



Review

Syndemic: A Synergistic Anthropological Approach to the COVID-19 Pandemic

Giuseppe Calcaterra ¹, Pier Paolo Bassareo ², Francesco Barilla ³, Francesco Romeo ⁴, Cesare de Gregorio ⁵, Paulette Mehta ⁶ and Jawahar L. Mehta ⁶*

- ¹ Faculty of Medicine and Surgery, Postgraduate Medical School of Cardiology, University of Palermo, 90127 Palermo, Italy; peppinocal?@gmail.com
- ² School of Medicine, University College of Dublin, Mater Misericordiae University Hospital, D07 R2WY Dublin, Ireland; piercard@inwind.it
- ³ Dipartimento Medicina dei Sistemi, University Tor Vergata, 00133 Rome, Italy; francesco.barilla@uniroma1.it
- ⁴ UniCamillus International Medical University, 00131 Rome, Italy; romeocerabino@gmail.com
- Department of Clinical and Experimental Medicine, University of Messina, 98122 Messina, Italy; cesare.degregorio@unime.it
- 6 Division of Cardiovascular Medicine, University of Arkansas for Medical Sciences and the Central Arkansas Veterans Healthcare System, Little Rock, AR 72205, USA; mehtapaulette@uams.edu
- * Correspondence: mehtajl@uams.edu; Tel.: +1-(501)-296-1426

Abstract: This review describes the relationship between the coronavirus-related pandemic and health inequities. The latter are linked to pre-existing social and economic discriminations in terms of access to healthcare for people affected by chronic diseases. We believe that we are living in a "syndemic pandemic". The term "syndemic" was originally developed by the medical anthropologist Merrill Singer in the 1990s in order to recognize the correlation between HIV/AIDS, illicit drug use, and violence in the United States. This complex interplay exacerbated the burden of the disease and the prognosis of the patient. Similarly, in COVID-19 infection, socio-economic, ethnic, and racial inequities result in higher morbidity and mortality in certain sections of society. Unfortunately, such differences are becoming too common during the COVID-19 pandemic, in terms of the incidence and prevalence of the disease, as well as inequal access to new medical advances and life-saving therapeutics for those with COVID-19, such as vaccines and monoclonal antibody treatment. Lockdown measures, imposed internationally as a response to the COVID-19 pandemic, are causing economic inequities, which complicate the issue even further. An appropriate syndemic anthropological approach is necessary to ensure that this pandemic does not increase health inequities in access to appropriate treatments.

Keywords: COVID-19; syndemic; pandemic; health; inequality

Citation: Calcaterra, G.; Bassareo, P.P.; Barilla', F.; Romeo, F.; de Gregorio, C.; Mehta, J.L. Syndemic: A Synergistic Anthropological Approach to the COVID-19 Pandemic. *Encyclopedia* 2022, *2*, 1344–1356. https://doi.org/ 10.3390/encyclopedia2030090

Academic Editors: Yury Zhernov, Milva Pepi and Raffaele Barretta

Received: 31 March 2022 Accepted: 7 July 2022 Published: 13 July 2022

Publisher's Note: MDPI stays neutral with regard to jurisdictional claims in published maps and institutional affiliations.



Copyright: © 2022 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (http://creativecommons.org/licenses/by/4.0/).

1. Introduction

A pneumonic illness of unknown origin was first identified in Wuhan, Hubei province, China, in December 2019. The World Health Organization (WHO) recognized the disease as being caused by severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2). The acronym COVID-19 (coronavirus disease-2019) was then coined to term the illness. Since its first appearance, it has spread all over the world, thus representing a true global public health challenge. The WHO declared that the disease was a "public health emergency of international concern" on 20 January 2020. Following that, it was declared as a "pandemic" on 11 March 2020. As of 22 January 2022, the pandemic has caused more than 346 million cases and 5.58 million deaths, undoubtedly making it one of the worst infections in human history [1]. However, the epidemiological study of the pandemic is

still ongoing, and, thus, it is affected by the need to steadily update data, which are often incomplete, owing to the emergency-induced chaos [2].

Of note, the COVID-19 pandemic has been found to have a strong link with socio-economic phenomena.

2. Syndemic Definition

Merrill Singer, an American medical anthropologist, coined the neologism "syndemic" by merging the two words synergy and epidemic in the mid-1990s [2,3]. "Syndemic" defines a theory according to which epidemics arise from the complex interplay between the spread of a disease and social/environmental/economic factors, which, in turn, negatively impact the disease itself. To date, the concept of "syndemic" has generally been applied to chronic medical conditions [2,3]. Conversely, disease co-occurrence, with or without any interaction, is known as "comorbidity".

The terms "comorbidity" and "syndemic" are different, since the first term tends to only focus on nosography, whereas "syndemic" mostly refers to the concomitant socio-economic factors. However, there is some overlap between the two terms. Two or more diseases can be comorbid with or without any social and/or economic influence. Conversely, the term "syndemic" is characterized by the influence of the accompanying economic and social features (such as poverty, exploitation, and oppressive social relationships), which, taken together, increase a person's susceptibility to an illness [4].

The *syndemic* approach to diseases is now gaining increasing recognition in the public domain and global health research settings.

Epidemiologists and medical anthropologists study health disparity, which is linked to poverty, stress, and violence. Racial and ethnic inequalities in access to healthcare are becoming increasingly more evident globally during the current COVID-19 pandemic, and they are having markedly negative long-lasting effects on health [5].

The term "syndemic" was used for the first time to describe community-level disparity patterns in substance abuse, violence, the occurrence of HIV/AIDS, and the occurrence of other conditions, and to help understand their relationships, especially as they are related to modifiable behaviors, in the USA [2,3]. Rather than being separate entities, substance abuse, violence, and AIDS—referred to as SAVA to stress their mutual relationship—represent a single syndetic (i.e., a closely interrelated complex of health and social crises) that continues to take a significant toll on the lives and well-being of the urban poor [6].

Specifically, a "syndemic" approach examines why certain diseases occur in clusters. This approach also studies the pathways through which a biological state or disease interacts with socio-economic conditions in individuals/groups of people, thereby exacerbating the overall disease burden. A degraded social environment, the conditions of community inequality and injustice, and disease are considered the worst features contributing to a patient's increased vulnerability.

Although often used to explain conditions such as sepsis, the term *syndemic* is not limited to infectious diseases. Non-communicable illnesses, such as cardiovascular disorders, diabetes, hypertension, chronic obstructive pulmonary disease, and cancer, also occur in the same settings as infectious diseases, that is, in low-income urban populations, and they affect the health of the individual and strains the healthcare system [7].

An example of this is type 1 diabetes in children. The immune basis of type 1 diabetes has been well defined. Since the autoimmune reaction and beta-cell loss start long before the diagnosis of type 1 diabetes, it is not surprising that the incidence of type 1 diabetes rose in concert with the peak occurrence of COVID-19, as well as with the pandemic containment measures in the 3 months following the appearance of COVID-19. Thus, it appears that type 1 diabetes is not caused by SARS-CoV-2 infection but, rather, is the result of environmental changes associated with the pandemic itself or pandemic containment measures [8].

With the aim of overcoming the lack of any national, state, and local public health data on the unbalanced economic and social impact of COVID-19, USA counties and zip codes in Illinois as well as New York City were matched with COVID-19 deaths, established COVID-19 cases and positive COVID-19 swabs. The given period was from 23 January 2020 to 5 May 2020. The use of zip codes allowed us to identify areas with high rates of poverty, crowding, and a high black population, as well as the Index of Concentration at the Extremes. The source of the data was the dataset USA Facts. By 5 May 2020, the COVID-19 death rates were 143.2/100,000/year vs. 83.3/100,000/year in low versus high poverty counties (≥20% vs. <5% of persons below poverty); 124.4/100,000/year vs 48.2/100,000/year in counties in the highest versus lowest quintile, relative to household crowding; and 127.7/100,000/year vs 25.9/100,000/year, relative to counties in the highest versus lowest quintile for the percentage of black population. As such, severe social inequities in the USA were found to relate to COVID-19 outcomes. This report supports the need for a change in policy and resource allocation [9].

Another example of *syndemic* inequality in access to care is related to COVID-19 therapy. Intravenous or subcutaneous anti–SARS-CoV-2 monoclonal antibodies are usually provided to non-hospitalized subjects suffering from mild-to-moderate positivity of SARS-CoV-2 infection and to those at increased risk of progressing to severe disease and admission to hospital. According to the Centers for Disease Control and Prevention, some racial and ethnic communities—namely Black, Hispanic or Latino, and American-Indian or Alaska-native subjects—are at an increased risk of requiring admission to hospital or passing away due to COVID-19 in comparison to Caucasians. The same groups are also those who are less often offered monoclonal antibodies [10].

3. Syndemic Pandemic

The harmful combination of the COVID-19 pandemic with widely diffused non-communicable diseases has put a strain on already existing social and economic discrepancies. In other words, access to the healthcare system for the treatment of non-communicable diseases, which was already significantly limited for lower classes, has become even more difficult in the COVID-19 era.

We believe that public health officials are taking the wrong approach in managing this outbreak. Unfortunately, public health officials have so far considered COVID-19 simply as an infectious disease. Richard Horton, Editor-in-chief of The Lancet, elegantly stressed this feature of the pandemic in a recent editorial. The link between COVID-19 and non-communicable illnesses with an underlying history of social and economic inequality brings out the negative consequences of each distinct illness [11]. The *syndemic* model is an emerging approach to healthcare in clinical practice. We should start thinking beyond the traditional historical concept of diseases as separate entities that are apart from other illnesses and not subject to social differences. Dr Horton concluded that "COVID-19 is a *syndemic*, not a pandemic" [11].

3.1. Theoretical Background

Applying the concept of *syndemic* to COVID-19 has the potential to help politicians and healthcare program implementers in their attempts to improve the health of the general population in view of facing other similar crises in the future.

Considering social factors, such as social disparity, differentiates syndemic processes from the classic biomedical notion of comorbidity and also differentiates syndemics among human beings from synergistic disease interplay among animals. Syndemic theory relies on recognizing the pivotal role of biosocial interrelationships. The latter is of particularly relevance to humans and it represents a new approach that has not been recognized in the past.

Concerning the word *syndemic*, which can also be spelled as *syn-demic*, the first of the two words comes from the Greek word $\sigma\acute{\nu}\nu$, meaning "together". It is used when two or more agents act together to generate a greater effect than each of them acting alone. The

second word, e.g., *demic*, is a suffix which derives from another Greek word, $\delta\tilde{\eta}\mu\rho\varsigma$, meaning "people". It has been previously used in three cornerstone concepts in public health: *epidemic*, a term that is used to describe greater than expected jumps in the frequency of an illness in a given population; *pandemic*, an epidemic spreading across multiple populations or even worldwide, such as COVID-19; and *endemic*, a disease that is well established in a population and remains year after year.

The syndemic perspective begs the questions of what predisposing factors have given rise to the spread of COVID-19 (for instance, overcrowded built environments, pollution, and climate change), what inequalities has COVID-19 exacerbated among vulnerable populations, and what strategies could be employed to detect and reduce such inequalities. To reach these aims, a scientific, rather than empirical, approach is preferred [12].

3.2. Lockdown

The first line approach suggested by governments and epidemiologists worldwide to hamper viral transmission and reduce contagion was based on non-pharmaceutical interventions, such as intensified hand hygiene, social distancing, self-isolation, quarantine, and lockdown of whole countries. This approach is quite similar to that used for plague containment, the old "quarantine" [13]. However, we now know that COVID-19 affects "fragile" people with non-communicable diseases the most. The state-imposed restrictions have various degrees of strictness, but all share a noteworthy increase in social isolation and discrimination. There is no doubt that overcrowded nursing homes with a lack of social distancing and isolation have represented the most dramatic outbreak during the COVID-19 pandemic and that elderly poor people were forced into this environment, thus representing an example of inequality [14].

The first consequences of the COVID-19 emergency restrictions are likely to have unequally triggered multiple health impacts. They will range from unbalanced experiences of lockdown (loss of jobs and income, living in overcrowded spaces, being confined in buildings with no access to leisure time activities) to how the restrictions themselves worsening the social features of health (e.g., very limited access to healthcare systems for patients with diseases other than COVID-19, since all hospitals are overcrowded with COVID-19 patients), and then to disparities in health consequences due to the lockdown (reduced sports activity, increased mental health depression and gender-based violence).

The world economy has been dramatically affected by COVID-19 and its complications. The aftermaths of economic disasters appear to be similarly unequally distributed, thus making heath disparities worse [15]. For example, in a study assessing the impact of COVID-19-related aftermaths on household income and food security in two African countries (Kenya and Uganda), a significant income drop was detected in approx. 70% of the sample. Food availability decreased by 38% and 44% in Kenya and Uganda, respectively, and dietary also quality worsened. In this respect, in both countries, the regular consumption of fruits decreased by around 30% during the pandemic compared to previously. The income-poor households and people depending on labor income without any savings were the most vulnerable. On the contrary, farmers were less likely to struggle with the same troubles. The findings suggest that in the future, the government should focus their efforts on building strong financial institutions to support the recovery of businesses in the medium term, ensuring the resilience of food supply chains [16]. The Covid-19 pandemic is challenging, not only for health sectors but also for public administration systems. As soon as the current unprecedented circumstances normalize, administrators and politicians should learn from the current crisis by gathering and analyzing data, building international networks, and preparing themselves to better cope with the next crisis [17].

Compliance with the COVID-19-related restrictive measures introduced by local governments to prevent contagion varies significantly depending on many factors, including different cultural features. In particular, individualism, which gives special value to personal freedom, makes government action more difficult, whilst collectivism, emphasizing the wellbeing of people, makes government measures easier. This is particularly evident when making a comparison between the US and China [18], but also within the US itself. In fact, analyzing a database of more than 3000 counties of the 50 American states, the percentage of those wearing masks was found to be higher in more collectivistic US states [19]. On the other hand, culture, identified as the existence of shared beliefs and values by members of the same nation or state, has a significant impact on the quality of the government itself. The impact is stronger than that exerted by other factors such as institutions and economic development [20].

In research carried out in the US, partisanship, defined as party identification, intended 2020 Presidential vote, and self-placed ideological positioning, proved to play a pivotal role in shaping individual responses to restrictions since the early beginning of the COVID-19 pandemic [21].

Additionally, social media has an influence on people's response to COVID-19-im-posed restricted mobility. In fact, the higher the number of Twitter messages containing words such as "stay home", "stay safe", "wear mask", "wash hands", and "social distancing", the more mobility was seen to decrease [22].

4. Vaccine Inequality

As stated above, during the early phases of COVID-19, the world community was totally unprepared to face the outbreak and the above-stated non-pharmaceutical interventions were the only possible way to reduce COVID-19 mortality and morbidity [23]. Fortunately, a number of vaccines have been developed, approved by governments and distributed throughout the world. Along with non-pharmaceutical interventions (such as masking and social distancing), there has been a reduction in the number of infections and deaths. On the other hand, COVID-19 vaccines represent a feasible solution and strategy for significantly limiting the spread of the disease, above and beyond what can be achieved with masking and social distancing in the long-term [24].

Nonetheless, despite the excellent efficacy and acceptable risk profiles of the vaccines, mass immunization campaigns are successful only when the vaccine rates are high, which results in herd immunity. Mass vaccination can control the viral transmission dynamics as well as provide fragile individuals with immune protection. Unfortunately, a significant percentage of individuals, despite being willing to receive a vaccination against COVID-19, are unable to do so because they are immune-deficient or immune-suppressed; their conditions do not allow them to achieve immunity levels which are sufficiently strong [25].

Apart from these clinical reasons, further heterogeneous circumstances, such as a lack of trust for scientific evidence and vaccination, beliefs concerning the supposed aggressiveness of COVID-19 and/or the effectiveness and adverse events of vaccines, and objective and perceived obstacles to immunization, can have negative consequences on the progress of mass vaccination campaigns, causing harmful health risks. In fact, as per the WHO, vaccine acceptance represents one of the more important challenges to public health [26]. Vaccine hesitancy is a multifaceted phenomenon which comes from a combination of different factors, ranging from socio-economic and educational variables to behavioral components [27,28]. For the sake of scientific honesty, the suspicion surrounding attempts to minimize some side effects to the vaccines did not help promote trust in vaccination [29–31].

Specifically, the COVID-19 pandemic has affected individuals in more than 220 countries and regions. It has by far adversely influenced healthcare sectors, the supply chains of production of anti-viral medications and vaccines and their delivery, logistics, and distribution. In the meantime, many wealthy nations started promoting immunization

campaigns, immunizing more than 50% of their entire population; some countries have even given third and fourth doses (a booster and a second booster, respectively) of COVID-19 vaccines. In contrast, the vast majority of developing countries, particularly African continent, are still having difficulty obtaining enough vaccines to start their own vaccination campaigns [32].

Low-income countries form collaborative networks to co-finance vaccine acquisition, such as the "COVID-19 Vaccines Global Access" (COVAX), the World Bank and the African Union's COVID-19 "Africa Vaccine Acquisition Task Team" (AVATT) platforms, with the aim of supporting balanced and sustainable access to COVID-19 vaccines [33,34].

It is interesting to cite the paper by Wagner and Coll., who studied the number of infections and potential risk of coronavirus variants emersion in two hypothetical regions, e.g., one with high access and the other with low access to vaccines. They found that for the first, sharing vaccines with the second would be much better with the aim of reducing COVID-19 burdens in countries with less access, lowering the number of case imports and minimizing the harmful risk of the occurrence of highly contagious viral mutations [35].

To objectively assess inequities in vaccine allocation, distribution, and uptake in low-and middle-income countries, a search was carried out, employing machine-learning [36]. This study has provided important information for those people working in global and public health systems, decision and policymakers, and all the important persons involved in vaccine roll-out. The most important aspects of the vaccine roll-out process have been identified, and these include: focusing on giving vaccines to children (children are a potential reservoir of coronavirus and may suffer from MIS-C/PIMS themselves) rather than expediting vaccination of the entire population [37]; increasing confidence in the COVID-19 vaccine with the use of famous figures as role-models and all the possible available promotional tools: training and recruiting more people to increase the vaccination process; and increasing government budgets for buying and distributing the vaccines [36]. Setting up databases such as the 'Response2covid19', a living dataset of governments' responses to COVID-19 involving more than 200 countries and which is updated every month can help in generating robust data to support public health and economic decision making [38].

5. Anthropological Approach

The *syndemic* perspective implies that human beings should be considered as the cornerstone of any pathological process ("anthropological approach"). It provides scientists with a biosocial conception of health. Therefore, a more philosophical perspective is necessary if we wish to stay free from COVID-19. Dealing with the SARS-CoV-2 infection means paying close attention to non-communicable diseases more than in the recent past. These diseases are not limited to wealthy countries. Conversely, they are diffused around the world, particularly among the poorest in the world.

By providing a framework of the interplay between diseases and socioeconomic features, an understanding of the *syndemic* theory can enhance our understanding of the pandemic we are living in, thus leading to major progress that cannot be reached when focusing on each disease as a separate entity [2].

Non-communicable diseases represent about a third of the total disease burden. The WHO has suggested several affordable, cost-effective interventions, which could prevent almost 5 million deaths from occurring, mostly among the poorest in the world, over the next decade. This is in addition to preventing deaths from COVID-19 [39].

Approaching COVID-19 as a *syndemic*, especially with the new variants threating nations, implies a comprehensive vision that includes educating people, reducing unemployment, providing affordable housing and food, and saving the countryside from pollution and property speculation. Considering COVID-19 simply as a pandemic rules out broader much needed approaches. Drugs or vaccines by themselves will not solve the future economic crisis that is arising. Increasing awareness of COVID-19 as a *syndemic* and the implementation of concrete measures to limit disparities are now needed to decrease

the onslaught and spread of the epidemic and the burden of health disparities globally [11]. As there are strong biological and environmental reasons to expect more serious epidemics than COVID-19, a *syndemic* approach can help researchers, clinicians, and other healthcare providers to identify the presence of socio-economic factors interacting with the disease and adversely affecting one another [40].

6. An Issue Coming from the Past

COVID-19, now in its third year, has brought forth a number of issues concerning balanced access to diagnostic tools and provision of what is necessary to heal, including vaccines. The lack of equity in the USA and other countries is not a new phenomenon. Social and financial was somehow embedded in the ecosystems a long time ago. Not all racial groups have the same opportunities in obtaining access to the best therapies for socioeconomic and other reasons. Health care providers also have to pull their weight. Examining the available data and evidence on health inequalities represents a valuable starting to point to identify weaknesses and try to fix them [41].

Historically, pandemics not only exacerbate marginalization but they also set the stage for inequalities that persist for centuries. For example, pathogens introduced during European colonialism contributed to indigenous population collapses throughout the Americas, thus making colonial exploitation easier. Even today, the Navajo population in the US, Mexican indigens in Central America, and Manaus people in Brazil are struggling to control the spread of viral diseases due to difficulties in obtaining access to healthcare facilities [42].

7. Monoclonal Antibodies and Equity

Many of us have already had the unpleasant experience of witnessing inequalities and that the most infected patients, those struggling for hospitalization, and having the worst outcomes are mostly of non-white race or belong to specific ethnic groups. This sad observation highlights the need to overcome these boundaries and to think about how we can provide people with access to health care facilities in an equal way. At the beginning of the outbreak, research helped scientists to identify some risk factors of those suffering from a more severe form of the coronavirus infection, namely the elderly, male gender, people with pre-existing lung disease, overweight/obesity, and diabetes [43]. Since then, other risk factors, even more harmful, have been identified, including being of non-White race or coming from other ethnic groups (including Black, Asian, Hispanic, American Indian or Alaska Native, Native Hawaiian or Other Pacific Islander, and multiple other races). This is particularly evident in access to and use of neutralizing monoclonal antibody therapy, as previously outlined. The latter are designed to mimic the body's natural immune response. Their use has been authorized on the basis of the emergency situation. When administered in the early phase of infection, monoclonal antibodies have proved to be effective in preventing COVID-19 from progressing to a severe form requiring hospitalization [44]. Specifically, the provision of monoclonal antibodies was significantly less common (4% of the population or less) in all non-White communities compared with White communities.

Hispanic patients received monoclonal antibodies 58% less often than non-Hispanic patients. Black patients, Asians, or individuals of other descent received monoclonal antibodies 22%, 48%, and 47% less often, respectively, than White patients (November 2020-August 2021) [45].

8. Other Supporting Evidence

The US Centers for Disease Control and Prevention have been continuing to collect data concerning COVID-19 infected, hospitalized, and deceased patients. However, data about race and ethnicity are missing. Notwithstanding this limitation, the gathered data show that in comparison with White Americans, Native Americans and Alaska Natives

are one and a half times more likely to suffer from COVID-19. Furthermore, rates of hospitalized and deceased patients are also much higher in these groups. Again, only about 10% of the American Black population have been vaccinated at least once against coronavirus, although they account for 12–13% of the whole US population. A more proactive attitude is needed to address these major issues in advance [46].

In a recently released paper [47], social determinants of health, which have been previously mentioned as important factors linked with health inequities, have been studied regarding their association with COVID-19 death rates among communities of different descent and rural, suburban, and urban areas during the first year of the pandemic (22 January 2022–28 February 2022) in the USA.

All the American states and counties, as well as the District of Columbia, were included. Areas with a high presence of a single race or ethnicity and a high COVID-19 death rate were named "concentrated longitudinal-impact counties". The three most represented racial and ethnic communities were observed to be: Black or African American, Hispanic or Latin, and non-Hispanic White populations.

Social determinants of health were identified, namely low income, lack of health insurance, lack of family physician, preventable hospital admissions, housing crisis, and limited access to internet. Their possible association with COVID-19 fatality (deaths per 100,000 population) was tested by using a complex statistical analysis. Again, four indexes were used to measure multiple dimensions of social determinants of health: socio-economic advantage, limited mobility, urban core opportunity, and mixed immigrant cohesion and accessibility.

More than three thousand counties were enrolled in the search and about 500 were concentrated in the longitudinal-impact counties. Among the counties, 11.0% of the population was Black or African American, 6.3% was Hispanic or Latin, and in 1.1% was non-Hispanic White. Approximately half a million COVID-19-related deaths were registered. Concentrated longitudinal impact counties with a prevalent Black or African American community were in urban, suburban, and rural areas and faced a great number of disadvantages such as low income (85.6%) and preventable hospitalization (81.0%). Concentrated longitudinal-impact counties with a prevalent Hispanic or Latin community were located in urban areas (57.6%), and in about two thirds of these counties most of the people did not have health insurance. Concentrated longitudinal-impact counties with a prevalent non-Hispanic White population were in rural areas (69.7%) the most. These people were old (78.8%) and had limited access to quality health care (72.7%). In urban areas, the mixed immigrant cohesion and accessibility index proved to be inversely linked with the COVID-19 deaths rate, thus showing the association between the latter and the presence of immigrant communities, whose social model was a traditional family-oriented society with accessibility stressors and living in overcrowded buildings. Higher COVID-19 mortality rates were associated with preventable hospitalization in rural areas and higher socioeconomic status vulnerability in suburban areas. Across all communities, limited internet access was correlated with a higher number of deaths, mostly in urban areas.

On balance, the research showed a dramatic association between different social determinants of health measures and COVID-19 fatality rate, with some fluctuations across different racial and ethnic communities and geographical areas [47].

Again, in another recent study, neighborhood conditions measured before the COVID-19 pandemic proved to have strong predictive power for subsequent incidence, with mobility-based disadvantages playing a pivotal role, even greater than residents' so-cioeconomic features [48].

However, it does not take a lot of research to see a connection between race and poor healthcare before and during the pandemic. This is exemplified by the case of Dr. Susan Moore, an African American physician from Indiana, a state in the United States, who was hospitalized for COVID-19. A physician herself, she immediately felt that her hospital and treatment were impacted by her race. She published her claims on Facebook Live and her

story quickly gripped the African American community. This community was already experiencing a huge disparity in terms of infection rates versus the rest of the American population. Unfortunately, Dr. Moore died a few weeks after her initial complaint about denial of care. This has been mentioned in the book *Heart Disease: It Is All in Your Head* [49].

An example of ethnic inequalities detected during the first wave of COVID-19, which is also the phase of the disease with the largest number of examined data, is reported in Table 1 [50].

Table 1. Ethnic differences in SARS-CoV-2 infection and COVID-19 outcomes during the first wave (1 February —3 August 2020) of the COVID-19 pandemic in the United Kingdom. All data are expressed in terms of hazard ratio [50].

Commu-	Likelihood of Being Tested Risk of Testing Positive		Risk of COVID-19 Related Risk of COVID-19 Related Risk of COVID-19 Re-		
nity	for SARS-CoV-2	for SARS-CoV-2	Hospitalization	ICU Admission	lated Death
Other eth- nicity	0.77 [95% CI 0.76–0.78]	1.20 [95% CI 1.14–1.28]	1.54 [95% CI 1.41–1·69]	3.18 [95% CI 2.58–3.93]	1.22 [95% CI 1.00–1.48]
South Asian	1.08 [95% CI 1.07–1.09]	1.99 [95% CI 1·94–2.04]	1.48 [95% CI 1.41–1.55]	2.18 [95% CI 1·92–2.48]	1.26 [95% CI 1.15–1.37]
Black	1.08 [95% CI 1.06-1.09]	1.69 [95% CI 1.62-1.77]	1.78 [95% CI 1.67-1.90]	3.12 [95% CI 2.65-3·67]	1.51 [95% CI 1.31-1.71]
Mixed eth- nicity	1.04 [95% CI 1.02–1.05]	1.49 [95% CI 1.39–1.59]	1.63 [95% CI 1.45–1.83]	2.96 [95% CI 2.26–3·87]	1.41 [95% CI 1.11–1.81]

Some progress has been noted. Since its introduction in December 2020, the UK vaccination program has helped to decrease deaths from COVID-19 in the majority of ethnic minority groups. This is important, since the risk of death from COVID-19 is more than 90% lower for individuals who have received a third vaccine dose than for unvaccinated subjects. Nevertheless, vaccine roll-out remains the lowest in ethnic minority groups with the highest risk of COVID-19 death in all three pandemic waves [51]. See Figure 1.

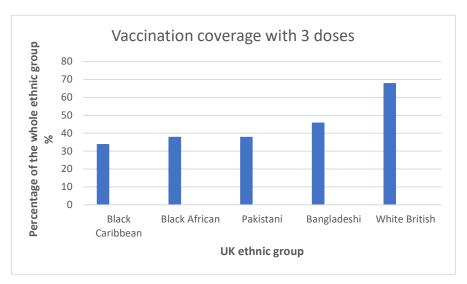


Figure 1. In the UK, only 34% of Black Caribbean, 38% of Black African, 38% of Pakistani, and 46% of Bangladeshi adult people have had three vaccine doses in comparison to 68% of White British adults [51]. White British vs. Black Caribbean p < 0.0001. White British vs. Black African p < 0.0004. White British vs. Pakistani p < 0.0004. White British vs. Bangladeshi p < 0.0006.

There are many stories about inequities in the distribution of COVID-19 vaccines. According to a report by the Centers for Disease Control and Prevention in February 2022, "of 42 million people receiving at least the first vaccine dose, only 6.3% were African Americans" [52].

Emerging statistics have proved without any doubt that COVID-19 disproportionately affects African Americans the most. The effects of COVID-19 on this specific population are inextricably related with four areas of systemic oppression and weakness, which are further exacerbated by COVID-19 itself: (1) healthcare inequality; (2) segregation, overall health, and food insecurity; (3) underrepresentation in government and the medical profession; and (4) inequalities in participatory democracy and public engagement. To overcome this no longer acceptable lack of equity, specific planned interventions are needed; for example, the development of a national, standardized database to monitor the demographic backgrounds of people suffering from COVID-19 with the aim of helping the American government to equitably and efficiently distribute the available human, economic, and medical resources; creating a partnership with nonprofit organizations, those run by African Americans and/or serving mostly African American people included; and developing specific strategies to safeguard the ability of African Americans to vote. This includes, but it is not limited to, expanding opportunities for them to register and vote remotely, as well as adopting and extending paid time off for voting for those involved in low-skill professions; addressing digital inequality to allow African Americans to gain access to potentially life-saving information, which is needed to mitigate the spread of COVID-19 [53].

9. Mental Health Issues as a Part of the Syndemic

The COVID-19 syndemic is accompanied and exacerbated by mental health issues in patients, family members, and healthcare workers themselves [54]. These mental health issues include anxiety, depression, and suicide. Mental health problems may arise from direct infection and inflammation of the brain from COVID-19 and/or concomitant vasculitis and thrombosis of blood vessels in the central nervous system. Psychological problems can also result from fear of the disease, proximity to death and dying, confinement in small spaces, and isolation. People who suffer from a mental health disease seem to have worse outcomes after COVID-19 infection compared to non-affected populations. There is a need for more studies in this area and for clinical trials to lessen and prevent the impact of these conditions, especially in the setting of long COVID-19 infection and in children [54]. Nonetheless, there is no doubt that the tighter the COVID-19-related restrictions, the poorer the mental health. The likelihood of mental issues related to the situation of emergency, such as anxiety, depression, post-traumatic stress disorder, and sleep disorders, is significantly increased in healthcare workers, mostly frontline workers, migrant workers, and workers in contact with the public. On the other hand, the risk of losing their job, long periods of quarantine, and uncertainty concerning the future make psychological conditions worse, especially in the youth and in people with a higher level of education [55].

10. Conclusions

The current COVID-19 syndemic/pandemic has highlighted chronic disparity in healthcare systems. Historically the highest rates of infection and mortality have been among the poorest and socially disadvantaged people around the world. COVID-19 morbidity and mortality are no exception. COVID-19 has revealed already existing social/economic/political discriminations in access to healthcare, vaccine administration, and monoclonal antibodies therapies, particularly in individuals affected by non-communicable diseases such as cancer and cardiovascular disease. COVID-19 has already dramatically reduced life expectancy in this subset of subjects [56].

The COVID-19 pandemic has made previous inequities clearer than ever. Three population groups, based on their historical inequities, can be identified: "The last" low- and middle-income countries whose healthcare systems run the risk of collapsing under the pressures of the pandemic; "The lost" refugees and migrants who have been left in even more precarious position than before the pandemic; and "The least" minority populations,

who are further prone to new detrimental socioeconomic consequences as a result of COVID-19 exacerbating the pattern of inequalities [57]

In addition, culture is crucial since it shapes how people react to crises such as the current COVID-19 pandemic. Understanding cultural differences not only provides a better understanding of the current pandemic, but also helps in preparing for future crises [19].

Without a strategic plan and a new anthropological *syndemic* approach to future pandemics, more and more people worldwide will unnecessarily suffer, health inequities will widen, and the world economies will slow down [58]. As a take home message learned from this global crisis, we should try our best to make it easier for everyone to access vaccines and other officially accepted therapies, and health in general. The distribution of limited and, at the same time, life-saving resources, such as vaccines, intensive care beds, and ventilators, should be global, equal, and fair. Any derogation from that is no longer acceptable in terms of ethics [59].

Funding: This research received no external funding.

Institutional Review Board Statement: Not applicable.

Informed Consent Statement: Not applicable.

Data Availability Statement: Not applicable.

Conflicts of Interest: The authors declare no conflict of interest.

References

1. Casti, G.; Bassareo, P.P.; Limone, M.; Pistolesi, F.; Fanos, V.; Marcialis, M.A. Epidemiology of SARS-CoV-2: Numbers matter! *J. Pediatr. Neonat. Individ. Med.* **2020**, *9*, e090203.

- 2. Singer, M.; Bulled, N.; Ostrach, B.; Mendenhall, E. Syndemics and the biosocial conception of health. Lancet 2017, 389, 941–950.
- 3. Singer, M. Introduction to Syndemics: A Critical System Approach to Public and Community Health; Wiley: Hoboken, NJ, USA, 2009; p. 304.
- 4. Jakovljevic, M.; Samarzija, M.; Milicic, D.; Reiner, Z.; Sartorius, N. Comorbidities and syndemicsin the COVID-19 Age: Challenges and opportunities for bringing separated branches of medicine closer to each other. *Psychiatr. Danub.* **2021**, *33*, 402–413.
- 5. Tsai, A.C.; Mendenhall, E.; Trostle, J.A.; Kawac, A. Co-occurring epidemics, syndemics, and population health. *Lancet* 2017, 389, 978–982.
- 6. Singer, M. A dose of drugs, a touch of violence, a case of AIDS: Conceptualizing the SAVA syndemic. *Free Inq. Creat. Sociol.* **2000**, *28*, 13–24.
- Niessen, L.W.; Mohan, D.; Akuoku, J.K.; Mirelman, A.J.; Ahmed, S.; Koehlmoos, T.P.; Trujillo, A.; Khan, J.; Peters, D.H. Tackling socioeconomic inequalities and non-communicable diseases in low-income and middle-income countries under the Sustainable Development agenda. *Lancet* 2018, 391, 2036–2046.
- 8. Kamrath, C.; Rosenbauer, J.; Eckert, A.J.; Siedler, K.; Bartelt, H.; Klose, D.; Sindichakis, M.; Herrlinger, S.; Lahn, V.; Holl, R.W. Incidence of type 1 diabetes in children and adolescents during the COVID-19 pandemic in Germany: Results from the DPV registry. *Diabetes Care* 2022, 17, dc210969.
- 9. Chen, J.T.; Krieger, N. Revealing the unequal burden of COVID-19 by income, race/ethnicity, and household crowding: US county versus zip code analyses. *J. Public Health Manag. Pr.* **2021**, *27*, S43–S56.
- 10. Stephenson, J. Racial, ethnic disparities found in receiving monoclonal antibodies for COVID-19. *JAMA Health Forum* **2022**, *3*, e220115.
- 11. Horton, R. Offline: COVID-19 is not a pandemic. Lancet 2020, 396, 874.
- 12. Tsai, A.C. Syndemics: A theory in search of data or data in search of a theory? Soc. Sci. Med. 2018, 206, 117–122.
- 13. Bassareo, P.P.; Melis, M.R.; Marras, S.; Calcaterra, G. Learning from the past in the COVID-19 era: Rediscovery of quarantine, previous pandemics, origin of hospitals and national healthcaresystems, and ethics in medicine. *Postgrad. Med. J.* **2020**, *96*, 633–638.
- 14. Sims, S.; Harris, R.; Hussein, S.; Rafferty, A.M.; Desai, A.; Palmer, S.; Brearley, S.; Adams, R.; Rees, L.; Fitzpatrick, J.M. Social Distancing and Isolation Strategies to Prevent and Control the Transmission of COVID-19 and Other Infectious Diseases in Care Homes for Older People: An International Review. *Int. J. Environ. Res. Public Health* 2022, 19, 3450.
- 15. Aspachs, O.; Durante, R.; Graziano, A.; Mestres, J.; Reynal-Querol, M.; Montalvo, J.G. Tracking the impact of COVID-19 on economic inequality at high frequency. *PLoS ONE* **2021**, *16*, e0249121.
- 16. Kansiime, M.K.; Tambo, J.A.; Mugambi, I.; Bundi, M.; Kara, A.; Owuor, C. COVID-19 implications on household income and food security in Kenya and Uganda: Findings from a rapid assessment. *World Dev.* **2021**, *137*, 105199.

17. Schomaker, R.M.; Bauer, M.W. What Drives Successful Administrative Performance During Crises? Lessons from Refugee Migration and the Covid-19 Pandemic. *Public Adm. Rev.* **2020**, *80*, 845–850. https://doi.org/10.1111/puar.13280.

- 18. Chen, C.; Frey, C.B.; Presidente, G. Culture and contagion: Individualism and compliance with COVID-19 policy. *J. Econ. Behav. Organ.* **2021**, *190*, 191–200.
- 19. Lu, J.G.; Jin, P.; English, A.S. Collectivism predicts mask use during COVID-19. Proc. Natl. Acad. Sci. USA 2021, 118, e2021793118.
- 20. Porcher, S. Culture and the quality of government. Public Adm. Rev. 2019, 81, 333–343.
- 21. Gadarian, S.K.; Goodman, S.W.; Pepinsky, T.B. Partisanship, health behavior, and policy attitudes in the early stages of the COVID-19 pandemic. *PLoS ONE* **2021**, *16*, e0249596.
- 22. Porcher, S.; Renault, T. Social distancing beliefs and human mobility: Evidence from Twitter. PLoS ONE 2021, 16, e0246949.
- 23. Duhon, J.; Bragazzi, N.; Kong, J.D. The impact of non-pharmaceutical interventions, demographic, social, and climatic factors on the initial growth rate of COVID-19: A cross- country study. Sci. Total Environ. 2021, 760, 144325.
- 24. Forni, G.; Mantovani, A. COVID-19 Commission of Accademia Nazionale dei Lincei, Rome. COVID-19 vaccines: Where we stand and challenges ahead. *Cell Death Differ.* **2021**, *28*, 626–639.
- 25. Watad, A.; De Marco, G.; Mahajna, H.; Druyan, A.; Eltity, M.; Hijazi, N.; Haddad, A.; Elias, M.; Zisman, D.; Naffaa, M.E.; et al. Immune-Mediated Disease Flares or New-Onset Disease in 27 Subjects Following mRNA/DNA SARS-CoV-2 Vaccination. *Vaccines* **2021**, *9*, 435.
- 26. Rosselli, R.; Martini, M.; Bragazzi, N.L. The old and the new: Vaccine hesitancy in the era of theWeb 2.0. Challenges and opportunities. *J. Prev. Med. Hyg.* **2016**, *57*, E47–50.
- 27. Kricorian, K.; Civen, R.; Equils, O. COVID-19 vaccine hesitancy: Misinformation and perceptions of vaccine safety. *Hum. Vaccin Immunother.* **2022**, *18*, 1950504.
- 28. Truong, J.; Bakshi, S.; Wasim, A.; Ahmad, M.; Majid, U. What factors promote vaccine hesitancyor acceptance during pandemics? A systematic review and thematic analysis. *Health Promot. Int.* **2022**, *37*, daab105.
- 29. Thacker, P.D. Covid-19, Researcher blows the whistle on data integrity issues in Pfizer's vaccine trial. BMJ 2021, 375, n2635.
- 30. Calcaterra, G.; Mehta, J.L.; de Gregorio, C.; Butera, G.; Neroni, P.; Fanos, V.; Bassareo, P.P. COVID 19 Vaccine for Adolescents. Concern about Myocarditis and Pericarditis. *Pediatr. Rep.* **2021**, *13*, 530–533.
- 31. Calcaterra, G.; Bassareo, P.P.; De Gregorio, C.; Barilla, F.; Romeo, F.; Mehta, J.L. COVID-19 Vaccine-Induced Pro-thrombotic Immune Thrombocytopenia (VIPIT): State of the art. Curr. Cardiol. Rev. 2022, 18, e210322202448. Epub ahead of print. https://doi.org/10.2174/1573403X18666220321105909.
- 32. Xu, Y.; Li, J.-P.; Chu, C.-C.; Dinca, G. Impact of COVID-19 on transportation and logistics: A case of China. *Econ. Res. Ekon. Istraživanja* **2021**, *13*, 251–257.
- 33. Tagoe, E.T.; Sheikh, N.; Morton, A.; Nonvignon, J.; Sarker, A.R.; Williams, L.; Megiddo, I. COVID- 19 Vaccination in Lower-Middle Income Countries: National Stakeholder Views on Challenges, Barriers, and Potential Solutions. *Front. Public Health* **2021**, *9*, 709127.
- 34. Available online: https://africacdc.org (accessed on 23 May 2022).
- 35. Wagner, C.E.; Saad-Roy, C.M.; Morris, S.E.; Baker, R.E.; Mina, M.J.; Farrar, J.; Holmes, E.C.; Pybus, O.G.; Graham, A.L.; Emanuel, E.J.; et al. Vaccine nationalism and the dynamics and control of SARS-CoV-2. *Science* **2021**, *373*, eabj7364.
- 36. Fischer, W.A., 2nd; Wohl, D.A. Inequities in access to diagnostics threatens global public health security. *Lancet Infect. Dis.* **2022**, 22, 754–756. https://doi.org/10.1016/S1473-3099(21)00806-9.
- 37. Calcaterra, G.; Mehta, J.L.; Fanos, V.; Bassareo, P.P. Insights on Kawasaki disease and multisystem inflammatory syndrome: Relationship with COVID-19 infection. *Minerva Pediatr.* **2021**, *73*, 203–208.
- 38. Porcher, S. Response2covid19, a dataset of governments' responses to COVID-19 all around the world. Sci. Data 2020, 7, 423.
- 39. Available online: https://www.who.int/news-room/fact-sheets/detail/noncommunicablediseases (accessed on 20 January 2022).
- 40. Rudd, K.E.; Mair, C.F.; Angus, D.C. Applying Syndemic Theory to Acute Illness. JAMA 2022, 327, 33–34.
- 41. You, J.; Zhang, J.; Li, Z. Consumption-Related Health Education Inequality in COVID-19, A Cross-Sectional Study in China. Front. Public Health 2022, 10, 810488.
- 42. Dávalos, L.M.; Austin, R.M.; Balisi, M.A.; Begay, R.L.; Hofman, C.A.; Kemp, M.E.; Lund, J.R.; Monroe, C.; Mychajliw, A.M.; Nelson, E.A.; et al. Pandemics' historical role in creating inequality. *Science* **2020**, *368*, 1322–1323.
- 43. Calcaterra, G.; Bassareo, P.P.; Barillà, F.; Sergi, D.; Chiocchi, M.; Romeo, F.; Mehta, J.L. The DeadlyQuartet (Covid-19, Old Age, Lung Disease, and Heart Failure) Explains Why Coronavirus- Related Mortality in Northern Italy Was So High. *Curr. Cardiol. Rev.* 2021, 17, 74–77.
- 44. Calcaterra, G.; Bassareo, P.P. A brief overview about the latest news on SARS-CoV-2 therapeutic management. *Curr. Top. Pharmacol.* **2021**, 25, 21–24.
- 45. Wiltz, J.L.; Feehan, A.K.; Molinari, N.M.; Ladva, C.N.; Truman, B.I.; Hall, J.; Block, J.P.; Rasmussen, S.A.; Denson, J.L.; Trick, W.E.; et al. Racial and Ethnic Disparities in Receipt of Medications for Treatment of COVID-19–United States, March 2020-August 2021. MMWR Morb. Mortal. Wkly. Rep. 2022, 71, 96–102.
- 46. Available online: https://covid.cdc.gov/covid-data-tracker/#datatracker-home. (accessed on 15 March 2022).
- 47. Lin, Q.; Paykin, S.; Halpern, D.; Martinez-Cardoso, A.; Kolak, M. Assessment of Structural Barriers and Racial Group Disparities of COVID-19 Mortality with Spatial Analysis. *JAMA Netw. Open* **2022**, *5*, e220984.
- 48. Levy, B.L.; Vachuska, K.; Subramanian, S.V.; Sampson, R.J. Neighborhood socioeconomic inequality based on everyday mobility predicts COVID-19 infection in San Francisco, Seattle, and Wisconsin. *Sci. Adv.* **2022**, *8*, eabl3825.

- 49. Available online: https://www.amazon.com/Heart-Disease-Your-Head-about/dp/B09NW8RTK7 (accessed on 18 March 2022).
- 50. Mathur, R.; Rentsch, C.T.; Morton, C.E.; Hulme, W.J.; Schultze, A.; MacKenna, B.; Eggo, R.M.; Bhaskaran, K.; Wong, A.Y.S.; Williamson, E.J.; et al. Ethnic differences in SARS-CoV-2 infection and COVID-19-related hospitalisation, intensive care unit admission, and death in 17 million adults in England: An observational cohort study using the OpenSAFELY platform. *Lancet* **2021**, 397, 1711–1724.
- 51. Raleigh, V.S. Ethnic differences in covid-19 death rates. BMJ. 2022, 376, o427.
- 52. Available online: https://www.cdc.org. (Accessed on 1 March 2022).
- 53. Wright, J.E., 2nd; Merritt, C.C. Social Equity and COVID-19: The Case of African Americans. *Public Adm. Rev.* **2020**, *80*, 820–826. 10.1111/puar.13251. https://doi.org/10.1111/puar.13251.
- 54. Magnúsdóttir, I.; Lovik, A.; Unnarsdóttir, A.B.; McCartney, D.; Ask, H.; Kõiv, K.; Christoffersen, L.A.N.; Johnson, S.U.; Hauksdóttir, A.; Fawns-Ritchie, C.; et al. Acute COVID-19 severity and mental health morbidity trajectories in patient populations of six nations: An observational study. *Lancet Public Health* **2022**, 7, e406–e416.
- 55. Giorgi, G.; Lecca, L.I.; Alessio, F.; Finstad, G.L.; Bondanini, G.; Lulli, L.G.; Arcangeli, G.; Mucci, N. COVID-19-Related Mental Health Effects in the Workplace: A Narrative Review. *Int. J. Environ. Res. Public Health* **2020**, *17*, 7857.
- 56. Kuhen, B.M. COVID-19 cuts life expectancy in dozen of countries. JAMA 2002, 327, 209.
- 57. Tan, M.K.I. COVID-19 in an inequitable world: The last, the lost and the least. Int Health 2021, 13, 493-496.
- 58. Emanuel, E.J.; Osterholm, M.; Gounder, C.R. A national strategy for the "new normal" of life with COVID. *JAMA* **2022**, 327, 211–212
- 59. Moodley, K. Vaccine inequity is unethical. Nat. Hum. Behav. 2022, 6, 168–169.