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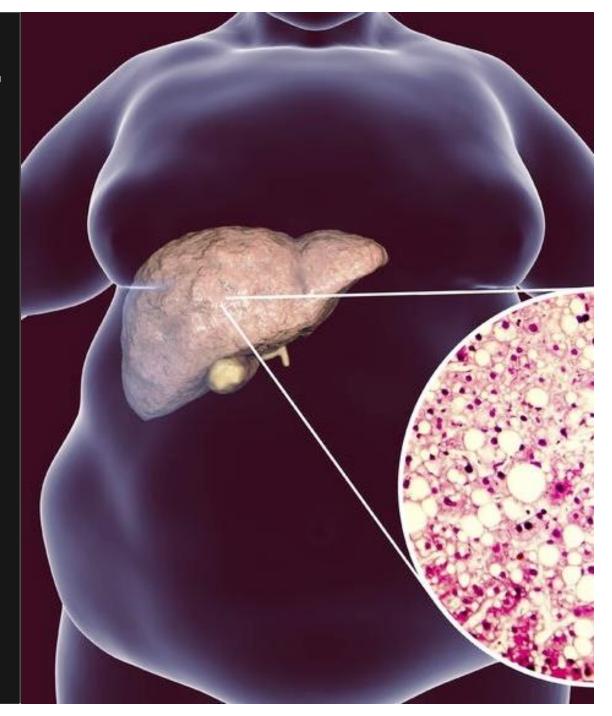
## WHO'S AT RISK?

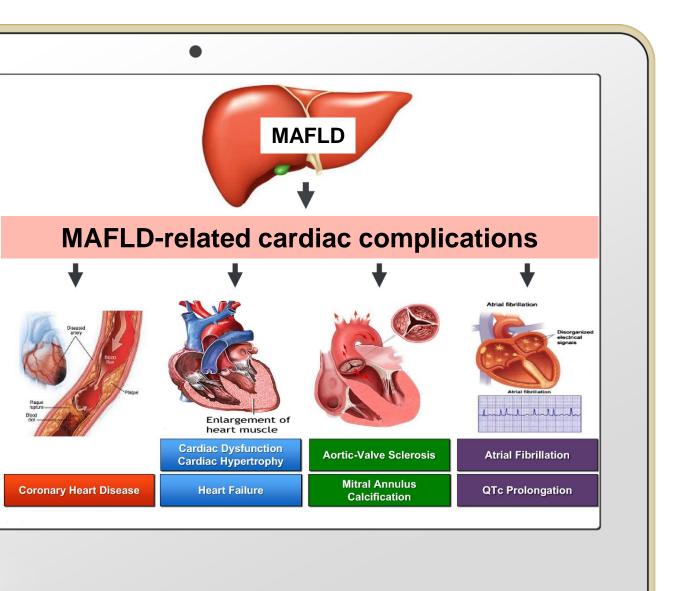
Improving Cardiovascular Outcomes in Patients with MAFLD

Professor Dr. Anis Safura Ramli MBBS (Newcastle, UK), MRCGP (UK), Fellow in Chronic Disease Management (Monash, AUS) Consultant Family Medicine Specialist Deputy Director & Fellow of I-PPerForM Research Centre of Excellence in Atherosclerosis and CVD Prevention Universiti Teknologi MARA

## PRESENTATION OUTLINE

- MAFLD and CVD: Partners in Crime?
- Who's at risk? Prevalence of Metabolic Syndrome in Malaysia
- How common is MAFLD in Primary Care?
- Who should be screened for MAFLD?
- How do we improve cardiovascular outcomes of patients with MAFLD?
- Take home message





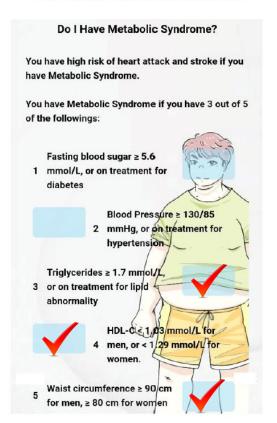


# MAFLD and CVD: Partners in Crime?

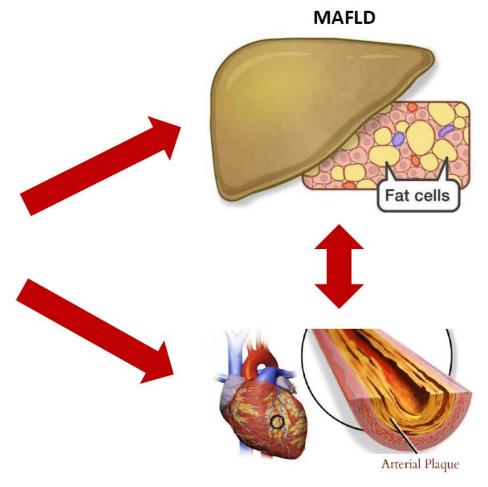
### MAFLD AND CVD: PARTNERS IN CRIME?



#### METABOLIC SYNDROME



JIS 2009 Definition



MAFLD and CVD are both manifestations of end-organ damage of the Metabolic Syndrome

**ASCVD** 

Medina-Santillán R, López-Velázquez J, Chávez-Tapia N, et al . Hepatic manifestations of metabolic syndrome.

Diabetes Metab Res Rev. 2013;7:1–16. https://doi.org/10.1002/dmrr.2410

## PATHOPHYSIOLOGICAL MECHANISMS LINKING MAFLD AND CVD



#### Lifestyle - high risk factors

- Sedentary lifestyle
- Western-style dietary pattern (high in energy, SFA, salt, trans-fatty acids, refined carbohydrates)
- Smoking
- Psychological stress
- •Sleep depriviation / disorders (e.g. OSAS)

#### **Body composition**

- Total body weight ↑
   Visceral fat ↑
- Peripancreatic fat ↑
   Epicardial fat ↑
  - •Hepatic fat ↑

#### Lifestyle - low risk factors

- Physical activity
- Balanced eating habbits
   (e.g. low-carb. Mediterranean diet, marine n-3 FA)
- •Restful/ restorative sleep pattern
- ·Low chronic stress
- Smoke-free environment
- ·Low ambient noise and air pollution

## **MAFLD**

#### Genetics/ Epigenetics

NAFLD risk ↑
PNPLA3
TM6SF

NAFLD risk ↓ HSD17B13 MARC1

Epigenetics

Altered DNA-methylation
(e.g. PGC1α)

Aberrant microRNA
-profile
(e.g. miR-122)

#### Endothelial dysfunction

ADMA ↑
eNOS ↓
Homocysteine ↑
↓
Vascular tone

dysregulation
Oxidative stress ↑

Platelet activation↑

#### Atherogenic dyslipidemia

Hepatic
VLDL, IDL, LDL
production↑

ApoB (ApoC III) ↑

Hepatic DNL↑ SFA (e.g. C16:0) ↑

Dyslipoproteinemia

VLDL↑,IDL↑,HDL↓ sdLDL ↑,LDL-P ↑

### Systemic/Vascular inflammation

NLRP3-Inflammasome activation (e.g. by C16:0)

> ↓ IL-1ß-IL-6-CRP

pathway 1

Inflammatory milieu

TNF-α↑,IL-1β↑,IL6↑ Adiponectin ↓

#### Altered glucose metabolism

Insulin resistance ↑
Insulin clearance ↓
Hyperinsulinemia ↑

Hepatic gluconeogenesis/

Dysmetabolic milieu

Hyperglycemia, AGEs↑, FFA's ↑

### Coagulopathy/ Plaque formation

Protein-C activity ↑
Factor VIII,IX,XI,XII ↑
Fibrinolysis ↓

▼ Procoagulatory imbalance ↑

VEGF ↑

Plaque formation/ vulnerability↑

### Altered gut microbiome

Gut-barrier dysfunction ("leaky gut")

> MAMP's (e.g. LPS)

Bacterial metabolites (e.g. SCFA,TMA)

Secondary bile acids

Kasper P, Martin A, Lang S, et al. NAFLD and cardiovascular diseases: a clinical review. Clinical Research in Cardiology. 2020.

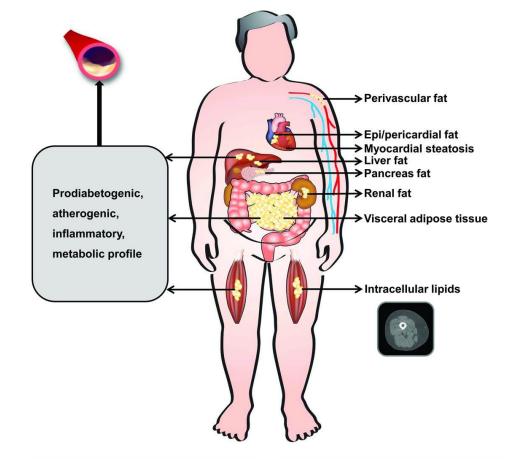
https://doi.org/10.1007/s003



## PATHOGENIC RELATIONSHIP BETWEEN

MAFLD AND CVD

 Ectopic fatty tissue depositions in the liver and the heart explain the central pathogenic relationship between MAFLD and CVD

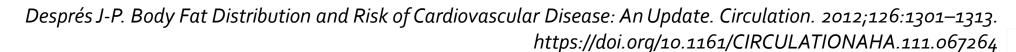


#### Ectopic fat depots with systemic effects:

- Liver fat
- Visceral adipose tissue
- Intracellular lipids
- Pancreas fat

#### Ectopic fat depots with local effects:

- Perivascular fat
- Epi/pericardial fat
- Renal fat
- Etc.

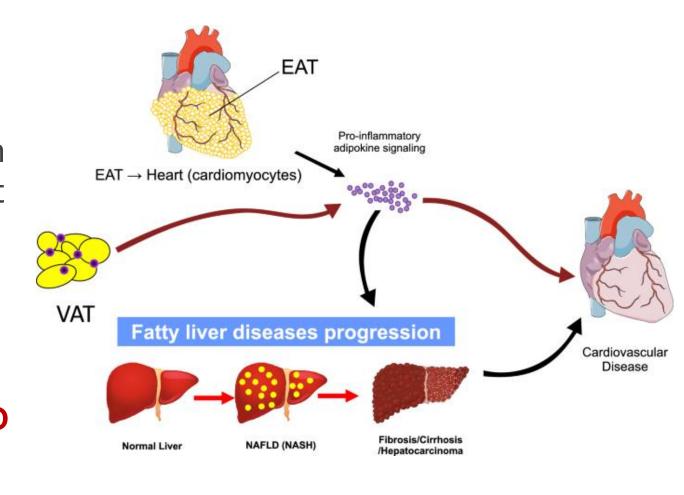


## PATHOGENIC RELATIONSHIP BETWEEN MAFLD AND CVD



 A recent meta-analysis of 2260 individuals found that Epicardial Adipose Tissue (EAT) was significantly increased in those with MAFLD compared to those without MAFLD.

 The increase in EAT was associated with the severity of hepatic steatosis, hepatic fibrosis and CVD in patients with MAFLD.



Liu B, Li YR, Li Y, et al. Association of epicardial adipose tissue with non-alcoholic fatty liver disease: A metaanalysis. Hepatol. Int. 2019; 13:757–765. https://doi.org/10.1007/s12072-019-09972-1 page 8

## IMPACT OF MAFLD ON CVD MORBIDITY AND **MORTALITY**



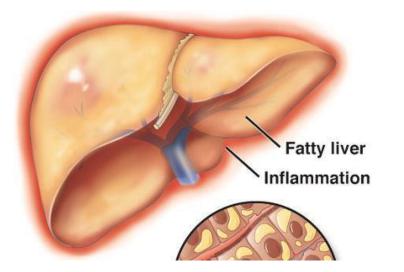
- A meta-analysis of 16 observational studies consisting of 34,043 patients with a median 7-year follow-up
- Patients with MAFLD had a higher risk of fatal and/or nonfatal CVD events than those without MAFLD [random effect OR 1.64; 95% CI 1.26-2.13]



## IMPACT OF MAFLD SEVERITY ON CVD MORBIDITY AND MORTALITY

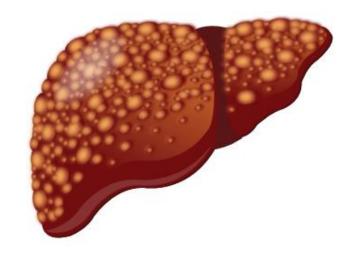


Patients with more 'severe'
 MAFLD were also more likely
 to develop fatal and non fatal CVD events [OR 2.58;
 95% Cl 1.78-3.75]



**MASH** 

**CIRRHOSIS** 



## IMPACT OF MAFLD ON CVD MORBIDITY AND MORTALITY



RESEARCH

Non-alcoholic fatty liver disease and risk of incident acute myocardial infarction and stroke: findings from matched cohort study of 18 million European adults

Myriam Alexander, <sup>1</sup> A Katrina Loomis, <sup>2</sup> Johan van der Lei, <sup>3</sup> Talita Duarte-Salles, <sup>4</sup> Daniel Prieto-Alhambra, <sup>5</sup> David Ansell, <sup>6,7</sup> Alessandro Pasqua, <sup>8</sup> Francesco Lapi, <sup>8</sup> Peter Rijnbeek, <sup>3</sup> Mees Mosseveld, <sup>3</sup> Paul Avillach, <sup>3,9</sup> Peter Egger, <sup>1</sup> Nafeesa N Dhalwani, <sup>10</sup> Stuart Kendrick, <sup>11</sup> Carlos Celis-Morales, <sup>12</sup> Dawn M Waterworth, <sup>13</sup> William Alazawi, <sup>14\*</sup> Naveed Sattar<sup>12\*</sup>

#### WHAT IS ALREADY KNOWN ON THIS TOPIC

Non-alcoholic fatty liver disease (NAFLD) is associated with metabolic syndrome and other risk factors for acute myocardial infarction (AMI) or stroke NAFLD is associated with increased risk of AMI and stroke and cardiovascular surrogate markers

The association between NAFLD and AMI and stroke after adjustment for established risk factors has yet to be fully established however

#### WHAT THIS STUDY ADDS

In four large European databases, the adjusted hazard ratios for incident AMI or stroke diagnoses in adults with NAFLD were modest and not significantly greater than those in age, sex, and general practice matched participants without NAFLD

- Design: Matched cohort study.
- Setting: Population based, electronic primary healthcare databases from four European countries: Italy, Netherlands, Spain and UK.
- Participants: 120 795 adults with a recorded diagnosis of MAFLD or MASH, matched by age, gender, practice site and visit, with patients without MAFLD or MASH in the same database.
- Mean follow-up: 2.1-5.5 years.

the**bmj** | *BMJ* 2019;367:15367 | doi: 10.1136/bmj.15367

## IMPACT OF MAFLD ON CVD MORBIDITY AND MORTALITY



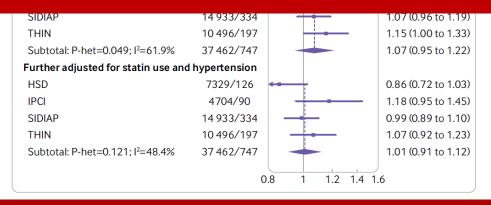
#### Hazard ratios for incident AMI

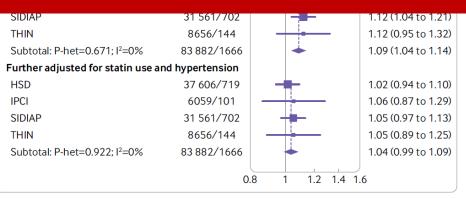
Database	Events in non- NAFLD/NAFLD	Hazard ratio (95% CI)	Hazard ratio (95% CI)
Total population			
Adjusted for age and smoking st	atus		
HSD	15 014/221	<del></del>	1.03 (0.90 to 1.18
IPCI	9625/137		1.27 (1.07 to 1.50
SIDIAP	23 238/414		1.11 (1.01 to 1.22
THIN	19 946/263	<del></del>	1.31 (1.16 to 1.49
Subtotal: P-het=0.032; I <sup>2</sup> =66.0%	67 823/1035		1.17 (1.05 to 1.30
Subset*			

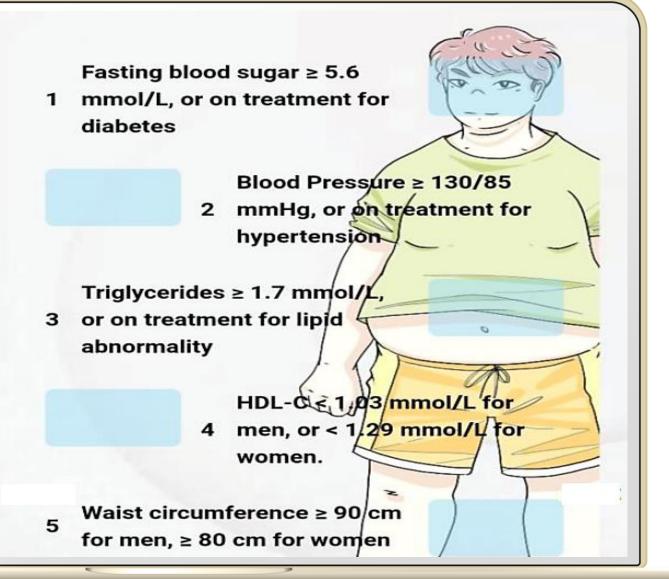
#### Hazard ratios for incident stroke

Database	Events in non- NAFLD/NAFLD	Hazard ratio (95% CI)	Hazard ratio (95% CI)
Total population			
Adjusted for age and smoking st	atus		_
HSD	60 082/962		1.16 (1.09 to 1.2
IPCI	11 902/156	+	1.15 (0.99 to 1.3
SIDIAP	45 658/854	-	1.14 (1.07 to 1.2
THIN	16 359/215	-	1.34 (1.17 to 1.5
Subtotal: P-het=0.236; I <sup>2</sup> =29.3%	134 001/2187	-	1.18 (1.11 to 1.2
Subset*			

- The diagnosis of MAFLD in 17.7 million patients in primary care appears not to be associated with AMI or stroke risk after adjustment for established CVD risk factors.
- Cardiovascular risk assessment in adults with a diagnosis of MAFLD is important but should be done in the same way as for the general population.









Who's at Risk?
Prevalence of
Metabolic
Syndrome in
Malaysia





#### Research Article

### JIS Definition Identified More Malaysian Adults with Metabolic Syndrome Compared to the NCEP-ATP III and IDF Criteria

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43.4% OF
MALAYSIAN
ADULTS ≥ 30
YEARS OF AGE
HAVE
METABOLIC
SYNDROME

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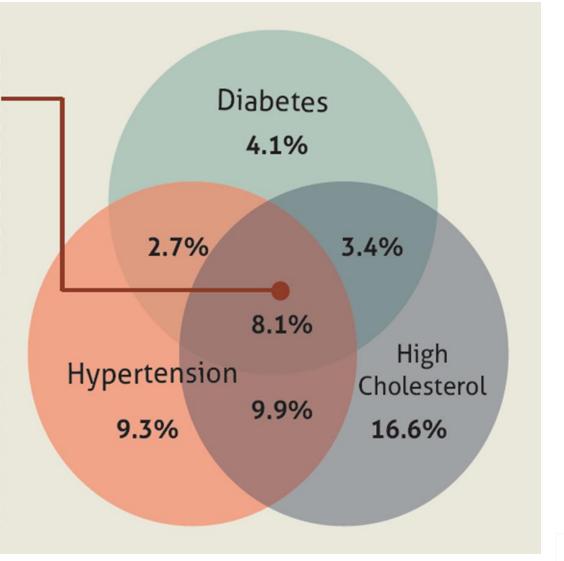
<sup>&</sup>lt;sup>6</sup>Cardiology Discipline, Faculty of Medicine, Universiti Teknologi MARA, 47000 Sungai Buloh, Selangor, Malaysia

## NHMS 2019: CLUSTERING OF RISK FACTORS





3.4 million people in Malaysia currently live with two major risk factors

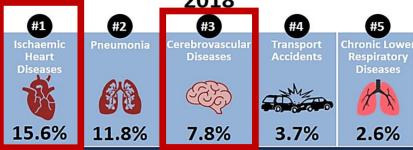


## CVD: PRINCIPAL CAUSE OF DEATH IN MALAYSIA

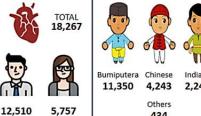




5 PRINCIPAL CAUSES OF DEATH 2018







15-40 years

#### Average death 12,101 Urban 6,166 Rural 434 persons

41-59 years

The principal causes of death by age group

0-14 years Pneumonia

Note: The analysis is based on medically certified



causes of death

Accidents

Transport **Ischaemic Heart** 

**Ischaemic Heart** Diseases Diseases

> Source: Statistics on Causes of Death, Malaysia, 2019 Department of Statistics Malaysia

60 years and over

### Heart attack leading cause of deat

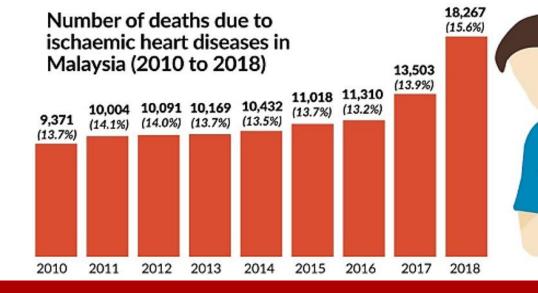
#### NATION

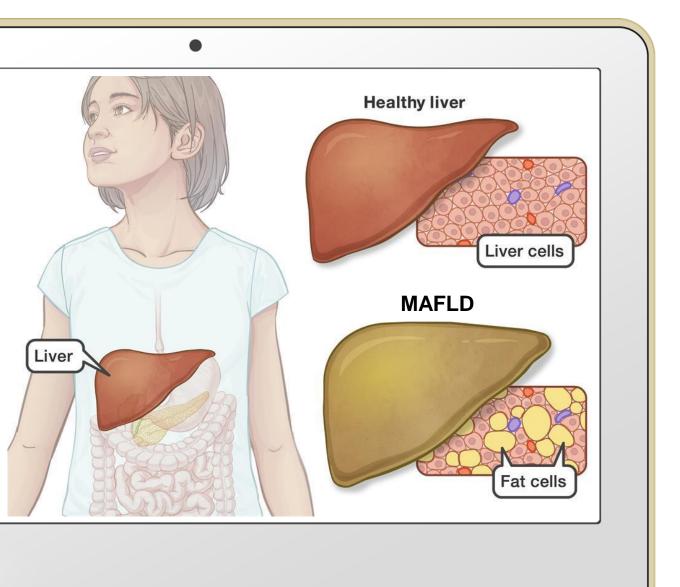
Thursday, 31 Oct 2019

By LOH FOON FONG

KUALA LUMPUR: Heart attack remains the leading cause of death in Malay

**AVERAGE AGE OF HEART ATTACK IN MALAYSIA IS** 58 YEARS OLD!







# How Common is MAFLD in Primary Care?

### PREVALENCE OF MAFLD IN PRIMARY CARE

Miptah et al. BMC Family Practice (2020) 21:238 https://doi.org/10.1186/s12875-020-01306-7

**BMC Family Practice** 

#### RESEARCH ARTICLE

**Open Access** 

Non-alcoholic fatty liver disease (NAFLD) and the cardiovascular disease (CVD) risk categories in primary care: is there an association?



Hayatul Najaa Miptah<sup>1</sup>, Anis Safura Ramli<sup>1,2\*</sup>, Mariam Mohamad<sup>3</sup>, Hilwati Hashim<sup>4</sup> and Zahirah Tharek<sup>1</sup>

#### **Abstract**

Background: Non-alcoholic fatty liver disease (NAFLD) is an emerging novel cardiovascular disease (CVD) risk factor. It's prevalence is increasing globally. However, there is paucity in the evidence showing the association between NAFLD and CVD risk in primary care setting. Therefore, the objectives of this study were to determine the prevalence and factors associated with NAFLD among patients with ≥1 risk factor for NAFLD or CVD attending primary care clinics.

Methodology: A cross sectional study was conducted in two clinics at a university primary care centre. Patients aged ≥18 years with ≥1 risk factor for NAFLD or CVD were recruited. Participants with history of established liver disease or chronic alcohol use were excluded. Socio-demographics, clinical related data, anthropometric measurements and blood investigation results were recorded in a proforma. Diagnosis of NAFLD was made using abdominal ultrasound. The 10-year CVD risk was calculated using the general Framingham Risk Score (FRS). Multiple logistic regression (MLogR) was performed to identify independent factors associated with NAFLD.

**Results:** A total of 263 participants were recruited. The mean age was  $52.3 \pm 14.7$  years old. Male and female were equally distributed. Majority of the participants were Malays (79.8%). The overall prevalence of NAFLD was 54.4% (95%CI 48,60%). Participants in the high FRS category have higher prevalence of NAFLD (65.5%), followed by those in the moderate category (55.4%) and the low category (46.3%), p = 0.025. From MLogR, independent factors associated with NAFLD were being employed (OR = 2.44, 95%CI 1.26,4.70, p = 0.008), obesity with BMI ≥27.5 (OR = 2.89, 95%CI 1.21,6.91, p = 0.017), elevated fasting glucose ≥5.6 mmol/L (OR = 2.79, 95%CI 1.445.43, p = 0.002), ALT ≥34 U/L (OR = 3.70, 95%CI 1.85,7.44, p < 0.001) and high FRS category (OR = 2.82, 95%CI 1.86,6.23, p = 0.010).

(Continued on next page)

#### **BMC FAMILY PRACTICE**

Impact Factor

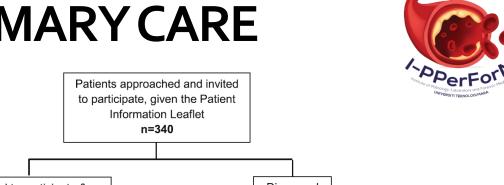
2.022 2.469

2019 5 year

JCR ® Category	Rank in Category	Quartile in Category
MEDICINE, GENERAL & INTERNAL	66 of 165	Q2







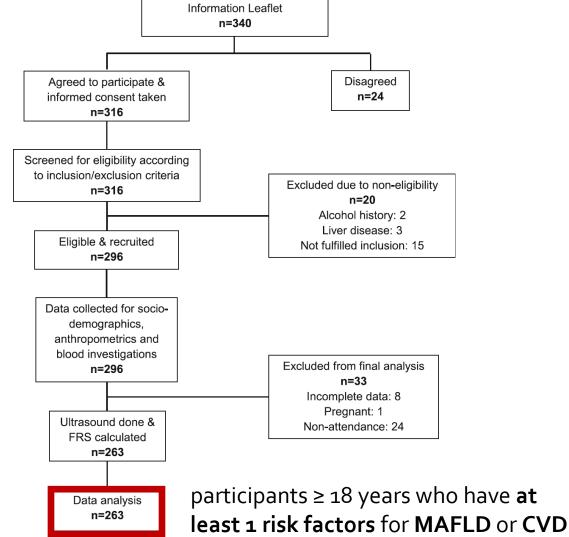


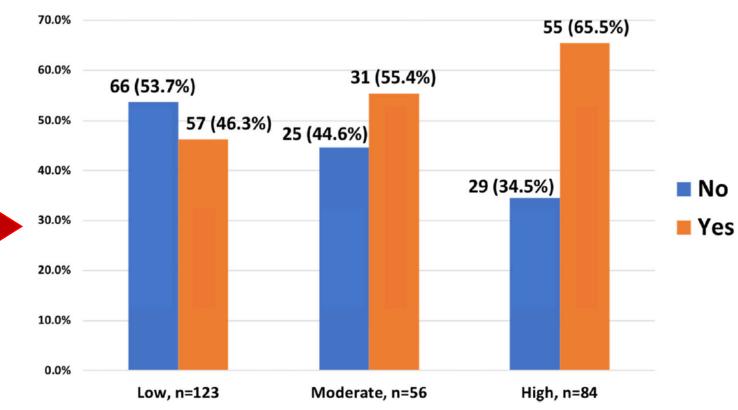
Fig. 1 Flow chart of conduct of the study

page **18** 

### PREVALENCE OF MAFLD IN PRIMARY CARE







- Fig. 2 Prevalence of NAFLD according to the FRS category
- Participants with high FRS category had a greater prevalence of MAFLD (p = 0.025)
- The mean FRS score was significantly higher in individuals with MAFLD compared to those without MAFLD (17.38  $\pm$  12.35 vs. 12.35  $\pm$ 12.89, p = 0.003)

## FACTORS ASSOCIATED WITH MAFLD

PerForm Recognition of the Control o

**Table 2** Factors independently associated with NAFLD (MLogR)

Variables	Adj Beta (SE)	Wald (df)	Adj. OR (95%CI)	<i>P</i> -value
Occupational sector:		_	-	_
Not working	REF		1.00	
Working	0.89 (0.335)	7.071 (1)	2.44 (1.26,4.70)	0.008
BMI:				
Not-obese	REF		1.00	
Obese	1.060 (0.445)	5.679 (1)	2.89 (1.21,6.91)	0.017
FPG				
< 5.6 mmol/L	REF		1.00	
≥ 5.6 mmol/L	1.027 (0.339)	9.169 (1)	2.79 (1.44,5.43)	0.002
ALT				
≤ 34 U/L	REF		1.00	
> 34 U/L	1.310 (0.355)	13.587 (1)	3.70 (1.85, 7.44)	< 0.001
FRS category				
Low	REF		1.00	
Moderate	0.388 (0.413)	0.884 (1)	1.47 (0.66,3.31)	0.347
High	1.038 (0.403)	6.620 (1)	2.82 (1.28,6.23)	0.010

Notes:

OR Odds Ratio, CI Confidence interval, df Degree of freedom, REF Reference group

The model reasonably fits well (Hosmer-Lemeshow test: p = 0.168)

Model assumptions were met

No significant interactions and multicollinearity problem

Model explained between 23.1% (Cox and Snell R Square) and 30.8% (Nagelkerke R Square) of the variance in NAFLD group and correctly classified 73.4% of cases

## SUMMARY OF MAIN FINDINGS AND IMPLICATIONS FOR CLINICAL PRACTICE



- MAFLD is highly prevalent (54.4%) in patients with at least one risk factor in our primary care setting.
- Patients with at least one CVD or MAFLD risk factor should be risk stratified using the 10-year general CVD FRS.
- If they are found to have **high FRS**, or **obese** or have **elevated FPG** or **elevated ALT**, they are recommended to have a **liver ultrasound to screen for MAFLD**.
- If they are found to have MAFLD, then the severity of the condition should be assessed using scoring such as NFS or FIB-4 to identify those who need referral to the hepatologist.
- Regardless of their MAFLD status, these patients should be targeted for aggressive lifestyle intervention and risk factor management.

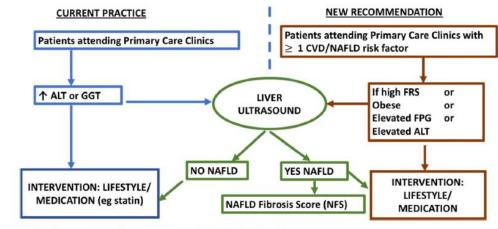
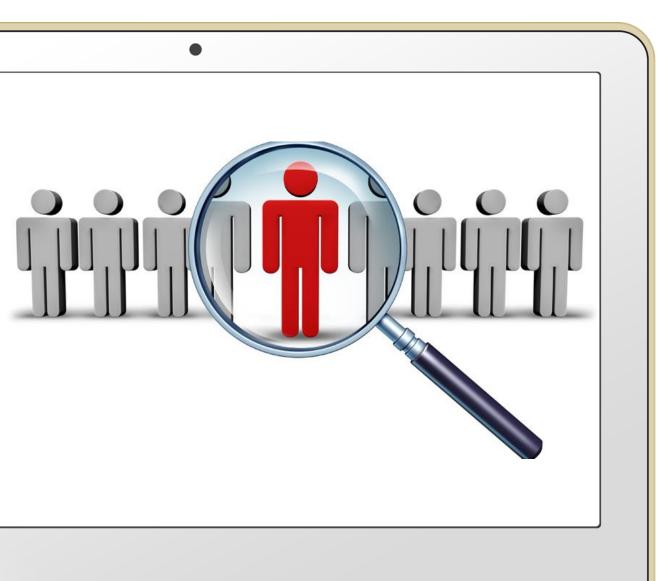


Fig. 3 Proposed algorithm for screening of NAFLD in the target groups in Primary Care

Miptah HN, Ramli AS, et al. Non-alcoholic fatty liver disease (NAFLD) and the cardiovascular disease (CVD) risk categories in primary care: is there an association? BMC Family Practice. 2020; 21:238 https://doi.org/10.1186/s12875-020-01306-7





Who Should Be Screened for MAFLD?

## RECOMMENDATIONS BY THE APASL GUIDELINE



Hepatology International (2020) 14:889–919 https://doi.org/10.1007/s12072-020-10094-2

#### **GUIDELINES**



The Asian Pacific Association for the Study of the Liver clinical practice guidelines for the diagnosis and management of metabolic associated fatty liver disease

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Received: 8 July 2020 / Accepted: 6 September 2020 / Published online: 1 October 2020 © Asian Pacific Association for the Study of the Liver 2020

#### Abstract

Metabolic associated fatty liver disease (MAFLD) is the principal worldwide cause of liver disease and affects nearly a quarter of the global population. The objective of this work was to present the clinical practice guidelines of the Asian Pacific Association for the Study of the Liver (APASL) on MAFLD. The guidelines cover various aspects of MAFLD including its

Electronic supplementary material The online version of this article (https://doi.org/10.1007/s12072-020-10094-2) contains supplementary material, which is available to authorized users.

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- Department of Gastroenterology, School of Medicine, Marmara University, Istanbul, Turkey

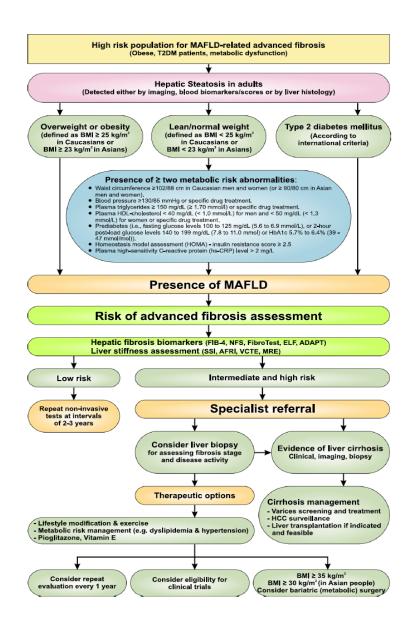
Should the high-risk population be screened for MAFLD?

#### Recommendations

- Screening for MAFLD by ultrasonography should be considered in at-risk populations such as patients with overweight/obesity, T2DM and metabolic syndrome (A1).
- Patients with MAFLD should be assessed for other components of metabolic syndrome and be treated accordingly (A1).
- Patients with MAFLD should receive advice and support for lifestyle interventions to reduce the risk of events from metabolic and cardiovascular disease, and to resolve fatty liver disease (A1).
- Screening for MAFLD in the population at risk should be in the context of the available resources, considering the burden for the national health care systems

## RECOMMENDATIONS BY THE APASL GUIDELINE



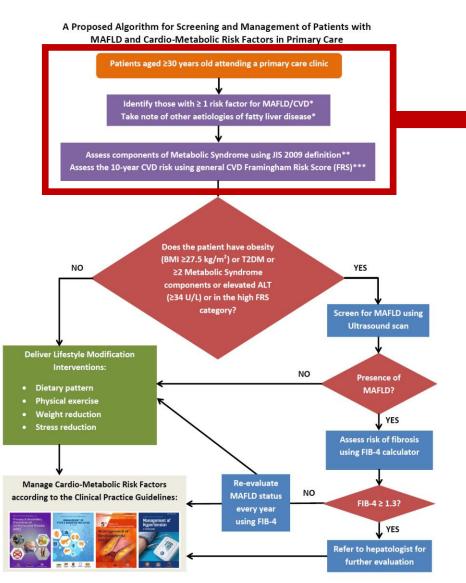


Targeted screening of high risk patients who **fulfil at least one** of these criteria:

- ☐ Overweight & Obese
- ☐ T2DM
- □ Presence of ≥ 2 Metabolic Syndrome components:
  - WC  $\geq$  90cm (men),  $\geq$  80cm (women)
  - BP ≥ 130/85 mmHg or on treatment
  - TG ≥ 1.7 mmol/L or on treatment
  - HDL ≤ 1.0 mmol/L (men), ≤ 1.3 mmol/L (women) or on treatment
  - FBS 5.6 6.9 mmol/L

## PROPOSED ALGORITHM FOR SCREENING AND MANAGEMENT OF MAFLD IN MALAYSIAN PRIMARY CARE





### Screen for \*risk factors in patients ≥30 years old:

- abnormal waist circumference (WC) ≥80 cm in women or ≥90 cm in men
- elevated blood pressure (BP) ≥130/85mmHg or on treatment for hypertension
- impaired fasting glucose (IFG) ≥5.6 mmol/L or random glucose ≥7.8 mmol/L or elevated HbA1c ≥7.0% or on treatment for elevated glucose or known Type 2 Diabetes Mellitus (T2DM)
- dyslipidaemia (TC ≥5.0 mmol/L, LDL-C ≥2.6, TG ≥1.7 mmol/L, HDL-C <1.0 mmol/L in men or HDL-C <1.3 mmol/L in women)</li>
- abnormalities of liver enzymes (ALT ≥34 U/L or GGT >60 U/L)

Identification of these patients has the potential to detect those at high cardio-metabolic risk who are candidates for therapeutic interventions aimed at prevention of MAFLD progression as well as ASCVD

## SCREENING FOR PATIENTS WITH METABOLIC SYNDROME IN MALAYSIAN PRIMARY CARE



Do I Have Metabolic Syndrome?

You have high risk of heart attack and stroke if you have Metabolic Syndrome.

You have Metabolic Syndrome if you have 3 out of 5 of the followings:

Fasting blood sugar ≥ 5.6 1 mmol/L, or on treatment for diabetes

Blood Pressure ≥ 130/85

2 mmHg, or on treatment for hypertension

Triglycerides ≥ 1.7 mmol/L,

3 or on treatment for lipid abnormality

HDL-Ci 1,03 mmol/L for

4 men, or < 1/29 mmol/L for women.

Waist circumference ≥ 90 cm for men, ≥ 80 cm for women Patients aged ≥30 years old attending a primary care clinic should be assessed for the presence of Metabolic Syndrome components using JIS 2009 definition

Alberti KGMM, Eckel RH, Grundy SM, Zimmet PZ, Cleeman JI, Donato KA, et al.
Harmonizing the metabolic syndrome: a joint interim statement of the
International Diabetes Federation Task Force on Epidemiology and Prevention;
National Heart, Lung, and Blood Institute; American Heart Association; World
Heart Federation; International Atherosclerosis Society; and International
Association for the Study of Obesity. Circulation. 2009; 120:1640–5

Ramli AS, Daher AM, Nor-Ashikin MN, et al. JIS Definition Identified More Malaysian Adults with Metabolic Syndrome Compared to the NCEP-ATP III and IDF Criteria. Biomed Res Int. 2013; 760963. https://doi.org/10.1155/2013/760963

## CVD RISK STRATIFICATION FOR PATIENTS AGED ≥30YEARS IN MALAYSIAN PRIMARY CARE



10-YEAR GENERAL	CVD FRAMINGHAM	<b>RISK SCORE</b>
TO- I LAV GLIJEVAL		VION OCOUL

Very High Risk	
■ 10-year CVD risk of > <b>30</b> %	
■ Established CVD	
■ Diabetes mellitus with proteinuria	
■ Stage 4 & 5 chronic kidney disease	
High Risk	
■ 10-year CVD risk of <b>21-29</b> %	
■ Diabetes mellitus without target of damage	
■ Stage 3 chronic kidney disease	
■ Very high levels of individual risk factors (LDL-C > 4.9 mmol/L, BP > 180/110 mmHg)	
Intermediate (Moderate) Risk	Clinic
■ 10-year CVD risk of <b>10-20</b> %	<b>-</b>

■ 10-year CVD risk of < **10**%

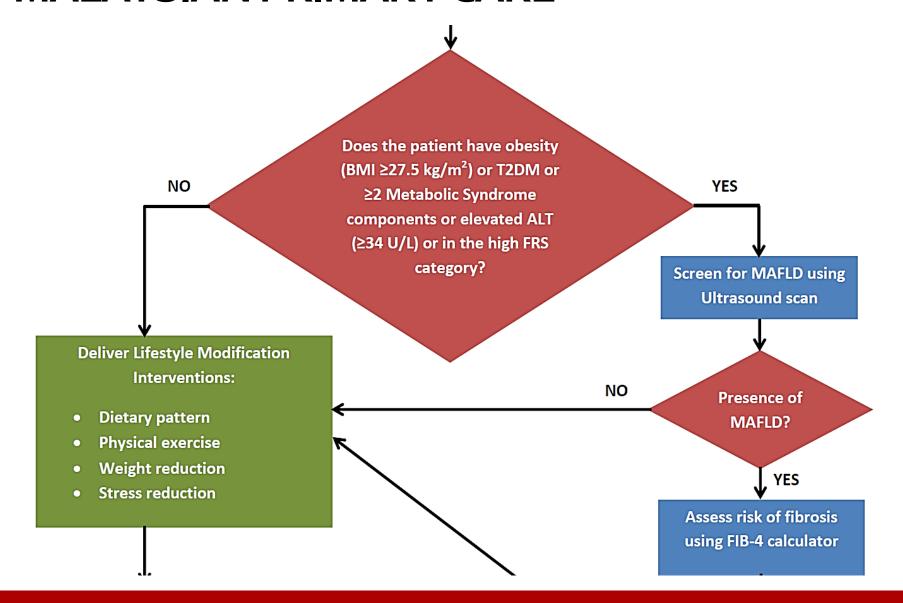
- Patients aged ≥30 years old attending a primary care clinic should be risk stratified using the 10-year general CVD Framingham Risk Score (FRS)
- The cut-off age of ≥30 years is recommended as the prevalence of cardio-metabolic risk factors rise exponentially in Malaysian adults aged ≥30 years

Clinical Practice Guidelines on the Primary & Secondary Prevention of Cardiovascular Disease 2017. Putrajaya: Ministry of Health Malaysia, 2017. https://www.moh.gov.my/moh/resources/Penerbitan/CPG/CARDIOVASCULAR/3.pdf

D'Agostino RB Sr, Vasan RS, Pencina MJ, Wolf PA, Cobain M, Massaro JM, et al. General cardiovascular risk profile for use in primary care: the Framingham Heart Study. Circulation. 2008; 117:743–53 page 27

## PROPOSED ALGORITHM FOR SCREENING OF MAFLD IN MALAYSIAN PRIMARY CARE

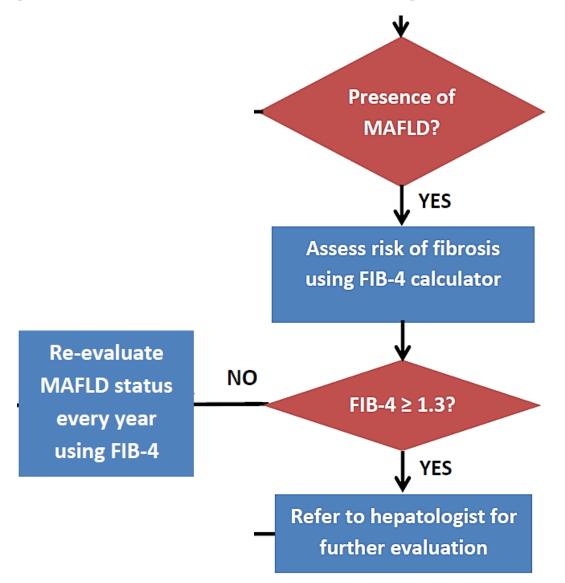




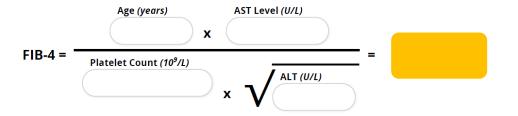
If they are found to have obesity (BMI ≥27.5 kg/m2) or T2DM or ≥2 Metabolic Syndrome components or elevated ALT (≥34 U/L) or in the high FRS category, they are recommended to have a liver ultrasound to screen for MAFLD

## PROPOSED ALGORITHM FOR ASSESSING MAFLD SEVERITY IN MALAYSIAN PRIMARY CARE



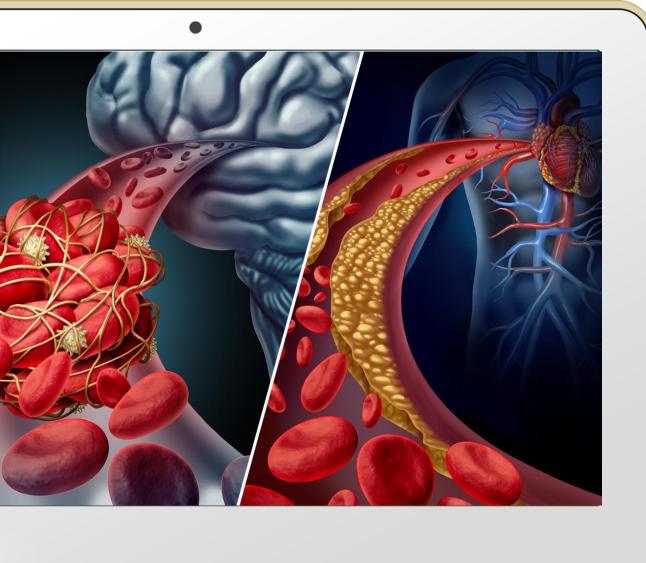


 If they are found to have MAFLD, then the severity of the condition should be assessed using FIB-4 scoring



- Those with FIB-4 of ≥ 1.3 should be referred to hepatologist for further evaluation
- Those with FIB-4 of < 1.3 should be reevaluated annually

Chan WK, Treeprasertsuk S, Goh GB, et al. Optimizing Use of Nonalcoholic Fatty
Liver Disease Fibrosis Score, Fibrosis-4 Score, and Liver Stiffness Measurement to
Identify Patients with Advanced Fibrosis. Clin Gastroenterol Hepatol. 2019;
17(12):2570-2580.e37. https://doi.org/10.1016/j.cgh.2019.03.006 page 29

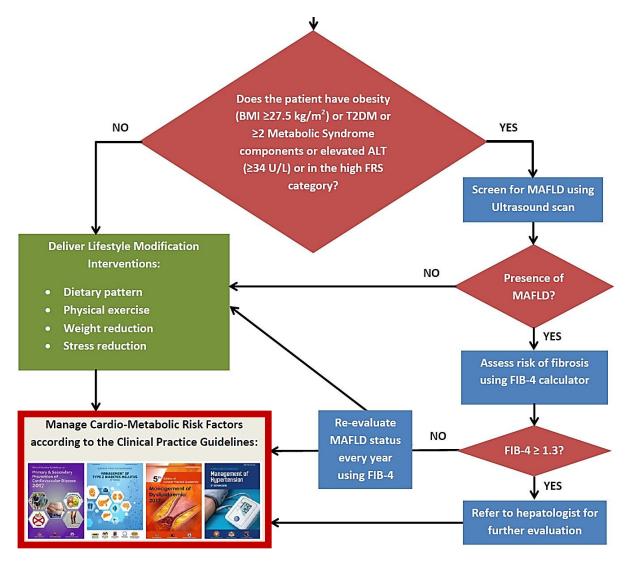




How do we improve cardiovascular outcomes of patients with MAFLD?

## MANAGEMENT OF PATIENTS WITH MAFLD TO IMPROVE THEIR CARDIOVASCULAR OUTCOMES

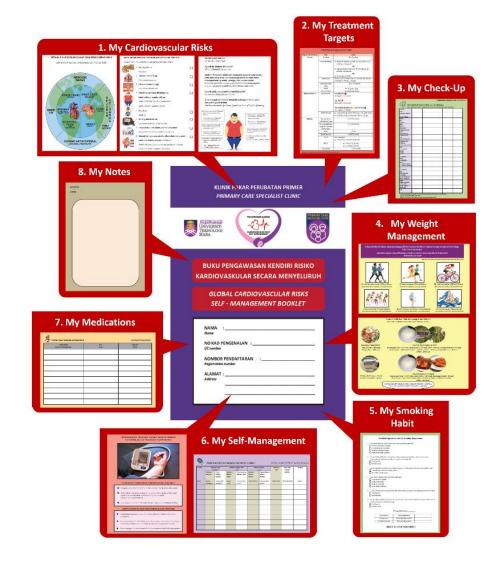




- Patients with MAFLD and the coexisting cardio-metabolic risk factors should be targeted for aggressive lifestyle intervention and risk factor management in accordance with the relevant Clinical Practice Guidelines
- The ultimate management goals for these patients are to prevent the progression of MAFLD and to prevent cardio-metabolic complications



KLINIK PAKAR PERUBATAN PRIMER PRIMARY CARE SPECIALIST CLINIC
UNIVERSITI TEKNOLOGI MARA  CABDIOVASCULAR RISKS SEL MANAGEMENT
BUKU PENGAWASAN KENDIRI RISIKO KARDIOVASKULAR SECARA MENYELURUH  GLOBAL CARDIOVASCULAR RISKS SELF - MANAGEMENT BOOKLET
NAMA :





RISIKO KARDIOVASKULAR SAYA | MY CARDIOVASCULAR RISKS

RISIKO KARDIOVASKULAR SAYA | MY CARDIOVASCULAR RISKS

#### SAYA MEMPUNYAI RISIKO KARDIOVASKULAR BERIKUT:

I HAVE THE FOLLOWING CARDIOVASCULAR RISKS



1. Kencing Manis



2. Tekanan darah tinggi



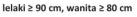


3. Paras kolesterol tinggi





Obesiti sentral ukur lilit pinggang:



Central Obesity (waist circumference:

men ≥ 90 cm, women ≥ 80 cm



Merokok

Smoking



6. Kurang aktiviti fizikal





7. Umur ( lelaki > 55 tahun, wanita > 65 tahun

Age (men > 55 years, women > 65 years)



8. Sejarah keluarga penyakit kardiovaskular yang awal

lelaki < 55 tahun, wanita < 65 tahun)

Family history of premature cardiovascular disease

(men < 55 years, women < 65 years)

#### SINDROM METABOLIK

**METABOLIC SYNDROME** 

#### Apakah itu Sindrom Metabolik?

What is Metabolic Syndrome?

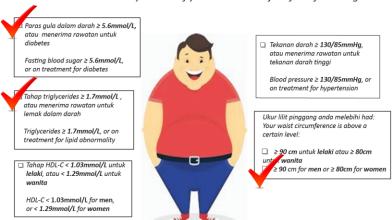
Sindrom Metabolik adalah satu kumpulan penyakit yang berlaku pada masa yang sama, dan ia meninggikan risiko anda untuk mendapat kencing manis, serangan jantung dan stroke Metabolic syndrome is a group of conditions occurring together that put you at risk of diabetes, heart disease and stroke

#### Adakah anda mengalami Sindrom Metabolik?

Do you have Metabolic Syndrome?

#### Anda mengalami Sindrom Metabolik sekiranya mempunyai 3 daripada 5 gejala berikut:

You have Metabolic Syndrome if you have 3 out of 5 of the following:



K. G. M. M. Alberti, R. H. Eckel, S. M. Grundy et al., "Harmonizing the metabolic syndrome: a joint interim statement of the international diabetes lederation task force on epidemiology and prevention, National Heart, Lung, and Blood Institute; American Heart Association; World Heart Federation; International Association (Atherosclerosis Society; and International Association for the Study of Obesity, "Circulation, vol. 126, no. 16, pp. 1640–1645, 2009



RISIKO KARDIOVASKULAR SAYA | MY CARDIOVASCULAR RISKS

RISIKO KARDIOVASKULAR SAYA | MY CARDIOVASCULAR RISKS

#### **SAYA MEMPUNYAI KOMPLIKASI BERIKUT:**

I HAVE THE FOLLOWING COMPLICATIONS



1. Serangan jantung/ penyakit jantung koronari Heart attack/ coronary heart disease





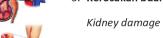
2. Strok/angin ahmar





3. Kerosakan buah pinggang



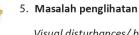


4. Penyakit vaskular periferi



Peripheral vascular disease





Visual disturbances/blindness



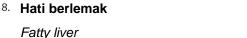
6. Hipertrofi ventrikal kiri





Peripheral neuropathy

7. Gangguan saraf periferi





#### SKOR RISIKO FRAMINGHAM

	Skor Risiko Framingham Saya (%
Risiko Yang Sangat Tinggi	
■ Risiko penyakit Kardiovaskular dalam 10 tahun > <b>30</b> %	
■ Disahkan mempunyai penyakit Kardiovaskular	
Kencing manis dan protein dalam urin	
■ Penyakit buah pinggang kronik tahap 4 & 5	
Risiko Tinggi	. /
<ul><li>Risiko penyakit Kardiovaskular dalam 10 tahun antara 21-29%</li></ul>	25%
■ Kencing manis tanpa kerosakan organ	04
■ Penyakit buah pinggang kronik tahap 3	25%
■ Tahap faktor risiko yang sangat tinggi (LDL-C > 4.9 mmol/L, BP > 180/110 mmHg)	
Risiko Sederhana	
Risiko penyakit Kardiovaskular dalam 10 tahun antara <b>10-20</b> %	
Risiko Rendah	
■ Risiko penyakit Kardiovaskular dalam 10 tahun < 10%	

\*D'Agostino RB Sr. Vasan RS. Pencina MJ. Wolf PA. Cobain M. Massaro JM. Kannel WB: General cardiovascular risk profile for use in primary care:the Framingham Heart Study. Circulation. 2008; 117(6):743



		SASARAN RAWATAN SAYA   MY TREATMENT TARGET
	KETAHUI SASA	RAN RAWATAN ANDA
Kategori Individu	Risiko	Sasaran
Umum	Merokok	Berhenti Merokok
	Aktiviti Fizikal	Aktiviti fizikal sederhana: 150 minit/ minggu iaitu 30 minit/hari, 5 hari/seminggu
		Gabungan kedua-duanya
<u> </u>	Penurunan Berat Badan	Sasarkan untuk menurunkan 5-10% berat badan dalam tempoh 6 bulan dan mengekalkan berat badan 1-2 tahun akan datang
	Indeks Jisim Tubuh (BMI)	18.5 – 22.9kg/m²
	Ukur lilit Pinggang	● < 90 cm untuk lelaki ● < 80cm untuk wanita
Tanpa Kencing Manis	Dislipidemia	Risiko Yang Sangat Tinggi  Sasaran LDL-C: < 1.8mmol/L  Risiko Tinggi  Sasaran LDL-C: < 2.6mmol/L  Pertengahan (Sederhana) dan Risiko Rendah  Sasaran LDL-C: < 3.0mmol/L
	Tekanan Darah	<ul> <li>◆ &lt; 140/90mmHg untuk kebanyakan individu &lt; 80 tahun</li> <li>◆ &lt; 150/90mmHg untuk individu &gt; 80 tahun</li> </ul>
Kencing Manis	Paras gula dalam darah sebelum makan atau semasa berpuasa	4.4 – 7.0 mmol/L
	Paras gula dalam darah selepas makan (90-120 minit selepas makan)	4.4 – 8.5 mmol/L
	HbA1c	≤ 6.5%
	Tekanan Darah	≤ 135/75 mmHg
	LDL-C	
	HDL-C	● > 1.0 mmol/L (lelaki) ● > 1.2 mmo/L (wanita)
	Triglycerides	≤ 1.7 mmol/L

TARIKH			Т	$\neg$
Date				
TEKANAN DARAH				П
Blood Pressure				
BERAT				
(kg)				
Weight				_
BMI (kg/m²)				
UKUR LILIT				
PINGGANG (cm) Waist				
Vircumference				
PEMERIKSAAN				_
KAKI				
Foot Assessment				
PEMERIKSAAN				_
FUNDUS				
Fundus Assessment				
FBS				_
< 6.1 mmol/L				_
HbA1c				
< 6.5 %				_
TC < 5 mmol/L				╝
LDL-C				
< 2.6 mmol/L				_
HDL-C				
> 1.0 mmol/L (male) > 1.2 mmol/L (female)				
TG (Terriale)				-
< 1.7 mmol/L				
ALT				_
(Liver Function)				
< 40 μmol/l				_
Serum Creatinine				
(Kidney Function)	 			
eGFR				
(Kidney Function) > 90 mL/min				
Urine Protein /				$\dashv$
Urine ACR				
ECG				

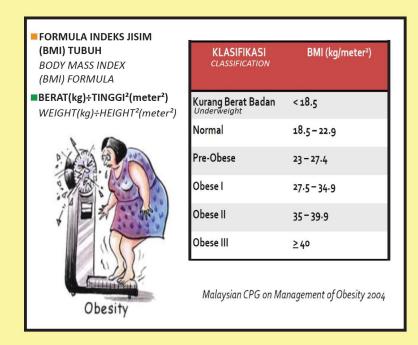
PEMERIKSAAN RAWATAN SAYA | MY CHECK-UF

Malaysian CPG on Primary & Secondary Prevention of Cardiovascular Disease, 2017



PENGURUSAN BERAT BADAN SAYA | MY WEIGHT MANAGEMENT

### INDEKS JISIM TUBUH SAYA MY BODY MASS INDEX



BERAT BADAN SAYA	-	85 <sub>kg</sub>
MY WEIGHT		
BMI SAYA	33	kg/m²
MY BMI		

PENGURUSAN BERAT BADAN SAYA | MY WEIGHT MANAGEMENT

### UKUR LILIT PINGGANG SAYA MY WAIST CIRCUMFERENCE



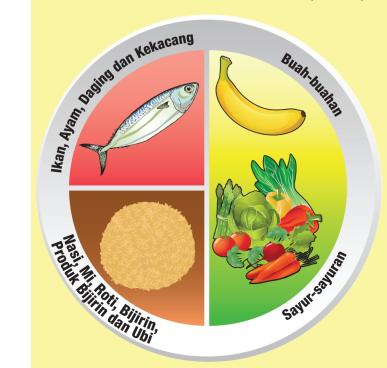
- < 90 cm untuk lelaki atau < 80 cm untuk wanita
- < 90 cm for men or < 80 cm for women

UKUR LILIT PINGGANG SAYA 98 \_\_\_\_cm



PENGURUSAN BERAT BADAN SAYA | MY WEIGHT MANAGEMENT

**Pinggan Suku Suku Separuh** Quarter Quarter Half Plate



- Makan 3 kali sel3 regular meals
- Hidangan tanpa Non-fried and so
- Makan malam s Have dinner before
- 1-2 hidangan sn 1-2 servings of h
- Makan makanar Have home cool
- Minum 8 gelas a Drink 8 glasses (

Malaysian CPG on Primary & Second

Setiap individu memerlukan sekurang-kurangnya 30 minit senaman berintensiti sederhana setiap hari selama 5 hari/minggu (150 minit setiap minggu)

Individuals require at least 30 minutes of moderate intensity exercise per day for 5 days/week

(150 minutes per week)

Malaysian CPG on Primary & Secondary Prevention of Cardiovascular Disease, 2017



30 minit berjalan pantas: 130 kcal dibakar 30 minutes brisk walking: 130 kcal burned



30 minit aerobik: 175 kcal dibakar 30 minutes aerobics: 175 kcal burned



30 minit berjoging: 215 kcal dibakar 30 minutes jogging: 215 kcal burned



30 minit bermain badminton: 135 kcal dibakar 30 minutes playing badminton: 135 kcal burned



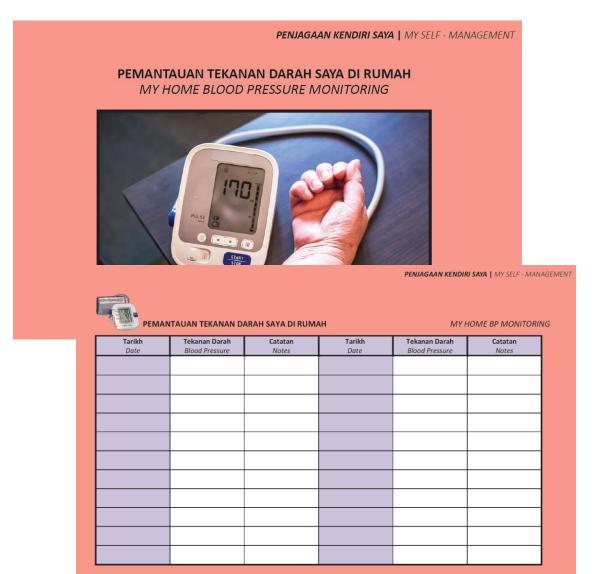
PENGURUSAN BERAT BADAN SAYA | MY WEIGHT MANAGEMENT

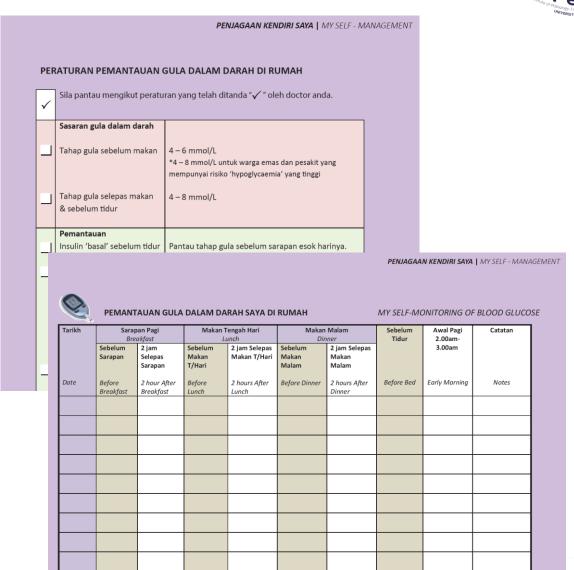
30 minit berbasikal: 160 kcal dibakar 30 minutes cycling: 160 kcal burned



30 minit berenang: 300 kcal dibakar 30 minutes swimming: 300 kcal burned







## EVIDENCE SUPPORTING THE USE OF SELF-MANAGEMENT BOOKLET IN PRIMARY CARE



Ramli et al. BMC Family Practice (2016) 17:157 DOI 10.1186/s12875-016-0557-1

**BMC Family Practice** 

Scientific Foundation SPIROSKI, Skople, Republic of Macedonia Open Access Macedonian Journal of Medical Sciences. 2020 May 30; 8(B):470-479. https://doi.org/10.3889/oamjms.2020.3764 elSSN: 1857-9655 Category: B - Clinical Sciences



#### **RESEARCH ARTICLE**

Open Access
CrossMark

Effectiveness of the EMPOWER-PAR Intervention in Improving Clinical Outcomes of Type 2 Diabetes Mellitus in Primary Care: A Pragmatic Cluster Randomised Controlled Trial

Anis Safura Ramli<sup>1,2\*</sup>, Sharmini Selvarajah<sup>3</sup>, Maryam Hannah Daud<sup>1,2</sup>, Jamaiyah Haniff<sup>4</sup>, Suraya Abdul-Razak<sup>1,2</sup>, Tg Mohd Ikhwan Tg-Abu-Bakar-Sidik<sup>4</sup>, Mohamad Adam Bujang<sup>4</sup>, Boon How Chew<sup>5</sup>, Thuhairah Rahman<sup>2</sup>, Seng Fah Tong<sup>6</sup>, Asrul Akmal Shafie<sup>7</sup>, Verna K. M. Lee<sup>8</sup>, Kien Keat Ng<sup>9</sup>, Farnaza Ariffin<sup>1</sup>, Hasidah Abdul-Hamid<sup>1</sup>, Md Yasin Mazapuspavina<sup>1</sup>, Nafiza Mat-Nasir<sup>1</sup>, Chun W. Chan<sup>8</sup>, Abdul Rahman Yong-Rafidah<sup>10</sup>, Mastura Ismail<sup>11</sup>, Sharmila Lakshmanan<sup>4</sup>, Wilson H. H. Low<sup>12</sup> and for the EMPOWER-PAR Investigators

Effectiveness of the EMPOWER-PAR Intervention on Primary Care Providers' Adherence to Clinical Practice Guideline on the Management of Type 2 Diabetes Mellitus: A Pragmatic Cluster Randomised Controlled Trial

Maryam Hannah Daud<sup>1,2</sup>, Anis Safura Ramli<sup>1,2</sup>\*, Suraya Abdul-Razak<sup>1,2</sup>, Jamaiyah Haniff³, Tg Mohd Ikhwan Tg Abu Bakar Sidik³, Nur Khairul Bariyyah Mohd Hatta³, Sarimah Mahmood¹, Sharmila Lakshmanan³

<sup>1</sup>Department of Primary Care Medicine, Faculty of Medicine, Universiti Teknologi MARA, Selangor, Malaysia; <sup>2</sup>Institute of Pathology, Laboratory and Forensic Medicine (I-PPerForM), Universiti Teknologi MARA, Selangor, Malaysia; <sup>3</sup>Clinical Epidemiology Unit, Clinical Research Centre, Ministry of Health, Kuala Lumpur, Malaysia

#### Abstract

Edited by: Slavica Hristomanova-Mitkovska Citation: Daud MH, Ramli AS, Abdul-Razak S, Haniff J, Tg-Abu-Bakar-Sidik TMI, Mohd-Hatta NKB, Mahmood S, Lakshmana S, Effectiveness of the EMPOWICE DAD AIM: The objective of this study was to evaluate the effectiveness of the EMPOWER- PAR intervention, a multifaceted strategy based on the chronic care model (CCM) on primary care providers' (PCP) adherence to type

Utilisation of the booklet as part of the multifaceted intervention has been shown to be effective in improving glycaemic control in patients with diabetes and in improving adherence to CPG among primary care providers in Malaysia

OPEN ACCESS

were designed based on four elements of the chronic care model i.e. healthcare organisation, delivery system design, self-management support and decision support. The primary outcome was the change in the proportion of patients achieving HbA1c < 6.5%. Secondary outcomes were the change in proportion of patients achieving targets for blood pressure, lipid profile, body mass index and waist circumference. Intention to treat analysis was performed for all outcome measures. A generalised estimating equation method was used to account for baseline differences and clustering effect.

(Continued on next page)

Hatts, Sammah Mahmood, Shammila Lakshmanan Funding: This work was supported by the Ministry of Higher Education (MOHE) Malaysia: Exploratory Research Grant Scheme (ERGS) no: ERGS/PHASE 1-2011Helean and Clinical Sciences/Universitä Teknologi MARAY (JPT S (BRV) 2000/690/1019 890 / 600-RMIECH SCIENCES 5/3 (28/2011) and by the MOH Malaysia: Major Research Grant Scheme (GMRRE ID-1125-88789) (CONCLINE) The EMPC/MER PAR intervention has been provided the property of the Park intervention has been provided the property of the property of the provided th

Competing interests: The authors have declared that no competing interests exist.

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CONCLUSION: The EMPOWER-PAR intervention has been proven to be effective in improving the PCPs' adherence to T2DM CPG in several indicators of care. Findings from this study provided objective evidence of the effectiveness of multifaceted intervention based on the CCM in the Malaysian public primary care setting.

TRIAL REGISTRATION: Registered with: ClinicalTrials.gov: NCT01545401. Date of registration: 1st March 2012.

## THE EMPOWER-SUSTAIN E-HEALTH INTERVENTION PROJECT



KLINIK PAKAR PERUBATAN PRIMER
PRIMARY CARE SPECIALIST CLINIC



BUKU PENGAWASAN KENDIRI RISIKO KARDIOVASKULAR SECARA MENYELURUH

GLOBAL CARDIOVASCULAR RISKS
SELF - MANAGEMENT BOOKLET

NAIVIA Name	:		 
NO KAD	PENGENALAN :_		 
I/C numbe	r		
NOMBO	R PENDAFTARAN	:	
Registrati	on number		
ALAMA	г:		
Address			

PRGS/MOHE 600-IRMI/PRGS 5/3

(003/2019)

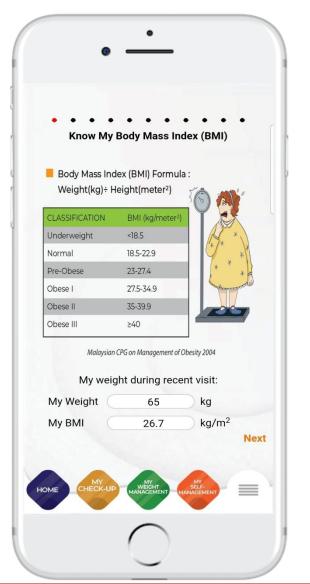


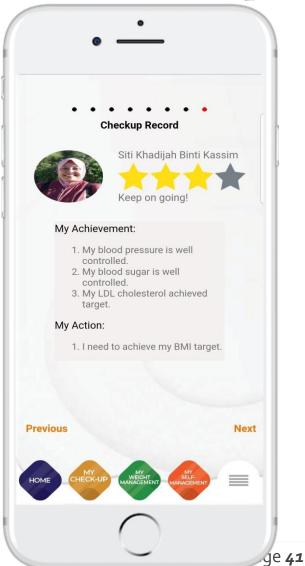
### THE EMPOWER-SUSTAIN MOBILE APP





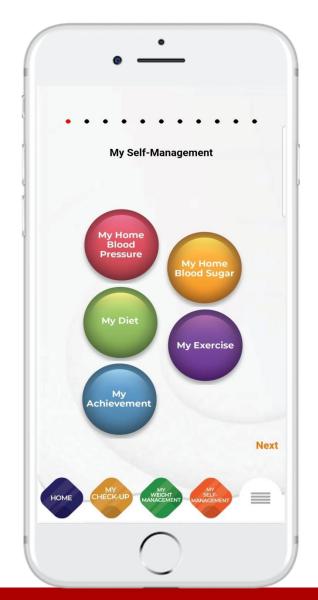




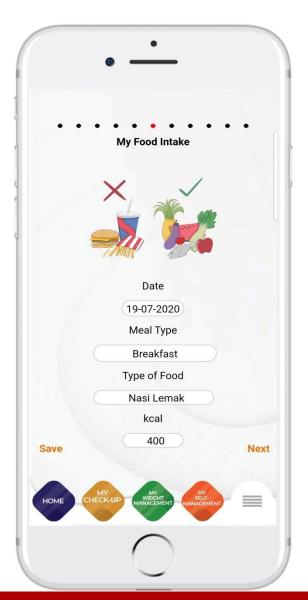


### THE EMPOWER-SUSTAIN MOBILE APP









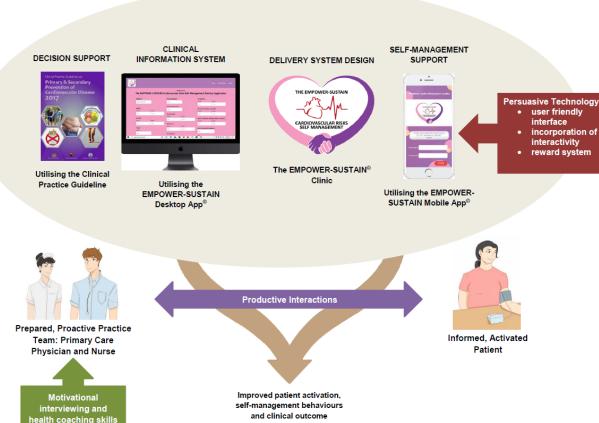


## THE EMPOWER-SUSTAIN E-HEALTH INTERVENTION

PROTOCOL PAPER

https://doi.org/10.1186/s13063-020-04237-x





https://trialsjournal.biomedcentral.com/articles/10.11 86/s13063-020-04237-x

#### **STUDY PROTOCOL**

**Open Access** 



The EMPOWER-SUSTAIN e-Health Intervention to improve patient activation and self-management behaviours among individuals with Metabolic Syndrome in primary care: study protocol for a pilot randomised controlled trial

Maryam Hannah Daud<sup>1,2</sup>, Anis Safura Ramli<sup>1,2\*</sup>, Suraya Abdul-Razak<sup>1,2</sup>, Mohamad Rodi Isa<sup>3</sup>, Fakhrul Hazman Yusoff<sup>4</sup>, Noorhida Baharudin<sup>2</sup>, Mohamed Syarif Mohamed-Yassin<sup>2</sup>, Siti Fatimah Badlishah-Sham<sup>2</sup>, Azlina Wati Nikmat<sup>5</sup> Nursuriati Jamil<sup>4</sup> and Hapizah Mohd-Nawawi<sup>1</sup>

#### **Abstract**

Background: Epidemiological studies conducted in various parts of the world have clearly demonstrated that metabolic syndrome (MetS) is an increasing global health problem, not only in Western societies but also in Asian populations. Web-based and mobile phone-based self-management applications have been proven to be effective in improving self-management behaviour of patients with MetS components (i.e., diabetes or hypertension). However, evidence is lacking in terms of their effectiveness specifically for patients with MetS. The aim of this pilot study is to evaluate the feasibility and potential effectiveness of the EMPOWER-SUSTAIN Self-Management e-Health Intervention in improving activation and self-management behaviours among patients with MetS. This paper presents the study protocol.

(Continued on next page)



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## TAKE HOME MESSAGE



01

MAFLD and CVD are both manifestations of end-organ damage of the Metabolic Syndrome

02

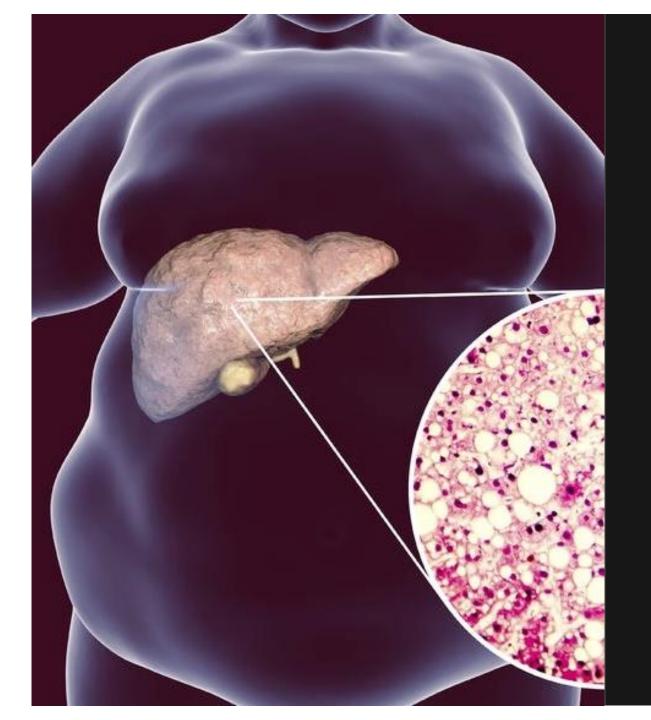
MAFLD is highly prevalent in Malaysian primary care

03

MAFLD should be screened in patients aged ≥30 years old who have cardiometabolic risk factors

04

The goal of management is to prevent cardio-metabolic complications and progression to liver fibrosis





## ThankYou

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