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The Impact of Financial Crisis in Coronary Artery Disease Burden in Greece

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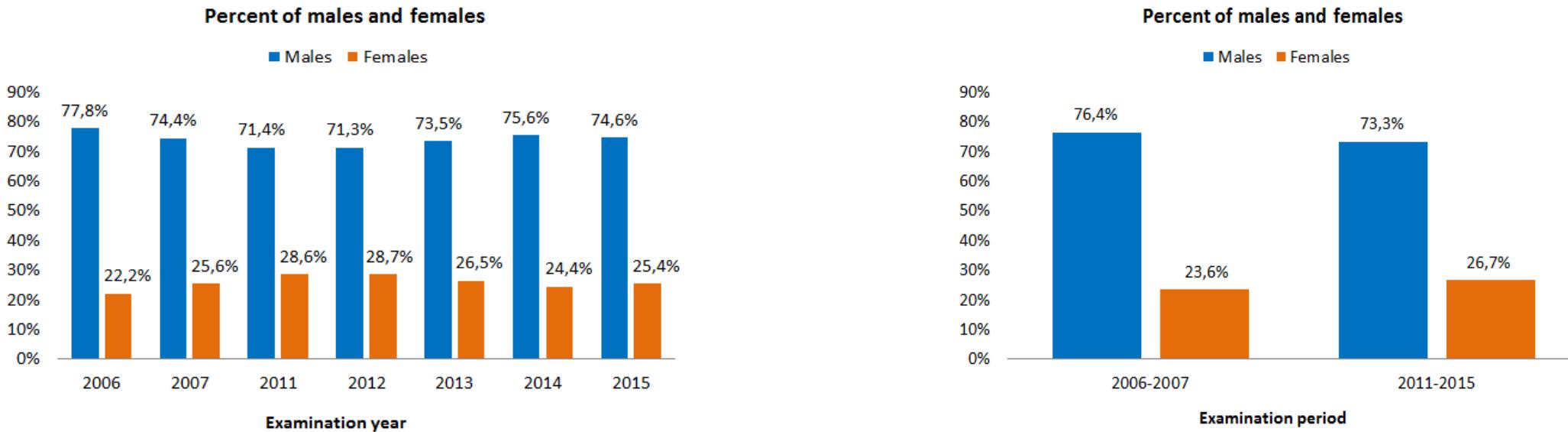
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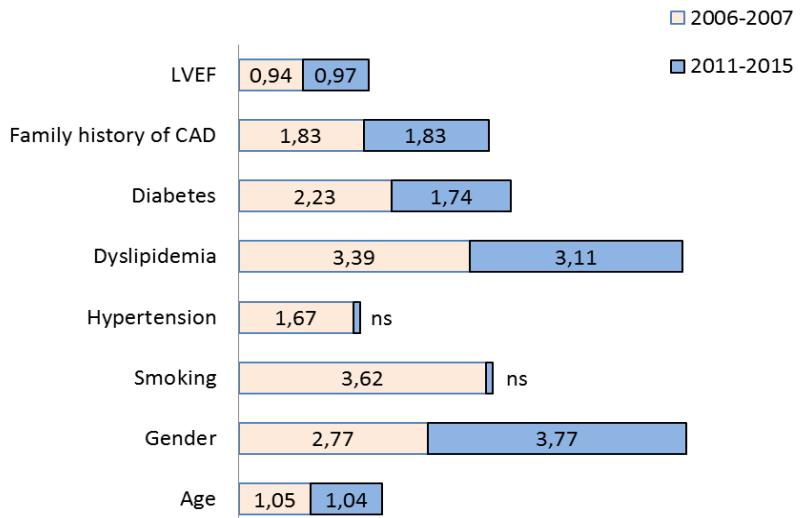
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A statistically significant increase in the percent of examined females was observed during the period 2011-2015 compared to 2006-2007 (26.7% vs 23.6%, p=0.038).



Odds ratios of independent risk factors associated with positive coronary artery disease findings for the two periods 2006-2007 and 2011-2015.

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2 **The Impact of Financial Crisis in Coronary Artery Disease Burden in Greece**

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Abstract

26 **Background.** Economic crisis poses an immense threat to public health worldwide
27 and has been linked to cardiovascular morbidity and mortality. Greece is facing a
28 distinctive recession over the recent years. However, the exact impact on coronary
29 artery disease (CAD) burden has not been adequately addressed.

30 **Methods.** Demographic, clinical and angiographic data of 3895 hospitalized patients
31 were retrospectively studied. Patients were classified in those before crisis (2006-
32 2007, n=1228) and those during crisis (2011-2015, n=2667).

33 **Results.** All data before and during crisis were compared. During crisis, patients
34 presented with less acute coronary syndromes (ACS - 45.5% vs 39.9%, p<0.001).
35 Subsequently, there were more patients without CAD (23.7% vs 35.1%, p<0.001) or
36 one vessel disease (20.5% vs 23%, p<0.001). The prevalence of traditional risk
37 factors decreased significantly or remained stable except obesity (26.3% vs 31.4%,
38 p=0.002). A significant increase of the examined females (23.6% vs 26.7%, p=0.04)
39 was also observed.

40 **Conclusions.** The extend of CAD during financial crisis was partially affected. Even
41 though the incidence of ACS was decreased, more women and more patients with
42 no or single vessel disease were led for cardiac catheterization. In addition, the
43 prevalence of traditional risk factors for CAD did not increase except obesity
44 confirming the “obesity paradox”. It seems that the impact of traditional risk factors
45 for CAD is not an immediate process and is somewhat related to living conditions or
46 other exogenous and social factors.

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50 **Introduction**

51 Financial crisis poses an immense threat to public health worldwide and has
52 been linked to cardiovascular morbidity and mortality [1-3]. Greece is facing a
53 distinctive recession over the last decade. However, the exact impact on coronary
54 artery disease (CAD) burden along with the possible negative health outcomes have
55 not been adequately addressed yet [4].

56 Historically, the major risk factors for cardiovascular disease include
57 advancing age, male gender, positive family history for premature CAD, smoking,
58 hypertension, lipid abnormalities, glucose metabolism disorders and obesity [5, 6].
59 These factors cluster and interact multiplicatively to promote vascular risk [7]. In
60 addition, the specific role of each factor has been incorporated in the development of
61 several multivariable risk prediction algorithms conduced to assess the risk of
62 disease in individual patients [8, 9]. Knowing the exact profile of each catheterized
63 patient (the number and the type of vessels diseased) we can obtain substantial
64 prognostic information for the outcome of CAD. Thus, it is of great importance to
65 determine the exact relationship between the clinical profile and the angiographic
66 characteristics [10].

67 The principal aim of the current study was to investigate the imminent impact
68 of economic crisis on patients who underwent diagnostic cardiac catheterization and
69 to observe any potential changes in the trends and the severity of the established
70 cardiovascular risk factors.

71

72 **Methods**

73 **Study Population.** Within two distinct periods of time namely 2006 to 2007
74 and from 2011 to 2015, 3895 patients in total were subjected to heart catheterization

75 due to typical or atypical clinical symptoms according to the records of Laiko General
76 Hospital of Athens, Greece and were classified in two groups, the group before
77 financial crisis (2006-2007, n=1228) and the group during crisis (2011-2015,
78 n=2667). Angiographic evaluation included a mandatory diagnostic coronary
79 angiography according to standard techniques for all incoming patients and, if
80 necessary, a percutaneous coronary intervention (PCI). Before examination a
81 detailed coronary risk profile for every patient was obtained as well as valid height
82 and weight measurements. The present study was approved by the Medical
83 Research Ethics Committee of our Institution and was carried out in accordance with
84 the declaration of Helsinki of the World Medical Association.

85 **Definitions.** Family history for premature CAD was considered positive when
86 there was a history of coronary manifestation in first-degree relatives before the age
87 of 55 for males and before the age of 65 for females. Dyslipidemia was defined by
88 total cholesterol levels >200 mg/dL and/or low-density lipoprotein cholesterol levels
89 >130 mg/dL and/or high-density lipoprotein cholesterol levels <40 mg/dL for men and
90 <50 mg/dL for women and/or triglyceride levels >150 mg/dL or current use of anti-
91 dyslipidemic agents. Patients that took antihypertensive medications or their blood
92 pressure levels were ≥140/90 mmHg were defined as hypertensive. Diabetes
93 mellitus (DM) was defined by fasting plasma glucose levels ≥126 mg/dL or 2-hour
94 values in the oral glucose tolerance test ≥200 mg/dL or levels of isolated elevation of
95 glycated hemoglobin ≥6.5% or current use of insulin or other oral hypoglycemic
96 agents. Current smokers were defined those that smoked at least one cigarette per
97 day and non-smokers those who did not smoke and/or had quit smoking for at least
98 one year. Body mass index (BMI) was calculated as bodyweight in kilograms divided

99 by height in meters squared and was estimated for all patients. Obesity was defined
100 as BMI $\geq 30 \text{ kg/m}^2$.

101 **Statistical analysis.** Continuous variables were expressed as mean values \pm
102 standard deviation (SD). Comparisons of continuous variables were performed by
103 one-way analysis of variance (ANOVA). Categorical data were expressed as
104 absolute and relative frequencies. Multivariable logistic regression analysis was
105 performed to examine the predictive value of the risk factors that are independently
106 related to the presence of CAD. All data were analyzed using Statistical Package for
107 the Social Sciences (SPSS) version 18.0 and all tests were 2-tailed with the 5%
108 indicating level of significance.

109

110 **Results**

111 **Demographic and clinical characteristics.** A total of 3895 patients were
112 angiographically evaluated for clinically suspected CAD. During crisis, less patients
113 with ACS (45.5% vs 39.9%, $p<0.001$) were presented compared to the pre-crisis
114 period. Even though the predominance of men over women was verified in both of
115 the examined periods (76.4% vs 73.3%, $p=0.04$) there was a significant increase of
116 the examined females (23.6% vs 26.7%, $p=0.04$) during crisis. Risk factors including
117 family history of CAD (17.7% vs 13%, $p<0.001$), smoking (45.4% vs 36.9%,
118 $p<0.001$), hypertension (69.2% vs 60%, $p<0.001$) and dyslipidemia (59% vs 48.1%,
119 $p<0.001$) were significantly decreased while the incidence of DM remained
120 practically stable (28.4% vs 27.2%, $p=0.44$). Conversely, only obesity was
121 considerably more prevalent (26.3% vs 31.4%, $p=0.002$). Data are presented in
122 Table 1.

123 Comparing patients with CAD before and during crisis a significantly lower
124 prevalence of all risk factors except DM was observed. Hypertension and
125 dyslipidemia were also reduced in those without CAD, but the differences were not
126 statistically significant. Nevertheless, only obesity was significantly increased in all
127 patients with or without CAD during economic downturn (Table 2).

128 **Angiographic characteristics.** The number of patients without CAD (23.7%
129 vs 35.1%, p<0.001) was increased during financial crisis. Furthermore, more
130 individuals with one vessel disease (20.5% vs 23%, p<0.001) and less with two
131 (20.2% vs 18.9%, p<0.001) or three vessels disease (35.6% vs 23%, p<0.001) were
132 examined (Table 3).

133 **Multivariable Analysis.** According to the multiple logistic regression
134 analysis, male gender (OR=3.77, CI=2.68-5.30), dyslipidemia (OR=3.11, CI=2.26-
135 4.28), family history (OR=1.83, CI=1.16-2.89) and DM (OR=1.74, CI=1.21-2.51) were
136 significantly associated with CAD during financial crisis. Before crisis, this correlation
137 was slightly different with smoking (OR=3.62, CI=2.51-5.21) to be considered as the
138 strongest predictive factor (Table 4).

139

140 **Discussion**

141 This study showed that the burden of CAD in Greece during financial crisis
142 was partially affected. Even though the incidence of ACS was decreased, more
143 women and more patients with no or single vessel disease were led for cardiac
144 catheterization. Moreover, the prevalence of traditional risk factors for CAD was
145 reduced or remained stable except obesity confirming the “obesity paradox”.

146 Lower socioeconomic status leads to higher rates of morbidity and mortality
147 [11, 12]. This relationship is particularly evident in the case of CAD [13]. A cross-

148 sectional study that was conducted to explore the current economic status of
149 patients with CAD revealed that most of the patients categorized in low income level
150 [14]. In addition, the PURE study that included citizens of 17 countries showed that
151 the rates of ACS were substantially higher in low income countries [15]. The results
152 of the present study suggest that the prolonged financial downturn might have led to
153 fewer patients with ACS. Furthermore, there were more patients without CAD or with
154 one vessel CAD whereas the incidence of complex CAD was reduced, presumably
155 because the majority of individuals presented with atypical symptoms or stable CAD
156 during crisis. Nevertheless, these patients were subjected to earlier diagnostic
157 catheterization and, if necessary, PCI.

158 Although CAD is mainly considered as men's disease, more women die
159 annually mainly due to the specific pathology differences of atherosclerosis [16, 17].
160 Females have a lower burden of obstructive CAD and a higher prevalence of angina
161 [18]. They are also two to three times more likely than men to experience a cardiac
162 event and have persistent chest pain symptoms within the first year following cardiac
163 catheterization [19]. Additionally, females with CAD are often less symptomatic or
164 have atypical symptoms emphasizing the need for gender-specific approaches to
165 CAD evaluation [20, 21]. Low socioeconomic status might also contribute to negative
166 health outcomes for women. Indeed, women with low socioeconomic status are at
167 significant greater risk of CAD compared with men according to a recent meta-
168 analysis [22]. In the current study, there was a significant increase of the examined
169 females during economic downturn compared with the previous period of time.
170 Consequently, the increasing awareness among physicians and general population
171 about clinical characteristics of female individuals with CAD might explain this
172 observation. To the best of our knowledge this is the first study in Greek population

173 that revealed an increase in the percentage of women that was evaluated for
174 clinically suspected CAD.

175 It is well established that the higher the BMI, the greater the risk of
176 cardiovascular diseases [23, 24]. However, a lot of studies have underlined the
177 paradoxical relationship between obesity and cardiovascular prognostic outcomes in
178 large number of patients [25, 26]. Lancefield et al in 2010 showed that overweight
179 and obese patients had a lower in-hospital and 1-year mortality rate after PCI
180 compared to normal weight patients [27]. The term used to describe this relationship
181 is “obesity paradox”. In this study there was a decline in the prevalence of all
182 traditional risk factors but the prevalence of obesity was increased. The increase in
183 obesity combined with the decrease of the total number of cases with CAD during
184 crisis could be a direct confirmation of the “obesity paradox”.

185 Tobacco use has been found to be reduced during period of economic
186 downturn [28-30]. Our results seem to support this finding. Moreover, smoking was
187 considered as the strongest predictive factor for CAD before financial crisis, but not
188 during crisis. Hence, it seems that economic downturn along with tobacco control
189 legislations that were enforced since 2010 in Greece led people to make healthier
190 lifestyle choices such as smoking cessation.

191 Last but not least, DM is associated with two to four-fold increased risk for
192 multivessel CAD and worse overall long-term prognosis even though after PCI [31-
193 34]. Moreover, there is also evidence that low income is related to higher incidence
194 of DM [35, 36]. Nevertheless, in the current study the prevalence of diabetes
195 remained stable during financial crisis. This finding might be consistent with the
196 decreased percentage of patients with multivessel CAD. Poor nutrition and

197 decreased adherence to medication along with poor monitoring of diabetic
198 complications during crisis could explain the stable prevalence of DM.

199

200 **Limitations**

201 The present study had several limitations. First, it was a retrospective single center
202 study including patients who underwent diagnostic cardiac catheterization. Our cath
203 lab is located in Athens and the patients admitted to our clinic were mostly Greek
204 natives with similar economic and social status. Second, the study population was
205 not propensity matched. Third, there were no follow up data.

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207 **Table 1.** Demographic and clinical characteristics of study population before and
 208 during financial crisis.

	Total population (n=3895)	Before crisis 2006 - 2007 (n=1228)	During crisis 2011 - 2015 (n=2667)	p-value
Age (yrs)	63.8±11.3	63.8±10.6	63.9±11.6	0.78
Males (%)	74.2	76.4	73.3	0.04
Females (%)	25.8	23.6	26.7	0.04
Weight (kg)	81.1±15.5	80.3±13.7	81.6±16.5	0.01
Height (cm)	169.5±8.5	169.2±8.4	169.7±8.5	0.08
BMI (kg/m ²)	28.1±4.6	27.9±4.0	28.2±4.9	0.04
Risk factors				
Family history (%)	14.5	17.7	13	<0.001
Smoking (%)	39.6	45.4	36.9	<0.001
Hypertension (%)	62.9	69.2	60	<0.001
Dyslipidemia (%)	51.5	59	48.1	<0.001
Diabetes mellitus (%)	27.6	28.4	27.2	0.44
Obesity (%)	29.5	26.3	31.4	0.002
ACS (%)	41.6	45.5	39.9	<0.001

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210 ACS: acute coronary syndrome, BMI: body mass index

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227 **Table 2.** Demographic and clinical characteristics of CAD-positive and CAD-negative
 228 patients before and during financial crisis.

Risk factors	Before crisis 2006 - 2007 (n=1228)		During crisis 2011 - 2015 (n=2667)		p-values * (p ₁ , p ₂)
	CAD (-)	CAD (+)	CAD (-)	CAD (+)	
Age (yrs)	61.7±11.6	64.4±10.2 [†]	61.8±12.0	65.0±12.2 [†]	0.9, 0.2
Gender (%), males	61.9	80.9 [†]	60.4	80.2 [†]	0.7, 0.6
BMI (kg/m ²)	28.0±4.3	27.9±3.9	28.5±5.2	28.1±4.7	0.01, 0.3
Family history (%)	14.4	18.7	12.8	13.1	0.5, <0.001
Smoking (%)	31.6	49.6 [†]	32.8	39.1 [†]	0.7, <0.001
Hypertension (%)	63.6	71.0 [†]	58.1	61	0.09, <0.001
Dyslipidemia (%)	39.9	64.9	36.2	54.6 [†]	0.3, <0.001
Diabetes mellitus (%)	18.2	31.6	20.5	30.9 [†]	0.4, 0.7
Obesity (%)	25.4	26.6	33.6	30.4	0.01, 0.05

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230 BMI: body mass index, CAD: coronary artery disease

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232 [†] significant difference (p<0.01) between positive versus negative CAD

233 * p-values (p₁, p₂) indicate significant (p<0.01) differences for subjects without CAD

234 (-) and those with CAD (+), between the two periods (2006-2007 vs 2011-2015),

235 respectively.

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242 **Table 3.** Angiographic characteristics of study population before and during financial
 243 crisis.

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	Before crisis 2006 - 2007 (n=1228)	During crisis 2011 - 2015 (n=2667)	p-value
Without CAD (%)	23.7	35.1	
1-vessel disease (%)	20.5	23.0	
2-vessels disease (%)	20.2	18.9	<0.001
3-vessels disease (%)	35.6	23.0	
Diseased vessel			
LAD (%)	63.5	49.4	
LCX (%)	49.3	38.7	<0.001
RCA (%)	54.6	41.4	
LM (%)	11.0	6.1	
Number of stented vessels	0.2±0.4	0.3±0.6	
0-vessel (%)	84.8	78.1	
1-vessel (%)	12.5	15.9	<0.001
2-vessels (%)	2.7	4.8	
3-vessels (%)	0	1.1	

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247 CAD: coronary artery disease, LAD: left anterior descending artery, LCX: left
 248 circumflex artery, LM: left main, RCA: right coronary artery

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258 **Table 4.** Multivariable analysis of the predictive value of each risk factor for CAD
 259 before and during financial crisis.

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Risk factors	Before crisis 2006-2007		During crisis 2011-2015	
	Odds Ratio (95% Confidence intervals)	p-values	Odds Ratio (95% Confidence Intervals)	p-values
Age (yrs)	1.053 (1.04-1.07)	<0.001	1.037 (1.02-1.05)	<0.001
Gender (males)	2.771 (1.97-3.89)	<0.001	3.772 (2.68-5.30)	<0.001
Family history	1.826 (1.20-2.79)	0.005	1.832 (1.16-2.89)	0.01
Smoking	3.619 (2.51-5.21)	<0.001	1.191 (0.85-1.66)	0.3
Hypertension	1.669 (1.19-2.33)	0.003	0.884 (0.63-1.24)	0.5
Dyslipidemia	3.386 (2.50-4.59)	<0.001	3.110 (2.26-4.28)	<0.001
Diabetes mellitus	2.234 (1.54-3.23)	<0.001	1.744 (1.21-2.51)	0.003
Obesity	0.924 (0.65-1.31)	0.6	0.833 (0.59-1.17)	0.3

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 262 CAD: coronary artery disease
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