ORIGINAL RESEARCH

Unconventional Cardiovascular Risk Factors and Systematic Coronary Risk Estimation (SCORE) in the Lebanese Rural Population: The Forgotten Factors

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Purpose: To evaluate the correlation between unconventional risk factors and the Systematic Coronary Risk Estimation (SCORE), and estimate the prevalence of conventional and unconventional cardiovascular (CV) risk factors in the rural Lebanese population in order to assess their CV risk.

Methods: This is a retrospective descriptive study conducted between November 2017 and June 2019 among the Lebanese rural population. The risk factors were analyzed from the files of the patients who presented for the CV disease screening days organized by a non governmental organization. The CV risk estimation tool is the SCORE. The classification of socio-economic level ranges from zero (low level) to 3 (high level).

Results: A total of 433 patients were included. The prevalence of hypertension, diabetes, dyslipidemia, smoking, and metabolic syndrome was 45.1%, 31.2%, 39.2%, 50% and 42.9% respectively. Only 13.6% of hypertensive patients and 6.7% of diabetics were controlled. A total of 0 or 1 point for the classification of socio-economic status was found in 62.6% of cases. A family history of CV diseases was present in 87.3% of participants. The SCORE was correlated with diabetes and metabolic syndrome (p = 0.000), without being correlated to socio-economic status (HR = -0.104; p = 0.059) or to family history (p = 0.834).

Conclusion: The socio-economic status and the family history of CV disease must be evaluated in addition to the classical risk calculation of the SCORE to better pinpoint the actual risk of the targeted population. The risk factors are prevalent but poorly controlled, hence the need for a national effort to ensure better care for the rural Lebanese population.

Keywords: screening, cardiovascular disease, conventional risk factors, unconventional risk factors, SCORE, Lebanon

Introduction

The synergistic nature of cardiovascular risk factors (CVRF) is well-established.¹ Therefore, there are tools used to calculate the risk of developing cardiovascular diseases (CVD), such as the Systematic Coronary Risk Estimation (SCORE) and the Pooled Cohort Equation (PCE).^{1,2}

These tools only consider conventional risk factors (CRF) which include age, sex, arterial hypertension (HTN), diabetes, dyslipidemia, and smoking status. However, there are also unconventional risk factors (URF) that significantly increase the risk of CV pathologies.^{2,3} URF such as a family history of early CV diseases, metabolic syndrome, lifestyle factors (physical activity or sedentary behavior, nutrition), socio-economic status, ethnicity, and coronary calcium score can be added to these CRF.

The prevalence of CVRF varies depending on genetic profile, environmental factors, socio-economic status, and access to healthcare.⁴ In Lebanon, in 2016, CV mortality was estimated to account for 47% of all causes of death

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combined, and 8% of the population was identified as being at high risk of CV disease. However, only 58% of them were being monitored by healthcare providers.⁵

To the best of our knowledge, there are no studies conducted on the rural Lebanese population to assess their CV risk and estimate the prevalence of CVRF. This study aims to specifically target the following CRF and URF: age, sex, hypertension, diabetes, dyslipidemia, smoking status, metabolic syndrome, socio-economic status, and familial history of early CV diseases. Our objectives are to evaluate the epidemiology of these CVRFs, calculate the CV risk using the SCORE, and establish correlations between this risk and the CVRFs included in our study, particularly the unconventional risk factors.

Methods

Study Design and Data Collection

This is a retrospective descriptive study. Data were collected with oral consent from the records of patients who attended cardiovascular disease screening days organized by the Lebanese NGO "Heartbeat" between November 2017 and June 2019 (see <u>Supplementary Material 1</u>).

The cardiovascular risk estimation tool used in this study was the SCORE, which estimates the risk of developing fatal cardiovascular disease over a 10-year period. It takes into account the following risk factors: age, gender, smoking status, systolic blood pressure (SBP), and total cholesterol levels.⁶ Since the Lebanese population is considered to be at high cardiovascular risk,⁵ the calculation and classification models used in this study are those employed for high-risk European countries⁷ (see Table 1). Table 2 and Table 3 provide a summary of the criteria used to define the conventional risk factors (URF) evaluated in this study.

It should be noted that triglycerides and total cholesterol levels were only measured during three out of the five Heartbeat screening days. Therefore, the SCORE calculation was performed based on data from those three days only.

Patients' Characteristics

We included patients who presented to the screening days, aged between 40 and 70 years old, and who gave their oral consent. Patients were excluded if they were non Lebanese or unable to give an oral consent.

Statistical Analysis

The handwritten data collected was manually computerized in May 2020 and analyzed using IBM-SPSS Statistics software, version 26.0 for Windows.

Risk of Fatal Cardiovascular Event Over 10 Years	Definition Criteria
Low	Calculated SCORE <1%
Moderate	Calculated SCORE ≥1% and <5%
High	 Calculated SCORE ≥5% and <10% Patients with a history of Cardiovascular Event Patients known to have Diabetes and 2 to 3 other Cardiovascular Risk Factors
Very High	 Calculated SCORE ≥10% Patients with uncontrolled risk factors (Total cholesterol levels ≥ 310mg/dL or Blood Pressure ≥ 180/110mmHg) Patients with uncomplicated Diabetes or with a Cardiovascular risk factor Patients taking lipid-lowering antidiabetic drugs Diabetic patients with a value of "HIGH" on hemoglucotest machine

Table I Classification of Cardiovascular Risk According to the SCORE

Risk Factors	Definition
Age and Sex	Men and Women aged between 40 and 70 years old
Systemic Hypertension	 Known hypertensive patients Patients taking anti-hypertensive drugs Hypertension defined by the European Society of Cardiology (ESC): Systolic blood pressure (SBP) ≥ 140 mmHg and/or (DBP) diastolic blood pressure ≥ 90 mmHg.¹
Diabetes	 Known Diabetic patients Patients with at least one diagnostic criteria according to the American Diabetes Association (ADA) A fasting plasma glucose (FPG) level of 126 mg/dL (7.0 mmol/L) or higher A 2-hour plasma glucose level of 200 mg/dL (11.1 mmol/L) or higher during a 75-g oral glucose tolerance test (OGTT) A random plasma glucose of 200 mg/dL (11.1 mmol/L) or higher in a patient with classic symptoms of hyperglycemia or hyperglycemic crisis (polyuria, polydipsia, unexplained weight loss, drowsiness, coma) A hemoglobin A1c (HbA1c) level of 6.5% (48 mmol/mol) or higher
Dyslipidemia	 Known dyslipidemic patients Patients taking lipid-lowering drugs (statins or Ezetimibe) SCORE ≥10%³
Smoking	Any patient who has smoked a cigarette or a hookah (water pipe) during the last 12 months is considered a smoker. A former smoker is someone who quit smoking more than a year ago. ⁴
Metabolic Syndrome	Combination of three or more of the following criteria: ⁵ (a) Waist circumference > 102cm in men or > 88cm in women (b) Triglycerides levels ≥150 mg/dL (c) Patients taking antihypertensive drugs and/or SBP ≥130 mmHg and/or (DBP) ≥85 mmHg (d) Patients taking antidiabetics or Fasting Blood Glucose ≥ 110 mg/dL.
Socio-Economic Status	The markers of Socio-Economic status that we have retained are: the work status, the educational level and the number of inhabitants per room. The elements in favor of a high socio-economic status are: self-employment or work for an employer, a high school or University diploma, and a number of inhabitants per bedroom below the median (4). The socio-economic status classification is calculated by assigning one point to each item in favor of a high level and zero points to those denoting a low level (see Table 3). The correlations are established according to the final result ranging from 0 to 3.
Family History	Occurrence of a cardiovascular event in a first-degree relative at a young age: before the age of 55 for men and before the age of 65 for women. ⁶ Family History includes Cardiovascular Risk Factors such as systemic hypertension, dyslipidemia and type 2 diabetes. ⁷

Table 2 Criteria Used in This Study for Defining the Assessed Conventional and Unconventional Risk Factors

Abbreviations: SBP, systolic blood pressure; DBP, diastolic blood pressure; ADA, American Diabetes Association; FPG, fasting plasma glucose; OGTT, oral glucose tolerance test.

Qualitative data is represented as frequencies with their percentage corresponding and the 95% Confidence Interval (CI). Continuous data not deviating from normal distribution are expressed as mean \pm standard deviation. Continuous data deviating from normal distribution and ordinal data are represented by median with its interquartile range 1 and 3 (Q1 and Q3). Spearman's rank correlation coefficient was used to correlate the continuous and ordinal data. Numerical and categorical variables were compared using the Mann–Whitney *U*-test and the Kruskal–Wallis test.

Ethical Considerations

The authors assert that all procedures contributing to this work comply with the ethical principles established in the World Medical Association's (WMA's) Helsinki Declaration, as revised in 2013, and in the Council for International Organizations of Medical Sciences (CIOMS) International Ethical Guidelines, as revised in 2016, in collaboration with the World Health Organization (WHO). Verbal informed consent was obtained because this study is done in the rural

		Point Allocated
Work Status	Self Employed	I
	Employee	I
	Unemployed	0
	Retired	0
Educational level	Illiterate	0
	Primary: no high school diploma	0
	Secondary: high school diploma	I
	University Diploma	I
Number of inhabitants per bedroom	< 4	I
	≥ 4	0

 Table 3 Classification of the Socio-Economic Status of the Studied Population

population among which many cannot read or write. Two oral consent were obtained by two different persons: the first upon arrival at the screening center and the second, prior to filling the medical files. To ensure confidentiality, all data was anonymous. The verbal informed consent process was approved by the ethics committee of Saint-Joseph University of Beirut (File code: CEHDF 1546).

Results

A total of 433 patients were included in the data analysis (see Figure 1). The average age of the participants was 55 years, with a majority of women (N=249, 57.5%). Table 4 provides a summary of the socio-demographic characteristics of the studied population.



Figure I Flowchart of the study population.

		Number of Patients	Mean	Standard Deviation		
Age		433	55	8		
		Number of Patients	Percentage	Lower Limit of Cl ^a	Higher of Cl ^a	Limit
Region	Kfarhabou	88	20.3%	16.7%	24.3%	
	Riyak	77	17.8%	14.4%	21.6%	
	Mtole	78	18.0%	14.6%	21.8%	
	Kfar-Dlaquous	94	21.7%	18.0%	25.8%	
	Amaz	96	22.2%	18.5%	26.3%	
Sex	Female	249	57.5%	52.8%	62.1%	
	Male	184	42.5%	37.9%	47.2%	
Marital Status	Single	58	13.4%	10.5%	16.9%	
	Married	356	82.4%	78.6%	85.8%	
	Divorced	3	0.7%	0.2%	1.8%	
	Widowed	15	3.5%	2.0%	5.5%	
Number of children	>4	41	15.7%	11.7%	20.5%	
	3-4	134	51.3%	45.3%	57.4%	
	I–2	52	19.9%	15.4%	25.1%	
	0	34	13.0%	9.4%	17.5%	
Work status	Self-Employed	89	21.9%	18.1%	26.1%	
	Employee	131	32.3%	27.9%	36.9%	
	Unemployed	157	38.7%	34.0%	43.5%	
	Retired	29	7.1%	4.9%	10.0%	
Education	Illiterate	29	8.4%	5.8%	11.7%	
	Primary	131	38.1%	33.1%	43.3%	
	Secondary	126	36.6%	31.7%	41.8%	
	University	58	16.9%	13.2%	21.1%	
Number of inhabitants per bedroom	Number of Patients	Mean	Standard Deviation	Median	Q١ ^ь	Q3 ^c
	340	4	2	4	3	5

Table 4 Socio-Demographic Characteristics of the Studied Population

Abbreviations: ^aCl, confidence interval; ^bQl, first quartile; ^cQ3, third quartile.

In addition to age and sex, the analyzed CRF were hypertension, diabetes, dyslipidemia, and smoking status (see Table 5). In the general population, the mean systolic blood pressure (SBP) and diastolic blood pressure (DBP) were 138 mmHg and 85 mmHg, respectively. Out of the total patients, 123 were known to be hypertensive (29.5%), and 116 were receiving antihypertensive drug therapy (27.8%). The prevalence of hypertension was 66% (N=279), but only 13.6% of hypertensive patients, as per the protocol, had their blood pressure controlled (N=37). Diabetes was present in 31.2% of

		Total Number of Patients	Number of Patients	Percentage	Lower Limit of Cl ^a	Higher Limit of Cl ^a
Hypertension	Per protocol	423	279	66.0%	61.3%	70.4%
	Known	273	123	45.1%	39.2%	51.0%
	Known and treated	123	99	80.5%	72.8%	86.7%
	Unknown and treated	150	17	11.3%	7.0%	17.1%
	Treated	273	116	42.5%	36.7%	48.4%
	Treated and controlled	116	30	25.9%	18.6%	34.4%
	Controlled	273	37	13.6%	9.9%	18.0%
Diabetes	Per protocol	420	131	31.2%	26.9%	35.7%
	Known	128	96	75.0%	67.0%	81.9%
	Known and treated	96	74	77.1%	68.0%	84.6%
	Unknown and treated	32	I	3.1%	0.3%	13.7%
	Treated	128	75	58.6%	49.9%	66.9%
	Treated and controlled	75	5	6.7%	2.6%	14.0%
	Controlled	128	8	6.3%	3.0%	11.4%
Dyslipidemia	Per protocol	263	103	39.2%	33.4%	45.2%
	Known	417	112	26.9%	22.8%	31.3%
	Treated		97	23.3%	19.4%	27.5%
Smoking	None	418	190	45.5%	40.7%	50.2%
	Current smokers]	209	50.0%	45.2%	54.8%
	Former smokers		19	4.5%	2.9%	6.9%

Table 5	Epidemiology	of	Conventional	Risk	Factors
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Abbreviation: ^aCl, confidence interval.

cases (N=131), and only 6.3% of diabetic patients, according to the protocol, had their diabetes controlled (N=8). The median levels of total cholesterol and triglycerides were 4.79 mmol/L (equivalent to 186.5 mg/dL) and 2.03 mmol/L (equivalent to 179.65 mg/dL), respectively. Based on our definition criteria, 103 patients were considered dyslipidemic (39.2%). Half of the patients were smokers (N=209, 50.0%). It is worth noting that 39 patients had previously experienced a cardiovascular event (9.4%).

The URF studied were the metabolic syndrome, socio-economic status, and family history (see Table 6 and Table 7). The prevalence of metabolic syndrome was 42.9% (N=115). The classification of socio-economic status, defined by the composite score, resulted in 182 patients with 1 point (42%) and 89 patients with 0 points (20.6%). The majority of participants had a family history of cardiovascular disease in their parents, children, or siblings (N=365, 87.5%), and 364 had a family history in either parents or children only (87.3%).

In our studied population, 67.2% of participants (N=163) were classified as being at high or very high cardiovascular risk (22.4%) (see Table 8). Correlations were examined between the SCORE and risk factors that were not included in the risk calculation. Diabetes and metabolic syndrome showed a strong positive relationship with the SCORE (p=0.000). This positive relationship was also observed with body mass index (BMI) (correlation coefficient CC = 0.179; p = 0.005). Socio-economic status did not show a significant correlation with cardiovascular risk (CC = -0.104; p = 0.059), and neither did family history of parents and children (p = 0.834) or family history including siblings (p = 0.975) (see Table 9).

		Total Number of Patients	Number of Patients	Percentage	Lower Limit of Cl ^a	Higher Limit of Cl ^a
Metabolic Syndrome		268	115	42.9%	37.1%	48.9%
Socio-Economic Status	0 point	433	89	20.6%	17.0%	24.5%
Classification	l point		182	42.0%	37.4%	46.7%
	2 points		125	28.9%	24.8%	33.3%
	3 points		37	8.5%	6.2%	11.5%
Family History	Father	417	185	44.4%	39.6%	49.2%
	Mother		232	55.6%	50.8%	60.4%
	Brother(s)		149	35.7%	31.2%	40.4%
	Sister(s)		102	24.5%	20.5%	28.8%
	Child(ren)	-	28	6.7%	4.6%	9.4%
	Parents' or children's cardiovascular events		364	87,3%	83,8%	90,2%
	Global familial cardiovascular events		365	87,5%	84,1%	90.4%

Table 6 Epidemiology of Unconventional Risk Factors

Abbreviation: ^aCl, confidence interval.

		Total Number of Patients	Mean	Standard Deviation	Median	Qlª	Q3 ^b
BMI ^c		419	28.04	4.78	27.41	24.69	30.63
Waist circumference	Female	171	96	12	95	87	104
	Male	125	104	12	104	96	111
Hip circumference	Female	170	98.9	14.2	100.0	89.0	109.0
	Male	125	105.0	10.8	106.0	98.0	111.0

Table 7 Anthropometric Measurements

Abbreviations: ^aQ1, first quartile; ^bQ3, third quartile; ^cBMI, body mass index.

Table 8 Cardiovascular Risk of the Rural Lebanese Population Studied

	Number of Patients	Percentage	Lower Limit of Cl ^a	Higher Limit of Cl ^a
Low	69	27.6%	22.3%	33.4%
Moderate	18	7.2%	4.5%	10.9%
High	107	42.8%	36.8%	49.0%
Very High	56	22.4%	17.6%	27.9%

Abbreviation: ^aIC, confidence interval.

		p-value			
Diabetes		0.000			
Metabolic Syndrome		0.000			
Familial History: parents and c	0.834				
Familial History: Parents, childi	Familial History: Parents, children and siblings				
Socio-Economic Status	Socio-Economic Status CC ^b				
	0.059				
BMI ^c	0.179	0.005			

Table 9 Correlations Between Risk Factors and SCORE^a

Abbreviations: ^aSCORE, Systematic Coronary Risk Estimation; ^bCC, correlation coefficient; ^cBMI, body mass index.

Discussion

CV diseases are the leading cause of morbidity and mortality worldwide, including in Lebanon.^{3,8} Given the lack of recent data in Lebanon, our study not only establishes an epidemiological database but also provides a roadmap for implementing a primary and secondary prevention strategy.

The SCORE is a tool that assesses the risk of fatal CV events, unlike the PCE, which calculates the total number of events (both fatal and non-fatal). The inclusion of non-fatal events is complicated due to varying definitions and evolving diagnostic methods, leading to confounding factors. Additionally, recalibrating the calculation models for total events is challenging for validation in each population. Efforts are underway to develop specific tables for different regions worldwide.⁶ In the absence of such tables, the risk calculation model for fatal CV events used in this study is based on that of high-risk European countries.

In our population, 65.2% of subjects have a high or very high risk of experiencing a CV event. This significant risk aligns with observations from daily medical practice, although recent publications lack a clear assessment. The World Health Organization (WHO) data from 2002 classified Lebanon as a country with a low risk of CV mortality.⁹ However, data published in 2018 indicates a CV mortality prevalence of 47%.⁵ Unfortunately, the WHO has not updated the risk status of the Lebanese population since 2002.

The pathogenesis of CV diseases is multifactorial, but the SCORE only considers age, sex, tobacco use, total cholesterol levels, and blood pressure. These factors are inherently interconnected, which is why they are not included in the correlation analyses.

The percentage of patients who experienced non-fatal CV events is low (9.4%), but both CRF and URF are common in our population. Limiting the study population to individuals aged between 40 and 70 years may explain this observation, as CV risk factors develop early in life, while their complications manifest later. The prevalence of cardiovascular diseases significantly increases with age in both men and women. Although advanced age exposes individuals to risk factors for a longer period, age itself is a major independent risk factor for cardiovascular disease.¹⁰ It's worth noting that women tend to develop cardiovascular diseases at an older age than men due to hormonal changes in postmenopausal women.^{11,12}

The prevalence of hypertension, diabetes, dyslipidemia, and smoking in our study is higher than the rates reported in a study conducted in the suburbs of Beirut (36.4%, 12.8%, 24%, and 43.1%, respectively). This difference can be attributed to the fact that our study defined CV risk factors based on patient-reported information rather than adhering strictly to protocol guidelines.¹³

Hypertension is the most significant risk factor for cardiac events,¹⁴ and it is highly prevalent in the rural Lebanese population, with 66% of participants being hypertensive. However, only 13.6% of hypertensive patients have controlled blood pressure. These percentages differ from those reported by Matar et al (36.9% and 54%, respectively).¹⁵ Factors such as the white coat effect, blood pressure measurement techniques, and limited access to healthcare in rural areas may

contribute to these variations. Blood pressure control remains a challenge, even in developed countries. For instance, the Health Study on the Environment, Biomonitoring, Physical Activity and Nutrition (ESTEBAN) conducted in metropolitan France estimated that 31.3% of the French population is hypertensive, of whom 50.0% are receiving antihypertensive drug therapy. Among hypertensive patients, only 24.3% have controlled blood pressure.¹⁶

Diabetes is also a major risk factor for CV events. In our population, the prevalence of diabetes is 31.2%, but only 6.3% of diabetic patients have their condition under control. However, a study published in 2019 reported that 30% of diabetics in Lebanon had their diabetes under control.¹⁷ Similar to hypertension, the lack of healthcare facilities, clinics, and physicians in rural areas may result in delayed diagnosis and inadequate management.

Dyslipidemia is present in 39.2% of participants. One noteworthy characteristic of the Lebanese population is the high prevalence of Familial Hypercholesterolemia, often referred to as the "Lebanese allele". This can be attributed to the heterozygosity of the gene and the high rates of consanguinity within the Lebanese population, particularly in rural areas.⁸ Additionally, educational level is inversely correlated with the risk of dyslipidemia.¹⁸ However, the study was unable to calculate the percentage of patients with controlled dyslipidemia since LDL-cholesterol, which is used for control assessment, was not measured.

According to WHO data, 33% of Lebanese individuals are smokers. In the rural population, smoking is the second most prevalent CRF, with a prevalence of 50%. When compared to smoking rates among men and women in Jordan, Syria, and Palestine, Lebanon ranks second among men and first among women in smoking prevalence.¹⁹ Smoking in Lebanon is more common among individuals with low economic status or educational level, and it is more prevalent in rural areas. Another notable aspect of the Lebanese population is the high prevalence of water-pipe smoking.⁸

The concept of URF emerged more than 30 years ago²⁰ and has gained importance due to advancements in research, diagnostics, and therapeutics. International recommendations emphasize the significance of considering URF without a specific decision-making algorithm.^{2,6} Metabolic syndrome is strongly associated with the occurrence of CV events and CV mortality.²¹ Its definition relies on several factors, including hypertension, diabetes, and hypertriglyceridemia, which explains its high prevalence and its correlation with the SCORE (as CRFs are correlated with the SCORE). Consequently, the CV risk associated with the metabolic syndrome is indirectly reflected in the SCORE calculation. The literature reports a metabolic syndrome prevalence of 20 to 30% in the general population.²²

Socio-economic status does not show a correlation with the SCORE. However, in Lebanon, individuals with low income and low educational level have a higher prevalence of CV diseases, independent of CV risk factors.²³ Therefore, socio-economic status should be considered as a risk factor independently of CV risk calculation.

Similarly, family history is not correlated with the SCORE. However, CV diseases have a significant genetic component, especially when there is a history of CV events in relatives.²⁴ Given the high rates of consanguinity, family history plays a notable role in CV risk. While not directly correlated with CV risk calculation, family history should be considered as an additional risk factor.

The two main limitations of this study are selection bias and recall bias. The non-random sampling method used in the study introduces selection error, as the sample was chosen from five Lebanese rural regions without proportional representation based on population size per district or governorate. Due to political uncertainties and the COVID-19 pandemic, expanding the study to other governorates was postponed. To minimize recall bias, particularly regarding drug intake, the questionnaire specified therapeutic classes.

Conclusion

Our study showed that diabetes and metabolic syndrome are correlated to SCORE and therefore indirectly reflected by it. The socio-economic status and the family history, as for them, are not correlated with the SCORE and put the affected population at risk; therefore, they should be included in the overall patient assessment in addition to the risk calculation. Moreover, CRF and URF are prevalent in our population but are poorly managed in the majority of cases. A national effort is required to ensure better control of the risk factors in the Lebanese population in general, and in the rural population in particular. Primary prevention should be the pillar of the strategy of the Ministry of Health in order to reduce the costs of the inevitable complications of CV risk factors. The involvement of non-medical staff could contribute to improving care and better patient compliance.²⁵

Disclosure

The authors report no conflicts of interest in this work.

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