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# Determinants of mortality rates from COVID-19: a macro level analysis by extended-beta regression model

# Determinantes de las tasas de mortalidad por COVID-19: un análisis de nivel macro mediante el modelo de regresión beta extendida

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# ABSTRACT

**Objective** The specific mortality rate (MR) due to COVID-19 is a useful indicator for monitoring and evaluating the health strategies of health systems in the pandemic era. The main objective of this study is to estimate the effects of social, health, and economic factors on MRs in 176 countries.

**Material and Methods** Beta regression models were used, and MRs were estimated as the total number of deaths divided by the total number of confirmed cases (infection fatality rate) until December 2, 2021.

**Results** The primary findings revealed heterogeneity in mortality rates between regions and countries. The estimated coefficients showed different patterns of association between the explanatory variables and mortality rates. In the American region, the results showed a strange pattern and nearly insignificant effect for almost all variables. In Asian countries, we found a significant effect of GDP per capita and the share of the population aged 65 years and older on mortality rates, whereas on the African continent, the significant variables affecting mortality rates were GDP per capita, human development index, and share of population aged 65 years and older. Finally, in the European region, we did not find clear evidence of an association between the explanatory variables and mortality rates.

**Conclusion** These results show, in a heterogeneous way among regions, the impact of aging, development level and population density (especially with forms of distancing) on increasing the risk of death from the coronavirus. In conclusion, the pandemic has succeeded in demonstrating chaotic patterns of associations with social, health, and economic factors.

Key Words: Mortality; COVID-19; risk factors (source: MeSH, NLM).

# ABSTRACT

**Objetivo** La tasa de mortalidad específica (TM) por COVID-19 es un indicador útil para monitorear y evaluar las estrategias de salud de los sistemas de salud en la era de la pandemia. El objetivo principal de este estudio es estimar los efectos de los factores sociales, de salud y económicos sobre las RM en 176 países.

**Materiales y Métodos** Se utilizaron modelos de regresión Beta y las RM se estimaron como el número total de muertes dividido por el número total de casos confirmados (tasa de letalidad por infección) hasta el 2 de diciembre de 2021.

**Resultados** Los principales hallazgos revelaron heterogeneidad en las tasas de mortalidad entre regiones y países. Los coeficientes estimados mostraron diferentes patrones de asociación entre las variables explicativas y las tasas de mortalidad. En la región americana, los resultados mostraron un patrón extraño y un efecto casi insignificante para casi todas las variables. En los países asiáticos, encontramos un efecto significativo del PIB per cápita y la proporción de la población de 65 años o más sobre las tasas de mortalidad, mientras que en el continente africano, las variables significativas CF: Statistician. Ph. D. Applied Statistics. Associate professor of Applied Statistics. Department of Basic Education, Ferhat Abbas University. Setif, Algeria. *fatih.chellai@univ-setif.dz*  que afectaron las tasas de mortalidad fueron el PIB per cápita, el índice de desarrollo humano y porcentaje de la población de 65 años y más. Finalmente, en la región europea, no encontramos evidencia clara de una asociación entre las variables explicativas y las tasas de mortalidad.

**Conclusión** Estos resultados muestran, de manera heterogénea entre regiones, el impacto del envejecimiento, el nivel de desarrollo y la densidad de población (especialmente con formas de distanciamiento) en el aumento del riesgo de muerte por coronavirus. En conclusión, la pandemia ha logrado demostrar patrones caóticos de asociaciones con factores sociales, de salud y económicos.

Palabras Clave: Mortalidad; COVID-19; factores de riesgo (fuente: DeCS, BIREME).

o date, no country has spared the spread of coronavirus. According to a recent report by the World Health Organization (1), the total number of confirmed cases is nearly 278 million and the cumulative deaths is 5.4 million which consists of a mortality rate of 0,019. Specifically, the high mortality rate associated with Covid-19 is a challenge for countries worldwide. Accordingly, figures show heterogeneity and dispersion in mortality rates among countries and regions, thus, factors likely affecting the levels of mortality rates and that can be sources of divergences are worth studying. In another context, to face the Covid-19 pandemic, each country chose its strategy and reacted more or less quickly and more or less clearly to an unprecedented health hazard. Certainly, Lockdown strategies that contain the population and close borders undoubtedly play an important role in the fight against epidemics (2). However, beyond these measures and before virus propagation, certain health systems seemed better prepared than others to confront the pandemic. This fact was confusing when we recorded the highest number of confirmed cases and deaths in some countries characterized by high-performing health systems, a strange matter that was discussed by (3).

In this context of damages caused by the pandemic, we believe that the most dangerous effects are public health problems (e. g., mortality and pressure on health systems) (4), moreover, we think that Covid-19 is not simply a public health challenge; indeed, it caused the most severe socio-political crisis, like the SARS epidemic (5). Another outcome of the pandemic is the economic impacts, globally, the spreading channels are mainly coming from the failure of China and USA, both made 35 % of global Gross Domestic Product (GPD) (6).

Specifically, the main objective of the current study was to estimate the determinants (or sources of variation) of mortality rates caused by coronaviruses at the macro level (among countries and regions). We can conceptualize specific mortality models on two levels: micro and macro. At a micro-level (clinical and epidemiological studies), several studies have been conducted to analyze and estimate the effects of socioeconomic, demographic and genetic factors on mortality from Covid-19. Some studies have attempted to analyze the association between diabetes and Covid-19, such as (7,8), and other studies have estimated the effects of cardiovascular diseases (9,10), and ageing (11,12).

Note that the mortality rates were estimated as the total deaths divided by the total number of confirmed cases from 1-1-2020 to 3-12-2021 in 176 countries. Estimating mortality rates based on the total number of infected cases is more objective and reliable than dividing it by the total population. Dividing by the total population is considered a biased measure, especially because this virus did not affect (except in rare cases) the category of people aged between 1 and 20 years (in some countries, from 1 to 25 years), especially if we consider countries with a large population size or vice versa. Thus, the main idea of this paper is to check whether what has been revealed at the micro-level remains valid at the macro-level. For example, previous work on a micro-level showed that diabetes and cardiovascular disease were risk factors for death by Covid-19, see (7,8) for diabetes factor, and (9) for cardiovascular disease; consequently, we attempted to test whether countries with a higher prevalence of diabetes and cardiovascular diseases recorded the highest mortality rates due to Covid19? The same idea for age factor, micro-level studies (or clinical analysis) depicted that mortality risk of Covid-19 vary between young and elderly patients, see (11-13), the aim of this study was to investigate the if countries with high percentage of elderly population recorded high mortality rates compared to countries with small percentage?

In the literature, we found three main studies that worked on this topic: the first was conducted by (14) to estimate the sources of variations among 196 countries in per capita mortality rates due to Covid-19, where they applied the multivariable linear regression model. The second study was conducted by (15), who applied a beta regression model to estimate the effects of health, social, and economic factors on mortality rates during the pandemic in 129 countries. The third was achieved by (16) to analyze the determinants of mortality rates from Covid-19 in Europe during the first wave of the pandemic, and the beta regression model was fitted. As a difference between these studies, we worked on mortality rates estimated using the total number of confirmed cases, and not by the volume of the population, as studied by (14). The second study was based on the European continent which focused on the first wave of the pandemic. At this point, the study by (15) covered a short period compared to our study; such a difference can reveal a likely instability in the estimated coefficients. However, in contrast to (15), when they considered the regions as dummy variables in modeling, we designed the beta regression models for each region, and fitted a beta model for Africa, Asia, America, and Europe; This design should allow us to highlight the difference between the regions in terms of the effects of each variable on the mortality rate.

In light of previous studies, some covariates were adjusted in the modelling process to estimate their effects on mortality rates. As a regression method, statistically, when performing regression on finite data as averages, probabilities, or ratios, it can create significant statistical and metric problems first, when predicting the values of the dependent variable the regression model can give values out of range (0.1), which makes the predictions unrealistic. Second, data structures, such as rates or ratios often do not follow a normal distribution, making the ordinary least squares approach unsuitable for model estimation. For this and specification for these two points, the Beta model came to provide optimal solutions and a mathematical and standard form suitable for this structure of data (17). As a drawback of a pervious study, unfortunately, this methodological point was not taken into account by (14) when they fitted a multivariable linear regression model rather than a Beta regression model.

The rest of the paper is organized as follows, section2 presents the theoretical aspect of the beta regression model and a wide description of the dataset. Section3 was devoted to the estimation results of the regression models; section4 covers the discussion of the results and finally section5 summarizes the results of the study and draws conclusions.

# MATERIAL AND METHODS

#### Methods and Data Beta Regression model

The dependent variable in this study is the mortality rate (estimated as total deaths divided by total confirmed cases) due to Covid-19 over the period (2020 to 2021). This variable is a continuous and takes its values in the unit interval [0, 1]. In the regression analysis, the more appropriate method to fit such a variable is fractional regression or more specifically the beta regression model (BR); this model is a type of the generalized linear family models. Historically, the BRs models were developed firstly by (17). With a slight modification, we define below the density of the beta regression model that takes in consideration a set of covariates x:

$$f(y; \mu_x, \vartheta_x) = \frac{\Gamma(\vartheta_x)}{\Gamma(\mu_x \vartheta_x) \Gamma(\vartheta_x - \mu_x \vartheta_x)} y^{\mu_x \vartheta_x - 1} (1 - y)^{(1 - \mu_x) \vartheta_x - 1} (1)$$

The conditional mean of the model as  $\mu_x = E(y|x)$ , generally we work on a logit link function between  $\mu_x$  and covariates, which is defined as  $g(\mu_x) = \ln(\mu_x(1-\mu_x)^{-1})$ . We define  $\vartheta$  as scale parameter, however, the conditional variance can be written as  $Var(y|x) = \mu_x(1-\mu_x)(1+\vartheta_x)^{-1}$ . As an extension of the BR models, we can further link the scale parameter to the set of covariates, for more detail see (18).

When generalizing the density in Equation (1) to the n-sample, we find the likelihood function, which can be written in its log-transformation as

$$\sum_{i=1}^{n} w_{i} \left( \ln \left\{ \Gamma \left( \mathcal{G}_{x,i} \right) \right\} - \ln \left\{ \Gamma \left( \mu_{x,i} \mathcal{G}_{x,i} \right) \right\} - \ln \left( \Gamma \left( 1 - \mu_{x,i} \right) \mathcal{G}_{x,i} \right) + \left( \mu_{x,i} \mathcal{G}_{x,i} - 1 \right) \ln \left( y_{i} \right) + \left( \left( 1 - \mu_{x,i} \right) \mathcal{G}_{x,i} - 1 \right) \ln \left( 1 - y_{i} \right) \right)$$

Consequently, the estimation was performed using the maximum likelihood (ML) method, where we used analytical gradients by taking the estimations of the linear regression of the transformed response as the starting values. Noted that a Bayesian inference of beta regression models has been developed and widely applied in different fields, see (19), currently the Bayesian inference is currently considered a competitive approach to the ML method. Similarly, (20) attempted to provide an overview and comparison between these two inference approaches for estimating beta regression.

#### Data and Software

The data from this study contain a final sample size of 176 countries from 4 continents (Africa, Asia, America (south and north) and Europe). We excluded ten countries due to missing data. However, missing values are estimated by the "multiple imputation" method based on the MCMC principle (21). Below, we have summarized all the regression variables.

The data sources were mainly the Our World In Data website, developed by (22), World Bank database (https://data.worldbank.org/), and World Health Organization (https://covid19.who.int/). For the statistical software used in this study, different R packages were used, mainly: ("fitdistrplus " v. 1.1-6) developed by (23) to fit a beta distribution to the mortality rates. ("betareg" v.3.1-4) developed by (24) to fit a beta regression model, and the ("corrplot" v. 0.92) developed by (25) to estimate and plot the correlation matrix among the variables (Cuadro 1).

	<b>v</b>
Variable	Description
Мо	Mortality rate (as dependent variable) that was estimated as: the number of totaldeathdivided by number of total confirmed cases
Popdens	Number of people divided by land area, measured in square kilometers, most recent year available
Plus65	Share of the population that is 65 years and older, most recent year available
Gdpc	Gross domestic product at purchasing power parity (constant 2011 international dollars), most recent year available
CVDR	Death rate from cardiovascular disease in 2017 (annual number of deaths per 100,000 people)
Diabet	Diabetes prevalence (% of population aged 20 to 79) in 2017
Beds	Hospital beds per 1,000 people, most recent year available since 2010
E0	Life expectancy at birth in 2019
HDI	A composite index measuring average achievement in three basic dimensions of human development—a long and healthy life, knowledge and a decent standard of living. Values for 2019, imported from http://hdr.undp.org/en/indicators/137506

#### RESULTS

#### Descriptive analysis

In this subsection, we describe and summarize the level and dispersion of the covariates during the study period. Table 1 summarizes the dataset based on the mean values and standard deviation (SD) worldwide and across four continents (Asia, Europe, Africa, and America).

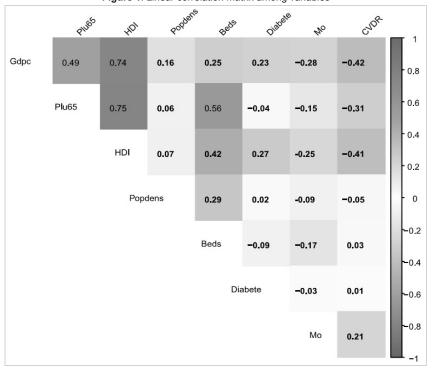
According to the results in tabel1, the mortality rate due to Covid-19 worldwide is 0.0206, which means that among 10,000 confirmed cases of Coronavirus 206 persons have died. However, this rate was not the same across the four continents. During the study period, the Americas (North and South) reported the highest mortality rate of 0.0259, followed by Africa, which reported a mortality rate of 0.0222. The Asian region was ranked third, with a mortality rate of 0.02; however, the lowest mortality rate was reported in the European region (0.0162). Note that these mortality rates were not homogenous among the countries in each region. We estimated the coefficient of variation ( $cv = (Mean/SD) \times 100$ ), and the results clearly showed that the rates were more dispersed in the European region (CV = 167.7 %), followed by the African region (CV=157.4 %), the Americas (CV=144.7%), and finally the Asian region (CV = 68.1 %). For some details, in the European region, Norway recorded the lowest mortality rate (0.0039), Bosnia-Herzegovina recorded the highest rate (0.0457); in the Asian region, Qatar first had the lowest mortality rate (0.0025), and Syria recorded the highest mortality rate (0.0571), followed by Twain and China; however, in the African region, Seychelles recorded the lowest mortality rate (0.0053) and Malawi had the highest mortality rate (0.0372); in the Americas, Cuba recorded the lowest rate (0.0086), and Peru had the highest mortality rate of 0.0898, approximately ten times the rate recorded in Cuba.

				•			,			
	Wo	orld	As	sia	Eur	оре	Afr	ica	Ame	erica
Variable	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD
Мо	0.0206*	0.0192	0.0200	0.0294	0.0162	0.0096	0.0222	0.0141	0.0259	0.0179
Popdens	347.41	1625.24	585.83	1537.95	587.59	2869.26	103.65	129.68	130.61	147.72
Plu65	8.80*	6.13	6.77	4.85	16.76	3.88	3.69	1.68	8.81	3.54
Gdpc	19070.2*	19378.9	23293.9	25194.7	32371.3	17839.1	5801.6	6513.5	16025.2	10646.8
CVDR	256.67*	115.76	286.97	144.10	232.94	122.52	286.91	73.89	196.30	68.79
Diabete	7.73*	3.78	9.37	3.67	6.56	1.99	5.70	4.06	9.73	2.78
Beds	3.05*	2.36	2.83	2.64	5.03	2.43	2.10	1.54	2.23	1.32
HDI	0.73*	0.15	0.74	0.12	0.87	0.08	0.57	0.11	0.76	0.08

Table 1. Description of the covariates in the study

Source: Author's estimation. (\*) means that the differences were statistically significant (at 0.05 level) between the Means of the variables across the four continents; noted that the Fisher-test was applied.

The summary statistics of the explanatory variables showed different levels between regions, indicating that the European region reported the highest share of the elderly population (65 years) with 16.8 % of the total population, followed by the Americas (8.8%). In terms of GDP per capita and HDI index, the countries of the European region recorded the highest levels, followed by the Asian countries, whereas the African regions were characterized by the lowest GDP per capita and HDI index. As health factors, cardiovascular diseases, and diabetes diseases, their prevalence is heterogeneous among the four regions, with a global prevalence of 256.67 (per 100,000 people) and 7.7% (% of the population aged 20 to 79), respectively, for CVDR and diabetes. However, the African region recorded the highest death rate from cardiovascular, whereas this continent showed the lowest prevalence of diabetes compared to the other regions.



#### Figure 1. Linear correlation matrix among variables

The bivariate analysis and associations among the study variables were conducted using the linear correlation matrix in Figure 1. We focused only on analyzing the linear correlation between the mortality rates from Covid-19 and the selected variables, and the estimated correlation coefficients showed a weak negative correlation for all variables. The sign of the coefficients was logical except for the share of the population aged 65 years or older, which is expected to be positive under the hypothesis that countries with a high proportion of the elderly population recorded the highest mortality rates due to Covid-19, this result was well analyzed in detail in the next section. However, we found the

same pattern of association between Mo and the set of covariates in the four regions. The next step in the analysis is to estimate beta regression models to measure the effects (or impact) of these factors on variations in mortality rates.

#### Results of Beta regression models

First, we tested if the data of mortality rates follow a beta distribution, for that purpose we used the Kolmogorov-Smirnov ( $\kappa$ s) test, this test is appropriate for continuous random variables. In application the "fitditrplus" R package was used and the results were reported in Table 2 and Figure 2.

Region	Calculated $D_{_n}$	Tabulated $D_{n,\alpha=0.05}$	Fitted Beta model $\mathit{Be}(\mu, artheta)$	Decision
World	0.0614	0.1008	Be(2.16, 107.41)	Accept
Africa	0.0646	0.1885	Be(2.61, 115.12)	Accept
America	0.2049	0.2242	Be(2.83, 102.47)	Accept
Asia	0.0889	0.2050	Be(1.61, 97.71)	Accept
Europe	0.1079	0.2027	Be(2.96, 179.75)	Accept

Source: Author's estimation. In a standard Kolmogorov-Smirnov table, the critical values were estimated as  $\frac{1.36}{\sqrt{n}}$  if n > 40 for the 0.05 level of significance.

Table 2 shows the results of the KS test for five samples of mortality rates: first, we test the hypothesis that the mortality rates from Covid-19 in 176 countries (world); then, we test this hypothesis for the four continents (Africa, America, Asia, and Europe). We found that the estimated values of the  $D_n$  test were all less than the tabulated values of the K-s test for a significance level of 0.05; thus, we accept that the mortality rates can be fitted by a beta distribution, where the estimated parameters of these distributions are shown in the fourth colon of Table 2.



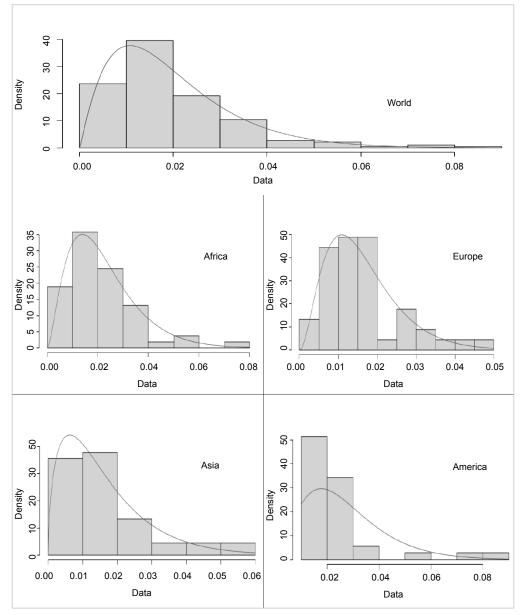


Figure 2 summarizes the test results of the five samples: full sample, Africa, Asia, America, and Europe. The dashed histograms represent the observed mortality rates, and the red curves are the fitted beta distributions. However, the density of the beta distribution  $Be(\mu, \theta)$  can take different forms depending on the values of the two parameters  $\mu$  and  $\theta$ . In our case as  $\mu > 1$ ,  $\theta > 1$ , so we have a uni-model beta distribution, note that when  $\mu = \theta$  then the density is symmetrical around  $\frac{1}{2}$ .

The results of the parameter estimates of the five beta regression models are presented in (Tables 3,4). In the whole sample (world model), the results showed that countries with a high GDP per capita, and a high share of the population aged 65 years or older were significantly associated with mean mortality rates. However, the prevalence

of diabetes and cardiovascular disease had a small positive but insignificant effect on the mean mortality rate. However, the population density and number of beds (hospital beds per 1,000 people) had a negative but insignificant effect on the mean mortality rate.

As cited in the Methods section, we deal with the extended beta regression model by adjusting some covariates to the precision function of the model (see scale components in (Tables 3,4). After some investigation in the bivariate analysis (using a scatter plot), the GDP per capita was included as a regressor in the precision function, which showed that it was highly significant. For example, in the estimated model for the full sample, for every additional unit of GDP per capita, the precision parameter increased by a factor of exp (0.0002) = 1.0002.

Table 3. Predictors of mortality	rate using beta regression	on modelin 176 countries

Table 5. Predictors of mortality rate using beta regression modeline 176 countries									
Variables	Coef.	Std. Err.	Z	P> z	[95% Conf.	Interval]			
Popdens	00002	.0000343	-0.580	0.561	0000873	.0000473			
Plu65	.0263977	.0138184	1.910	0.056	0006858	.0534811			
Gdpc	0000166	0.00000343	-4.830	0.001	0000233	-0.00000985			
CVDR	.0004843	.0004957	0.980	0.329	0004872	.0014559			
Diabete	.0055054	.0149592	0.370	0.713	0238141	.0348249			
Beds	0188098	.0258823	-0.730	0.467	0695381	.0319186			
HDI	5780984	.7900581	-0.730	0.464	-2.126584	0.970387			
Intercept	-3.556776	.4770825	-7.460	0.001	-4.491841	-2.621712			
Scale									
Gdpc	.0000262	0.0000067	3.910	0.001	.0000131	.0000393			
Intercept	4.480005	.1960587	22.850	0.001	4.095737	4.864273			

Source: Author's estimation.

Continent	Variables	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]
	Popdens	.0000106	.0008169	0.010	0.991	0015905	.0016116
	Plu65	0893938	.0359814	-2.480	0.013	1599161	0188716
	Gdpc	-0.000008	.0000136	-0.640	0.523	0000352	.0000179
	Cvdr	0005873	.0024211	-0.240	0.808	0053325	0.00416
	Diabete	.0088194	.0387557	0.230	0.821	0671404	.0847793
America (n=35)	Beds	0497579	.0986112	-0.500	0.614	2430324	.1435165
	Hdi	2.61447	3.67012	0.710	0.476	-4.57883	9.80778
	Intercept Scale	-4.54444	2.87	-1.580	0.113	-10.17	1.08187
	Gdpc	.0001324	.0000475	2.790	0.005	.0000393	.0002254
	Intercept	2.94	.8510032	3.460	0.001	1.27334	4.60921
	Popdens	0000326	.0000526	-0.620	0.535	0001358	.0000705
	Plu65	.0612469	.0305125	2.010	0.045	.0014435	.1210503
	Gdpc	0000157	0.00000458	-3.430	0.001	0000247	-0.00000673
	Cvdr	0.001017	.0007384	1.380	0.168	0004302	.0024641
	Diabete	.0302921	.0413358	0.730	0.464	0507247	.1113088
Asia (n=44)	Beds	-0.045213	.0500125	-0.900	0.366	1432357	.0528097
	Hdi	-0.812942	1.742048	-0.470	0.641	-4.227293	2.601409
	Intercept Scale	-4.15825	1.080728	-3.850	0.001	-6.276437	-2.040063
	Gdpc	0.000035	.0000115	3.040	0.002	.0000124	.0000576
	Intercept	4.576361	.4077611	11.220	0.001	3.777164	5.375559
	Popdens	0009652	.0009848	-0.98	0.327	0028954	0.000965
	Plu65	1659363	.0715376	-2.32	0.020	3061474	0257253
	Gdpc	0000521	.0000266	-1.96	0.050	0001042	-8.27E-08
	Cvdr	.0013007	.0014533	0.89	0.371	0015477	0.004149
	Diabete	.0601155	.0336385	1.79	0.074	0058147	.1260457
Africa (n=52)	Beds	0804401	.0856124	-0.94	0.347	2482373	.0873572
	Hdi	4.00206	1.663957	2.41	0.016	.7407639	7.263355
	Intercept Scale	-5.651476	.8140203	-6.94	0.000	-7.246926	-4.056025
	Gdpc	.0000651	.0000622	1.05	0.295	0000568	0.000187
	Intercept	4.760653	.4908875	9.70	0.000	3.798531	5.722775
	Popdens	.0000106	.0008169	0.010	0.990	0015905	.0016116
	Plu65	0893938	.0359814	-2.480	0.013	1599161	0188716
	Gdpc	-0.00000865	.0000136	-0.640	0.523	0000352	.0000179
	Cvdr	0005873	.0024211	-0.240	0.808	0053325	0.00416
	Diabete	.0088194	.0387557	0.230	0.820	0671404	.0847793
Europe (n=45)	Beds	0497579	.0986112	-0.500	0.614	2430324	.1435165
	Hdi	2.61447	3.67012	0.710	0.476	-4.57883	9.80778
	Intercept Scale	-4.54444	2.87	-1.580	0.113	-10.17	1.08187
	Gdpc	.0001324	.0000475	2.790	0.005	.0000393	.0002254
	Intercept	2.94	.8510032	3.460	0.001	1.27334	4.60921

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Among the regions, the parameter estimates showed different patterns of association between explanatory variables and mortality rates. In the American region, the results showed a strange pattern and nearly insignificant effect for almost all variables. In Asian countries, we found a significant effect of GDP per capita and share of population aged 65 years and older on mortality rates, whereas on the African continent, the significant variables affecting mortality rates were GDP per capita, the human development index and share of population aged 65 years and older. Finally, in the European region, we do not find clear evidence of an association between explanatory variables and mortality rates.

 Table 5. Predictive accuracy of the beta regression

Model	R <sup>2</sup>	LL	GD	AIC	BIC
World	0.1945	575.8	-1169.16	-1145.52	-1106.72
Africa	0.2006	168.1	-335.58	-311.55	-287.1
Asia	0.3021	158.8	-322.24	-298.24	-276.56
Americas	0.1857	106.6	-228.65	-204.65	-185.99
Europe	0.4454	168.2	-335.42	-311.43	-289.75

Source: R<sup>2</sup>: ;proportion of the square error predicted by the model; LogLik: log likelihood function of the estimated models; GD: Global deviance of the model (-2\*log(likelihood)); AIC: Akaike information criterion; BIC: Bayesian information criterion.

Table 5 summarizes the goodness-of-fit and predictive accuracy of the five models. In terms of pseudo-R<sub>2</sub>, the model in the European region performed the best compared to the other models, the lowest estimated value of pseudo-R<sub>2</sub> was for the American region. However, the remaining accuracy measures are mainly used in nested models to select the optimal model. Moreover, by estimating an increment function, we found heterogeneity in the contribution of each individual observation (country) to the global deviance in each model.

#### DISCUSSION

The main aim of this study was to analyze the pattern of mortality rates due to Covid-19, where particular attention was paid to estimating the effects of social, health, and economic factors on mortality rates across 176 countries. For that, Beta regression models were used across four regions (Africa, Asia, America, and Europe), this statistical method is the optimal tool for dealing with fractional data bounded in the unit interval (0,1).

Is population density a factor affecting the specific mortality rate of Covid19? We can compare and contrast the results of our study with those of (26,27) despite that they have included the factor of "population density" in the context of virus spreading and not fatality. We believe that the lack of significance of this factor in estimated models was perhaps due to the fact that the population density for a country can hide the heterogeneity between the states and regions in the same country. Furthermore, it's revealed that population density is a major factor (or predictor) of coronavirus mortality rates among counties in the USA (27).

What about the effect of ageing on mortality rates? As shown in (Tables 3,4), we can confirm that the aging covariate at the macro-level is not a prominent factor contributing to the high death rate due to Coronavirus. This result is very clear in Figure 3, which depicts the association between mortality rates and the proportion of the population aged  $\geq$  75 years across the four continents. from block (A) of Europe, we can reveal two clusters of countries, the first cluster contains the developed countries of this continent (Western Europe) of which we see a certain homogeneity of association between the levels of mortalities and share of the elderly population, except Italy which is an outlier in this block. The other cluster is that of developing countries (especially in Eastern Europe) that have recorded very high mortality rates, despite the fact that the share of the population aged 75 years and older is very low compared to the first cluster.

However, in the Asian region (B in Figure 3), there was a certain concentration of these countries in one block, except for certain aberrant observations, namely Japan, which had a very low mortality rate despite its large share of the population aged 75 years and older. In contrast, countries such as Syria and Afghanistan recorded high mortality rates despite their very low percentage of population aged 75 years or older. The same pattern of comparison can be performed in African (bloc C) and American regions (bloc D).

The above discussion was at the macro level, despite the fact that we can discuss the indirect effect of aging on the worse outcomes of the pandemic via these chronic diseases, (28). In the pandemic era, (11) revealed that aging is a generator factor for severe disease and death, and vulnerability to Covid-19 outcomes among older adults could occur via the decline and dysregulation of immune function. This hypothesis was tested by (12,13), who compared the clinical manifestation of Covid-19 in young and elderly patients.

Does the prevalence of diabetes and cardiovascular disease affect mortality rates? Our analysis is based on a macro level. Therefore, the results of this study should be considered in this context. First, we hypothesized that countries with a higher prevalence of diabetes and cardiovascular diseases would have the highest mortality rates due to Covid19. The estimated results revealed insignificant coefficients across almost all regions. In contrast, at a micro level, several studies reported that patients with diabetes (7,8) and with cardiovascular disease (9,10) have an increased risk of mortality from Covid-19.

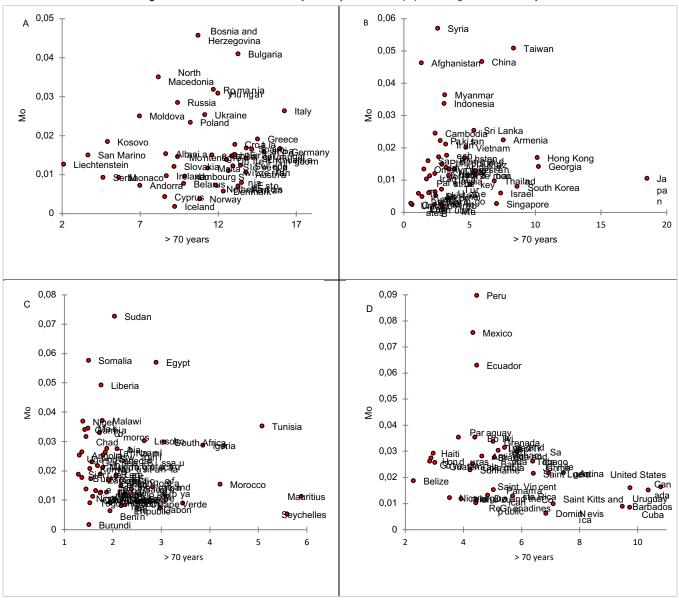


Figure 3. distributions of the mortality rates by the share of population aged 65 and elderly

On the other hand, the well-established impact of Covid-19 on the GDP has been confirmed in several studies (29). However, inverse causality was not confirmed, and the estimated results showed that the gross domestic product and HDI index had a weak significant effect on mortality rate from Covid-19 across the four regions (Africa, Asia, America, and Europe).

At this level, we can answer the following question: Why is what is validated at the clinical level not validated at the macro-and regional levels? Thus, we must maintain objectivity regarding the challenge of passing from a micro-level to a macro-level conceptualization of the determinants (or risk factors) of mortality rates due to Covid-19. A challenge for this econometric model is the problem of multicollinearity among explanatory variables, which was depicted in our case (see the correlation matrix in Table 2). A better solution for future studies is the application of the ridge-regression model, which was proposed by (30), they showed the performance of the ridge-estimation over maximum likelihood estimation. In the same context, and despite the computational ease of the likelihood-based method, the problems of non-convergence and biasing of estimation make the Bayesian inference an optimal alternative to estimate the beta regression models (20). Moreover, the Bayesian method can provide some new insights into the estimation process by exploring the prior information about some covariates (for example we can deal with an informative prior for certain parameters or hyper-parameters).

A proper pattern of the pandemic is its dynamic and perhaps chaotic behavior in some contexts; this idea was depicted by (15). Perhaps what is found to be significant over one period (or during a wave of this virus) might not be significant over other time periods Here, we can talk about the heterogeneity of the pandemic behavior across the first, second and third wave and currently the fourth wave. Another challenge is the differences and accuracy of data recording (for new cases and deaths), particularly in the underdeveloped countries of Africa and Asia. The problem of competing risks of death could be a hidden factor that can bias the specific mortality rates from Covid-19. In this context some researchers asked a challengeable question about the factors making death and mortality differ between states, countries and regions (31). In a specific context, the existence of competing risk factors can be remedied (at the micro level) using multistate models (32).

Summing up the results, it can be concluded that further investigation is needed to analyze the behavior of gthe pandemic and the patterns of associations with social, economic, and health factors at the macro level  $\blacklozenge$ 

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#### REFERENCES

- World Health Organization. Weekly epidemiological update on CO-VID-19 - 28 December. [Internet]. Geneva: WHO; 2021 [cited 2021 Dec 1]. https://bityl.co/CAuO.
- Alfano V, Ercolano S. The efficacy of lockdown against COVID-19: a cross-country panel analysis. Appl Health Econ Health Policy. 2020; 18:509-517. https://doi.org/10.1007/s40258-020-00596-3.
- Legido-Quigley H, Asgari N, Ying Teo Y, Leung GM, Oshitani H, Fukuda K, et al. Are high-performing health systems resilient against the CO-VID-19 epidemic? The Lancet. 2020; 395(10227):848-50. https://doi.org/10.1016/s0140-6736(20)30551-1.
- Blumberg S, Borlase A, Prada JM, Solomon AW, Emerson P, Hooper PJ, et al. Implications of the COVID-19 pandemic in eliminating trachoma as a public health problem. Trans R Soc Trop Med Hyg. 2021; 115(3):222-8. https://doi.org/10.1093/trstmh/traa170.
- Chang HJ, Huang N, Lee CH, Hsu YJ, Hsieh CJ, Chou, YJ. The impact of the SARS epidemic on the utilization of medical services: SARS and the fear of SARS. American journal of public health. 2004; 94(4);562-4. https://dx.doi.org/10.2105%2Fajph.94.4.562.
- Carlsson-Szlezak P, Reeves M, Swartz P. What coronavirus could mean for the global economy. Harvard Business Review [Internet]. 2020 [cited 2022 Mar 1]; 3(10). https://bityl.co/CAus.
- Rubino F, Amiel SA, Zimmet P, Alberti G, Bornstein S, Eckel RH, et al. New-onset diabetes in Covid-19. N Engl J Med. 2020; 383(8):789-90. https://doi.org/10.1056/nejmc2018688.
- Cuschieri S, Grech S. COVID-19 and diabetes: The why, the what and the how. Journal of Diabetes and its Complications. 2020; 34(9):107637. https://dx.doi.org/10.1016%2Fj.jdiacomp.2020.107637.

- Bansal M. Cardiovascular disease and COVID-19. Diabetes & Metabolic Syndrome: Clinical Research & Reviews. 2020; 14(3):247-50. https://doi.org/10.1016/j.dsx.2020.03.013.
- Barison A, Aimo A, Castiglione V, Arzilli C, Lupón J, Codina P, et al. Cardiovascular disease and COVID-19: les liaisons dangereuses. Eur J Prev Cardiol. 2020; 27(10):1017-25. https://doi.org/10.1177/2047487320924501.
- Chen Y, Klein SL, Garibaldi BT, Li H, Wu C, Osevala NM, et al. Aging in COVID-19: Vulnerability, immunity and intervention. Ageing Res Rev. 2021; 65:101205. https://doi.org/10.1016/j.arr.2020.101205.
- Mori H, Obinata H, Murakami W, Tatsuya K, Sasaki H, Miyake Y, et al. Comparison of COVID-19 disease between young and elderly patients: Hidden viral shedding of COVID-19. J Infect and Chemother. 2021; 27(1):70-75. https://doi.org/10.1016/j.jiac.2020.09.003.
- Liu K, Chen Y, Lin R, Han K. Clinical features of COVID-19 in elderly patients: A comparison with young and middle-aged patients. J Infect. 2020; 80(6):e14-e18. https://doi.org/10.1016/j.jinf.2020.03.005.
- 14. Leffler CT, Ing E, Lykins JD, Hogan MC, McKeown CA, Grzybowski A. Association of country-wide coronavirus mortality with demographics, testing, lockdowns, and public wearing of masks. The American journal of tropical medicine and hygiene. 2020; 103(6):2400-11. https://doi.org/10.4269/ajtmh.20-1015.
- De Oliveira RP, Achcar JA, Nunes AA. Modeling the incidence and death rates of COVID-19 pandemic in different regions of the world. Epidemiologic Methods. 2020;9(s1). https://doi.org/10.1515/em-2020-0017.
- Cordeiro GM, Figueiredo D, Silva L, Ortega EM, Prataviera F. Explaining COVID-19 mortality rates in the first wave in Europe. Model Assist Stat Appl. 2021; 16(3):211-21. https://doi.org/10.3233/MAS-210534.
- Ferrari S, Cribari-Neto F. Beta regression for modelling rates and proportions. Journal of Applied Statistics. 2004; 31:799-815. https://doi.org/10.1080/0266476042000214501.
- Smithson M, Verkuilen J. A better lemon squeezer? Maximum-likelihood regression with beta-distributed dependent variables. Psychological Methods. 2006; 11:54-71. https://doi.org/10.1037/1082-989x.11.1.54.
- Branscum AJ, Johnson WO, Thurmond MC. Bayesian beta regression: applications to household expenditure data and genetic distance between foot-and-mouth disease viruses. Aust N Z J Stat. 2007; 49(3):287-301. https://doi.org/10.1111/j.1467-842X.2007.00481.x.
- Liu F, Eugenio EC. A review and comparison of Bayesian and likelihood-based inferences in beta regression and zero-or-one-inflated beta regression. Stat Methods Med Res. 2018; 27(4):1024-44. https://doi.org/10.1177/0962280216650699.
- Schafer JL. Multiple imputation: a primer. Statistical methods in medical research. 1999; 8(1):3-15.

https://doi.org/10.1177/096228029900800102.

- Hannah R, Mathieu E, Rodés-Guirao L, Appel C, Giattino C, Ortiz-Ospina E, Hasell J, Macdonald B, BeltekianD, and Roser M. Coronavirus Pandemic (COVID-19). Our World in Data [Internet]. 2020[cited 2021 Nov 28]. https://bityl.co/CAvk.
- Delignette-Muller ML, Dutang C. Fitdistrplus: An R package for fitting distributions. Journal of statistical software. 2015; 64(4):1-34. https://doi.org/10.18637/jss.v064.i04.
- Zeileis A, Cribari-Neto F, Gruen B, Kosmidis I, Simas AB, Rocha AV, Zeileis MA, Package 'betareg'. R package [Internet]. 2016[cited 2021 Nov 28]; 3(2). https://bityl.co/CAvr.
- Wei T, Simko V, Levy M, Xie Y, Jin Y, Zemla J. Package 'corrplot'. Statistician [Internet]. 2017[cited 2021 Nov 28]; 56(316):e24. https://bitvl.co/CAvw.
- Adekeye O, Olawole-Isaac A, Fasina F, AdekolaPO, Samuel GW, Akanbi MA. Nigeria and Italy divergences in coronavirus experience: impact of population density. The Scientific World Journal. 2020. https://doi.org/10.1155/2020/8923036.

- Zhang CH, Schwartz GG. Spatial disparities in coronavirus incidence and mortality in the United States: an ecological analysis as of May 2020. J Rural Health. 2020; 36(3):433-445. https://doi.org/10.1111/jrh.12476.
- Kennedy BK, Berger SL, Brunet A, Campisi J, Cuervo AM, Epel ES, et al. Geroscience: linking aging to chronic disease. Cell. 2014; 159(4):709-13. https://doi.org/10.1016/j.cell.2014.10.039.
- 29. Maliszewska M, Mattoo A, van der Mensbrugghe, D. The potential impact of COVID-19 on GDP and trade: A preliminary assessment [Internet]. Washington: World Bank Policy Research Working Paper. 2020 [cited 2021 Nov 28]. https://bityl.co/CAwA.
- Abonazel MR, Taha IM. Beta ridge regression estimators: simulation and application. Commun Stat Simul Comput. 2021; 1-13. https://doi.org/10.1080/03610918.2021.1960373.
- 31. Henriques M. Coronavirus: Why death and mortality rates differ. BBC Future; 2020 [cited 2021 Nov 28]. https://bityl.co/CAwG.
- Andersen PK, Keiding N. Multi-state models for event history analysis. Statistical methods in medical research. 2002; 11(2):91-115. https://doi.org/10.1191/0962280202sm276ra.