QRISK2 Score in CABG Patients Correlated with Risk Factors

AMIN BAZYANI^{3,4}, RAZAN AL NAMAT¹*, MAURA GABRIELA FELEA¹*, IRINA IULIANA COSTACHE¹*, MIHAI CONSTANTIN², VICTORITA SORODOC², LAURENTIU SORODOC², PAUL SIMION^{3,4}, MIRELA MIHAELA MIHALCIA¹, FLORIN MITU¹, GRIGORE TINICA^{3,4}

¹Grigore T. Popa University of Medicine and Pharmacy, Faculty of Medicine, 1st Medical Department, 16 Universitatii Str., 700115, Iasi, Romania

²Grigore T. Popa University of Medicine and Pharmacy, Faculty of Medicine, 3rd Medical Department, 16 Universitatii Str., 700115, Iasi, Romania

³Grigore T. Popa University of Medicine and Pharmacy, 16 Universitatii Str., 700115, Iasi, Romania ⁴Institute for Cardiovascular Diseases, 50 Carol I Blvd., 700503, Iasi, Romania

Guidelines for primary prevention suggested using any risk score, among those QRISK2, identifying the high-risk populations. The purpose of this study was to determine whether the QRISK2 Score would register changes in patients with coronary artery disease demanding acute or postponed CABG intervention. The QRISK2 Score was performed the day of admission after the clinical examination and blood test results, and immediately after CABG surgery (in the first week post-CABG, in an interval of 24 hours to 7 days) having another blood test evaluation. The 120 patients admitted in the Clinic of Cardiovascular Surgery of the Institute of Cardiovascular Disease met the inclusion criteria: CABG patients (less than 1 week), aged 40-85 years old, BMI > 25 kg/m², and mixed dyslipidemia. In both phases, for every patient, it was performed a clinical examination, a set of hematological, biochemical, lipid, coagulation and inflammatory profile, and ECG and echocardiography. Our research on hospitalized patients undergoing CABG, by comparing the Phase I and Phase III results, revealed that the median 10-year QRISK2 score gives a more appropriate risk estimation based on the social component, thus identifying high risk patients associating social deprivation. Comparative to Framingham risk score, QRISK2 score, by including additional variables, proves the efficacy of lifestyle changes and management decisions, and sustaines the treatment directed towards modifying variables or risk factors.

Keywords: QRISK2 score, coronary artery disease, CABG, cardiovascular disease prevention

According to the European Guide 2016 on cardiovascular disease prevention, the benefit of simple risk scores was to predict long-term morbidity and subsequent mortality: Framingham Risk Score (FRS), ASSIGN, QRISK1 & QRISK2, PROCAM. The most common clinical tool, used to estimate the risk level of coronary artery disease (CAD), to identify men and women at high risk susceptible to changing risk factors in order to prevent future cardiovascular events, is the Framingham Risk Score (FRS). FRS components comprise gender, age, smoking status, systolic blood pressure and lipid profile. FRS can indicate the possible benefits of prevention, being useful for the patient and clinician to choose lifestyle changes and/or preventive medical treatment [1].

A large number of risk assessment scores have been developed to help clinicians determine the long-term risk of cardiovascular diseases (CVD). In primary prevention, the Framingham, ASSIGN and QRISK2 scores are widely used to predict CVD risk at 10 years. The Framingham risk score is based on an American cohort a few decades ago. The ASSIGN risk score was derived from the Scottish Extended Health School and the QRISK risk score from a large primary care database in England and Wales. These scores were based on risk factors that can be easily assessed and measured in the general population. Framingham, ASSIGN and QRISK2 risk scores were validated by comparison with the predicted risks in the total population [2]. There is no consensus on which risk score to be used for CVD risk assessment. Guidelines for primary prevention suggested using any risk score [3]. These three risk scores are currently used in the UK to determine the risk of CVD. Two validation studies for QRISK2 reported that the predicted and observed risks were on average similar and concluded that QRISK2 was accurate in identifying a high-risk population [4, 5].

In 1968, Rene Favaloro introduced the Coronary Artery Bypass Grafting (CABG) as the first technique for myocardial revascularization, which suffered some methodological changes later on [6]. Another change was the indication of CABG for particular groups of patients, such as patients with more complex coronary anatomy, defined by a Syntax score greater than 22, and patients presenting comorbidities like diabetes mellitus and chronic kidney disease in stage 4 and 5 [7, 8].

Experimental part

Material and methods

The 120 individuals comprised in the study group were admitted in the Clinic of Cardiovascular Surgery of the Institute for Cardiovascular Diseases.

All subjects who participated in the research signed in the informed consent, certifying that they agree with the investigations, blood tests and treatment mandatory to be performed, as well as with the final publication of the data in scientific form and under permanent protection of anonymity. The study was endorsed by the Research Ethics Committee of the Grigore T. Popa University of Medicine and Pharmacy of Iasi.

The purpose of this study was to determine whether the QRISK2 Score would register changes in patients with coronary artery disease demanding acute or postponed CABG intervention. The QRISK2 Score was performed the

* email: dr.razan_romania@yahoo.com; feleamag@yahoo.com irinaiulianacostache@yahoo.com day of admission after the clinical examination and blood test results, and immediately after CABG surgery (in the first week post-CABG, in an interval of 24 hours to 7 days) having another blood test evaluation.

Statistical analysis

The database was compiled in Microsoft Office Excel 2010 version, and statistical analysis was performed in the IBM SPSS Statistics v.20. We computed the averages, frequencies, standard deviations, differences between the maximum and minimum values of the numerical parameters.

The statistical significance of the difference between two frequencies was determined by the Chi-square test of independence. The t-Student test was used to reveal the significance of the difference between two average values. The threshold values for p were considered < 0.05, providing a statistical significance level of the test. The regression equations and correlation coefficients were also calculated.

Results and discussions

The 120 patients admitted in the Clinic of Cardiovascular Surgery of the Institute of Cardiovascular Disease met the inclusion criteria: CABG patients (less than 1 week), aged 40-85 years old, $BMI > 25 \text{ kg/m}^2$, and mixed dyslipidemia.

In both phases, for every patient, it was performed a clinical examination, a set of hematological, biochemical, lipid, coagulation and inflammatory profile, and ECG and echocardiography.

Blood pressure and heart rate were monitored daily in the Intensive Care Unit, and then twice a day.

Glucose plasma level was monitored once daily and more frequent in case of diabetic patients. Renal function was evaluated by serum urea, creatinine and uric acid level. Creatinine (2-Amino-1-methyl-5H-imidazol-4- one), an important indicator of renal function, is byproduct of the muscle metabolism that is excreted unchanged. Glomerular filtration rate (GFR) was calculated according to the most accurate formula of CKDEPI (Chronic Kidney Disease Epidemiology Collaboration) [9]:

Disease Epidemiology Collaboration) [9]: $eGFR = 141 \text{ x min}(SCr/k,1)\alpha \text{ x max}(\alpha Cr/k,1)-1.209 \text{ x}$ 0.993Age x [1.018 if Female]

[SCr = serum creatinine (mg/dL); k = 0.7 for females and 0.9 for males; $\alpha = -0.329$ for females and -0.411 for males; min = the minimum of SCr/k or 1, and max = the maximum of SCr/k or 1).

QRISK®2 is a well established cardiovascular disease (CVD) risk score used in the NHS since 2009, that is designed to identify people at high risk of developing CVD, who need to be more thoroughly assessed to reduce their risk of developing CVD [10,11].

The QRISK[®]2 score, estimating the risk of a person to develop CVD over the next 10 years, was specifically developed by physicians and academics to be used in the UK. In the British Medical Journal in July 2007 and in the journal Heart in January 2008, it was published the original research that substantiates QRISK score [10,11]. The **QRISK2** score includes age, gender, ethnicity, smoking status, and presence/ absence of diabetes, chronic kidney disease (stage 4 or 5), atrial fibrillation, antihypertensive treatment, rheumatoid arthritis, and family history of angina pectoris or myocardial infarction in a first degree relative younger than 60 years old; laboratory item includes ratio of total serum cholesterol to high density lipoprotein (HDL)cholesterol ratio; the functional items include the values for systolic blood pressure, height and weight, thus alowing to calculate the body mass index (BMI). The laboratory and functional parameters are more variable and susceptible to changes.

A score is assigned to each level of every risk factor, and the total score is calculated by adding all the points. The

- About you
Age (25-84): 64
Sex: Male Female
Ethnicity: White or not stated v
UK postcode: leave blank if unknown
Postcode:
Clinical information
Smoking status: non-smoker
Diabetes status: none 🔻
Angina or heart attack in a 1st degree relative < 60?
Chronic kidney disease (stage 4 or 5)?
Atrial fibrillation?
On blood pressure treatment?
Rheumatoid arthritis?
Leave blank if unknown
Cholesterol/HDL ratio:
Systolic blood pressure (mmHg):
Body mass index-
Height (cm):
Weight (kg):
Calculate risk over 10 Vears. Calculate risk
Calculate fisk over 10 • years. Calculate fisk

Fig. 1. The QRISK2 Score (https://qrisk.org/ 2016/)

sum of these points provides cardiovascular risk assessment, estimated over the next 10 years.

If someone has a 10 years -QRISK®2 score of 20%, then, out of a group of 100 people like these, on average, 20 people would develop cardiovascular disease over the next 10 years, or else they have one chance of five to be affected by one or more CVD over the next 10 years.

Statistical analysis

The database was compiled in Microsoft Office Excel 2010 version, and statistical analysis was performed in the IBM SPSS Statistics v.20. We computed the averages, frequencies, standard deviations, differences between the maximum and minimum values of the numerical parameters. The statistical significance of the difference between two frequencies was determined by the Chi-square test of independence. The t-Student test was used to reveal the significance of the difference between two average values. The threshold values for p were considered < 0.05, providing a statistical significance level of the test. The regression equations and correlation coefficients were also calculated.

The patients included in the group were aged between 41 and 85 years. The group characteristics at the admission, Phase I, before the CABG was performed, are detailed in table 1.

In Phase III, patients' characteristics (table 2) suffered significant changes, with amelioration of blood pressure control, lipid profile and body mass index. It is to be mentioned that those who were smokers at the admission, ceased to smoke after the cardiac surgery.

Our research on hospitalized patients undergoing CABG, by comparing the Phase I and Phase III results, revealed that the median 10-year QRISK2 cardiovascular risk score was approximately 47.88 % lower (p=0.000) in the first week after cardiac surgery (fig. 2).



Fig. 2. Box-plot Diagram – Patients distribution on the QRISK2 cardiovascular score in Phase I and III after CABG

The mean risk score value (50.7) in Phase I was similar to the median value (49.5), the same concordance being revealed also for Phase III risk scores (28.05 vs. 25.08) (table 3). The difference between the phases, as shown in the box-plot above is about 50%.

The mean and median changes of QRISK2 was more important in patients previously diagnosed with type 2 Diabetes Mellitus, or in patients with chronic kidney disease (from stage 2 to stages 4 and 5) (table 4).

QRISKŽ Score indirectly revealed another major change that is not previewed in the score components: the glomerular filtration function (fig. 3).

The study age group was ranged between 40 to 85 years old. It was considered the optimal range because cardiovascular risk in this age groups should be determined steadily, and possible further investigations are needed to evaluate true vascular age.

Framingham Score was the early well documented strategy to identify cardiac and cardiovascular risk factors,

Table 1

CHARACTERISTICS OF PATIENTS UNDERGOING CABG, AGED 41 TO 85 YEARS, AND ELIGIBLE FOR THE STUD

Characteristics	Women n= 29	Men n= 91
Mean (SD) age (years)	(24.17%) 67.34 (8.67)	(75.83%) 65.47 (10.18)
Mean (SD) body mass index (kg/m ²) – Phase I	28.76 (4.40)	29.63 (2.43)
Mean (SD) systolic blood pressure (mm Hg) – Phase I	151.03 (6.18)	157.42 (6.72)
Mean (SD) diastolic blood pressure (mm Hg) – Phase I	85 (6.81)	90.71 (6.26)
Mean (SD) total cholesterol to HDL cholesterol ratio – Phase I	5.96 (3.79)	5.25 (2.72)
Mean (SD) total cholesterol – Phase I	193.8 (61.04)	177.64 (40.24)
Mean (SD) HDL cholesterol – Phase I	38 (10.65)	40.89 (23.68)
Mean (SD) LDL cholesterol – Phase I	147.86 (19.98)	143.11 (33.43)
Mean (SD) TG – Phase I	142.76 (61.55)	149.60 (55.66)
Smoking status:		
Non-smoker	19 (15.83%)	28 (23.33%)
Current smoker (cigarettes/day):		
Light (<10)	10 (8.33%)	0
Heavy (≥20)	0	63 (52.5%)
Ethnic group: White	120 (100%)	120 (100%)

Table 2

CHARACTERISTICS	OF	PATIENTS'	PARAMETERS	ONE-WEEK	AFTER	CABG

Characteristics	Women n= 29	Men n= 91
Characteristics	(24.17%)	(75.83%)
Mean (SD) age (years)	67.34 (8.67)	65.47 (10.18)
Mean (SD) body mass index (kg/m ²) – Phase III	27.66 (4.61)	28.33 (0.03)
Mean (SD) systolic blood pressure (mm Hg) – Phase III	128.28 (6.98)	130.49 (5.17)
Mean (SD) diastolic blood pressure (mm Hg) – Phase III	73.62 (5.33)	76.43 (8.31)
Mean (SD) total cholesterol to HDL cholesterol ratio - Phase III	3.77 (1.64)	4.33 (3.82)
Mean (SD) total cholesterol – Phase III	175.76 (47.56)	165.66 (49.28)
Mean (SD) HDL cholesterol – Phase III	51.23 (14.39)	50.04 (28.19)
Mean (SD) LDL cholesterol – Phase III	126.48 (22.72)	120.60 (30.52)
Mean (SD) TG – Phase III	143.41 (57.60)	129.20 (56.68)

Table 3
DESCRIPTIVE STATISTICS OF QRISK2 CHANGES BETWEEN PHASE I AND PHASE III

Descriptive		Statistic							
statistics	Mean	95% Confidence Interval for Mean		Interval for Mean		Std. Deviation	Minimum	Maximum	
		Lower Bound	Upper Bound						
QRISK2-I	50.70	46.42	54.98	49.50	120	23.67	1.50	121.80	
QRISK2-III	28.05	25.16	30.93	25.80	120	15.96	2.00	67.00	

Table 4

DESCRIPTIVE STATISTICS OF QRISK2 CHANGES IN PATIENTS WITH TYPE 2 DIABETES MELLITUS AND IN PATIENTS WITH CKD

		Descriptive Statistics						
				nfidence for Mean				
		Mean	Lower Bound	Upper Bound	Median	Std. Deviation	Minimum	Maximum
Dif. QRISK2 III - I	No DM	-20.01	-24.01	-16.01	-18.40	14.81	-101.40	4.80
	T2DM	-24.89	-29.31	-20.47	-21.30	17.85	-66.70	6.00
Dif. QRISK2 III - I	No CKD	-20.89	-24.68	-17.10	-16.00	17.57	-101.40	4.80
	CKD	-26.94	-31.54	-22.33	-25.80	13.40	-66.70	6.00



distribution on the QRISK2 cardiovascular score in Phase I and III after CABG

		-	
	SCORE	FRS	QRISK2
Age			
Sex			
Ethnicity			
Systolic Blood Pressure			
Total Cholesterol			
HDL Cholesterol			
Total Chol to HDL-Chol Ratio			
Smoking Status			
Diabetes			
Antihypertensive Treatment			
Family History of CVD			
Chronic Kidney Disease (stage 4 or 5)			
Atrial Fibrillation			
Rheumatoid Arthritis			
Weight, Height, Body Mass Index			
High Sensitivity C-Reactive Protein			

Table 5 ITEMS INCLUDED IN SCORE, FRS AND QRISK2

thus estimating and stratifying the 10-year cardiovascular

risk of an individual to develop coronary heart disease. By comparison, QRISK2 Score includes more components than FRS and SCORE (table 5).

Along the years of medical research, a great number of diagnostic tools, as coronary angiography, and methods were created and used on patients suffering from coronary artery disease (CAD) [12]

Although it is costly and invasive, coronary angiography remains the gold standard for diagnosing the acute form or the chronic but significant coronary obstructive disease, and pointing out toward the optimal treatment choice.

As medical and surgical revolution in cardiovascular disease showed us along the years, and because of the multiple risk factors for CAD, several risk assessment tools may be used to try and estimate the risk of this pathology within the different age groups [13].

CABG proved to be effective in reducing the value of different parameters reflecting cardiac ischemia, and also in ameliorating the cardiovascular risk scores [14,15].

Conclusions

QRISK2 score gives a more appropriate risk estimation based on the social component, thus identifying high risk patients associating social deprivation. Comparative to Framingham risk score, QRISK2 score, by including additional variables, proves the efficacy of lifestyle changes and management decisions, and sustaines the treatment directed towards modifying variables or risk factors.

Limitations

Encountered limitations reffered to the follow-up of the study group for more than to years, in order to review the QRISK2 Score evolution, and to stratify the patients' compliance to lifestyle changes and medical treatment.

References

1.YOUSEFZADEH, G., SHOKOOHI, M., NAJAFIPOUR, H., et al. ARYA Atheroscler, 2015, 11(3), p. 179-185.

2.SIONTIS, G.C.M., TZOULAKI, I., IOANNIDIS, J.P.A., BMJ, 2012, 3318: 1-11.

3.DAVIES, M., KHUNTI, K., WEBB, D., et al. Updated. The handbook for vascular risk assessment, risk reduction and risk management. Leicester, 2012.

4.COLLINS, G.S., ALTMAN, D.G., BMJ, 2012, 4181: 1-12.

5.VAN STAA, T.P., GULLIFORD, M., NG, E.S., et al. PLoS One, 2014, 9(10): 106455

6.HU, S., LI, Q., GAO, P., et al. Ann Thorac Surg, 2011, 91, p. 432-438. 7.DEB, S., WIJEYSUNDERA, H.C., KO, D.T., et al., JAMA, 2013, 310, p. 2086-2095.

8.WINDECKER, S., KOLH, P., ALFONSO, F., et al. Eur Heart J, 2014, 35, p. 2541–2619.

9.LEVEY, A.S., STEVENS, L.A., SCHMID, C.H., ZHANG, Y.L., CASTRO, A.F., FELDMAN, H.I., KUSEK, J.W., EGGERS, P., VAN LENTE, F., GREENE, T., CORESH, J., Ann. Intern. Med., 150, no. 9, 2009, p. 604–612.

10.HIPPISLEY-COX, J., COUPLAND, C., VINOGRADOVA, Y., ROBSON,

J., MAY, M., BRINDLE, P., et al., BMJ 2007; 335:136.

11.HIPPISLEY-COX, J., COUPLAND, C., VINOGRADOVA Y., et al., Heart 2008;**94**:34-39

12.JENSEN, J.M., VOSS, M., HANSEN, V.B., ANDERSEN, L.K., JOHANSEN, P.B., et al., Atherosclerosis, 2012, 220: 557-562.

13.GRIMA, K.B., BEZZINA, P., RAINFORD, L. J Clin Diagn Res, 2017, 5:135. doi: 10.4172/2376-0311.1000135

14.AL NAMAT, R., AURSULESEI, V., FELEA, M.G. et al. Rev. Chim. (Bucharest), **68**, no. 6, 2017, p. 1485-1489.

15.COSTACHE, I.I., AL NAMAT, R., MITU, F., CIOCOIU, M., AURSULESEI, V., MITU, O., COSTACHE, A.D., MARCU, D., BUBURUZ, A.M., Rev. Chim. (Bucharest), **68**, no. 12, 2017, p. 2967.

Manuscript received: 7.10.2018