may be initiated due to existing cardiovascular risk factors (such as a family history of cardiovascular diseases or raised LDL-c). The data was not nationally representative as the selection of the clinics was purposely undertaken for the FAMCAT study,<sup>7</sup> and involved three out of 14 states and federal territories in Malaysia. Additionally, only data on age and sex were extracted from the EMR for this study, it was therefore impossible to assess the relationship between age and/or sex and LDL-c levels, adjusting for other confounding variables such as dietary habits, physical activity, and medication use, which are known to impact LDL-c levels.<sup>10</sup> Given these limitations, the interpretation and generalisation of the results to the broader Malaysian population should be made with caution.

## Clinical Implications

Our study findings have significant clinical implications for the management of cardiovascular health in Malaysia. Firstly, the age-specific variations in LDL-c levels underscore the importance of tailoring screening and intervention strategies to different age groups. For instance, healthcare professionals could closely monitor LDL-c levels in individuals aged 30 to 59, as they exhibited the highest mean LDL-c levels. Additionally, the reversal of sex-related LDL-c differences after age 50 suggests that post-menopausal women may require closer attention regarding their cardiovascular risk factors, including LDL-c levels. Furthermore, the declining trend in LDL-c levels after the age of 60 raises questions about the potential benefits of adjusting cholesterol-lowering interventions in older individuals. Future research should investigate whether the age-related decrease in LDL-c levels in this population is associated with improved cardiovascular outcomes or if additional factors play a role.

In conclusion, our study provides valuable insights into the distribution of mean LDL-c levels across different age groups and sex in the Malaysian primary care population. Due to the limitations of our data, further research is warranted to comprehensively understand the complex relationship between age, sex, and LDL-c levels in the Malaysian population. These findings could, however, help in developing tailored strategies for cardiovascular risk assessment and management.

## Ethics

The study was approved by the respective research ethics committees in Malaysia, that is, the UiTM Research Ethics Committee [(REC/03/2020) (FB/48)] and the Medical Research Ethics Committee of the Ministry of Health Malaysia [NMRR20-272-52797 (IIR)].

## Acknowledgment

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## Data Sharing Statement

Raw data are kept at the Institute of Pathology, Laboratory and Forensic Medicine (I-PPerForM), Universiti Teknologi MARA, Selangor, Malaysia and encrypted data are kept at the University of Nottingham, United Kingdom in secured databases. Unidentified data will be shared by the corresponding authors upon request after all the results of this study have been published, and the data are subjected to the data protection regulations of Malaysia and the United Kingdom.

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## Disclosure

The authors report no conflicts of interest in this work.

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