## Cardiovascular features and risk of mortality in COVID-19 hospitalized patients in Spain during 2020. A nationwide study from the Minimum Basic Data Set

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**Introduction:** During the last two years scientific evidence has been gathered regarding the cardiovascular complications of Covid-19. Nevertheless nationwide studies are still required to better understand both the incidence of less frequent clinical findings, and the prognostic implications of cardiovascular COVID-19 complications.

**Purpose:** The aim of this study was to estimate the incidence of cardiovascular diseases among COVID-19 hospitalized patients in Spain during 2020, as well as their association with mortality, besides other clinical and epidemiological factors.

**Methods:** We used the Minimum Basic Data Set from the Spanish Ministry of Health (RAE-CMBD) to analyze the data of all COVID-19 hospitalized patients in Spain during 2020. This national database includes concurrent diagnostics of all studied patients codified according to the Tenth International Classification of Diseases (ICD-10).

Logistic regression analysis was performed to evaluate the influence of the different clinical and epidemiological variables in the evolution of COVID-19 hospitalized patients. Odds ratios were obtained for each variable adjusting by age and sex, and also adjusting by the rest of clinical factors. The software used for analysis was STATA v 16.1.

**Results:** 75585 men (55.15%) and 61468 women (44.85%) were hospitalized due to COVID-19 during 2020 in Spain. The median age was 66 in men and 71 in women. Mortality was 14.92% in males, and 13.81% in females. 9.62% of patients were admitted to intensive care unit (ICU). Mortality in ICU was 29.13%.

Table 1 - Clinical characteristics of COVID-19 hospitalized patients - Spain 2020

| [                                     | Men   |         | Women |         | Total  |        |
|---------------------------------------|-------|---------|-------|---------|--------|--------|
|                                       | N     | %       | N     | %       | N      | %      |
|                                       | 75585 | 55,15   | 61468 | 44,85   | 137053 | 100    |
| Arterial Hypertension                 | 25902 | 34,27   | 21522 | 35,01   | 47424  | 34,6   |
| Diabetes Mellitus                     | 18853 | 24,94   | 13265 | 21,58   | 32118  | 23,43  |
| Hypercholesterolemia                  | 2727  | 3,61    | 2498  | 3,93    | 5145   | 3,75   |
| Chronic Obstructive Pulmonary Disease | 6472  | 8,56    | 1370  | 2,23    | 7842   | 5,72   |
| Asthma                                | 2632  | 3,48    | 4482  | 7,29    | 7114   | 5,19   |
| Obesity                               | 7700  | 10,19   | 7870  | 12,8    | 15570  | 11,36  |
| Chronic Kidney Disease                | 4157  | 5,5     | 3364  | 5,47    | 7521   | 5,49   |
| Acute Renal Failure                   | 8933  | 11,82   | 5703  | 9,28    | 14636  | 10,68  |
| Hypertransaminasemia/Coagulopathy     | 3066  | 4,06    | 1594  | 2,59    | 4660   | 3,4    |
| Alzheimer/Vascular Dementia           | 2730  | 3,61    | 4494  | 7,31    | 7224   | 5,27   |
| Delirium                              | 3718  | 4,92    | 2663  | 4,33    | 6381   | 4,66   |
| Acute Myocardial Infarction           | 511   | 0,68    | 274   | 0,45    | 785    | 0,57   |
| Angina pectoris                       | 242   | 0,32    | 165   | 0,27    | 407    | 0,3    |
| Atrial Fibrillation                   | 5786  | 7,65    | 4764  | 7,75    | 10550  | 7,7    |
| Atrial Flutter                        | 681   | 0,9     | 394   | 0,64    | 1075   | 0,78   |
| Ventricular Fibrillation              | 25    | 0,03    | 13    | 0,02    | 38     | 0,03   |
| Ventricular Taquicardia               | 110   | 0,15    | 45    | 0,07    | 155    | 0,11   |
| Supraventricular Taquicardia          | 213   | 0,28    | 192   | 0,31    | 405    | 0,3    |
| Ventricular Flutter                   | 3     | 0       | 1     | 0       | 4      | 0      |
| Sinus Disease                         | 59    | 0,08    | 68    | 0,11    | 127    | 0,09   |
| Pulmonary Embolism                    | 2715  | 3,59    | 2031  | 3,3     | 4746   | 3,46   |
| Ischemic Stroke                       | 275   | 0,36    | 183   | 0,3     | 458    | 0,33   |
| Heart Failure                         | 5038  | 6,67    | 5648  | 9,19    | 10686  | 7,8    |
| Cardiomyopathy                        | 1017  | 1,35    | 440   | 0,72    | 1457   | 1,06   |
| Pericarditis                          | 48    | 0,06    | 34    | 0,06    | 82     | 0,06   |
| Myocarditis                           | 93    | 0,12    | 66    | 0,11    | 159    | 0,12   |
| Γakotsubo Disease                     | 19    | 0,03    | 51    | 0,08    | 70     | 0,05   |
| Adult Respiratory Distress Syndrome   | 4193  | 5,55    | 2113  | 3,44    | 6306   | 4,6    |
| Intensive Care Unit Admission         | 9019  | 11,93   | 4162  | 6,77    | 13181  | 9,62   |
| Invasive Mecanical Ventilation        | 4764  | 6,3     | 1971  | 3,21    | 6735   | 4,91   |
| Death                                 | 11277 | 14,92   | 8242  | 13,81   | 19519  | 14,24  |
| Age (IQR)                             | 66    | (53-79) | 71    | (54-84) | 68     | (53-81 |

Heart Failure (7.8%), Atrial Fibrillation (7.7%), Pulmonary Embolism (3.46%), Supraventricular Arrythmias (1.18%), Cardiomyopathy (1.06%), Acute Coronary Syndrome (0.87%), Ischemic Stroke (0.33%), Myocarditis (0.12%) Pericarditis (0.06%), or Takotsubo Disease (0.05%), were relevant cardiovascular findings in COVID-19 hospitalized patients (Table 1).

In the logistic regression multivariate analysis in COVID-19 patients we found epidemiological predictors of in-hospital mortality such as age (OR 2.38 for each decade), or male sex (OR 1.39). Among the clinical predictors of mortality we differentiated cardiovascular ones as Acute Coronary Syndrome (OR 1.51), Ischemic Stroke (OR 1.46), or Heart Failure (OR 1.43); and non cardiovascular ones such as admission to ICU (OR 3.12), Adult Respiratory Distress Syndrome (OR 2.74), need for Mecanical Ventilation (OR 2.52), Acute Kidney Failure (OR 2.07), Liver damage (OR 1.67), or Dementia (OR 1.66), (Table 2)

Conclusion(s): Heart Failure, Pulmonary Embolism, Ischaemic Heart Disease, Atrial fibrillation, Ischemic Stroke, or Cardiomyopathy were among the main cardiovascular diseases associated to COVID-19. They increased in a different measure the risk of mortality in COVID-19, together with factors such as Mecanical ventilation, ICU admission, Acute kidney failure, Dementia, Liver damage, Adult Respiratory Distress Syndrome, older age, or male sex.

Table 2- Odds Ratio (OR) for in-hospital Mortality in COVID-19 hospitalized patients. Spain 2020

|   | OR Adjusted by Age & Sex |            |        | OR Adjusted by Age, Sex, and Comorbidities |           |        |  |
|---|--------------------------|------------|--------|--|-----------|--------|--|
|   | OR                       | CI 95%     | р      | OR   | CI 95%    | р      |  |
| Arterial Hypertension                             | 0,75                     | 0,72-0,78  | <0,001 | 0,83                                       | 0,80-0,86 | <0,001 |  |
| Diabetes Mellitus                                 | 1,09                     | 1,05-1,13  | <0,001 | 1,05                                       | 1,01-1,09 | <0,001 |  |
| Hypercholesterolemia                              | 0,84                     | 0,77-0,91  | <0,001 | 0,84                                       | 0,77-0,91 | <0,001 |  |
| Obesity   | 1,2                      | 1,14-1,27  | <0,001 | 1,06                                       | 0,99-1,12 | 0,066  |  |
| Chronic Obstructive Pulmonary Disease             | 1,13                     | 1,07-1,20  | <0,001 | 1,17                                       | 1,10-1,24 | <0,001 |  |
| Asthma  | 0,76                     | 0,70-0,83  | <0,001 | 0,76                                       | 0,70-0,83 | <0,001 |  |
| Chronic Kidney Disease                            | 1,38                     | 1,30-1,45  | <0,001 | 1,01                                       | 0,95-1,07 | 0,697  |  |
| Acute Kidney Failure                              | 2,9                      | 2,78-3,02  | <0,001 | 2,07                                       | 1,98-2,17 | <0,001 |  |
| Hypertransaminasemia/Coagulopathy/Hepatic Failure | 2,43                     | 2,23-2,65  | <0,001 | 1,67                                       | 1,51-1,84 | <0,001 |  |
| Alzheimer/Vascular Dementia                       | 1,43                     | 1,35-1,51  | <0,001 | 1,66                                       | 1,56-1,75 | <0,001 |  |
| Acute Isquemic Cardiomyopathy                     | 2,05                     | 1,79-2,33  | <0,001 | 1,51                                       | 1,31-1,74 | <0,001 |  |
| Atrial Fibrillation                               | 1,4                      | 1,33-1,46  | <0,001 | 1,1  | 1,03-1,18 | 0,004  |  |
| Acute Pulmonary Embolism                          | 1,61                     | 1,47-1,77  | <0,001 | 1,19                                       | 1,08-1,31 | <0,001 |  |
| Ischemic Stroke                                   | 1,5                      | 1,20-1,87  | <0,001 | 1,46                                       | 1,15-1,84 | <0,001 |  |
| Heart Failure                                     | 1,73                     | 1,65-1,81  | <0,001 | 1,43                                       | 1,35-1,50 | <0,001 |  |
| Cardiomyopathy                                    | 1,45                     | 1,28-1,65  | <0,001 | 1,25                                       | 1,09-1,44 | <0,001 |  |
| Adult Respiratory Distress Syndrome               | 6,37                     | 6,00-6,77  | <0,001 | 2,74                                       | 2,56-2,93 | <0,001 |  |
| Intensive Care Unit Admission                     | 7,1                      | 6,75-7,47  | <0,001 | 3,12                                       | 2,91-3,34 | <0,001 |  |
| Invasive Mecanical Ventilation                    | 10,01                    | 9,42-10,63 | <0,001 | 2,52                                       | 2,33-2,74 | <0,001 |  |
| Age (decade)                                      | 2,08                     | 2,05-2,11  | <0,001 | 2,38                                       | 2,34-2,42 | <0,001 |  |
| Sex (Male)  | 1,13                     | 1,10-1,17  | <0,001 | 1,39                                       | 1,34-1,44 | <0,001 |  |