

Quantifying Effects, Forecasting Releases, and Herd Immunity of the Covid-19 Epidemic in S. Paulo – Brazil

Sergio Celaschi

CTI Renato Archer, Rod. D Pedro I (SP-65), Km 143,6 - CEP 13069-901, Campinas – SP, 13069, Brazil

Corresponding Author: Sergio.celaschi@cti.gov.br

ABSTRACT

A simple and well known epidemiological deterministic model was selected to estimate future outcomes for the 2020 dynamics of the Covid-19 epidemic breakout in the city of São Paulo – Brazil with a population of 12.2 million inhabitants. A time-dependent incidence weight on the reproductive basic number accounts for modeling social distancing, and local Mitigation Policies (MP). Official published data from January 1st to July 7th, 2020 was used to adjust all the model parameters aiming to forecast the evolution of the COVID-19 epidemic outbreak. Data points for the last seven weeks, prior July 7th, were added after the model was complete, granting confidence on the outcomes. The insights gained from analysis of these successful interventions were used to quantify shifts and reductions on active cases, casualties, and estimative on required medical facilities (ICU). By beginning November 2020, the predicted numbers of symptomatic infected individual in the city of S. Paulo may reach 200 thousands and fatalities would accounts for 18 thousands. The total number of non notified infected hosts was considered to be twice the number of notified symptomatic infected. The analysis was applied to forecast the consequences of releasing the MP over specific periods of time. Herd Immunity (HI) analysis allowed estimating how far we are from reaching the HI threshold value, and the price to be paid. This knowledge can be applied to other Brazilian areas.

KEYWORDS; COVID-19; Brazil; epidemiological model; mitigation policy; social distancing; herd immunity.

Date of Submission: 27-07-2020

Date of Acceptance: 11-08-2020

I. INTRODUCTION

This work aims to shed some extra light in understanding the dynamics of the COVID-19 pandemic in Brazil, particularly in the city area of São Paulo with its 12.2 million inhabitants. The first patient in Brazil was tested positive in São Paulo who had returned from Italy. Since then, were officially confirmed 1,668,589 cases, and 66,741 deaths in Brazil (July, 07 2020), and 142,502 cases, and 7,743 deaths in S. Paulo city. The public response to the pandemic has been the introduction of mitigation policies to ensure quarantine social distancing, such as closing schools, restricting commerce, and home office. No country knows the true number of people infected with COVID-19. All is known is the infection status of those who have been tested. The total number of people that have tested positive – the number of confirmed cases – is not the total number of people who have been infected. The true number of people infected with COVID-19 is much higher. The number of those who have been tested positive (by May 1st) in Brazil, 0.46/thousand, is very low compared to other countries [1]. The response to a new pandemic, such as Covid-19, can be based on four major actions: 1) surveillance and detection; 2) clinical management of cases; 3) prevention of the spread in the community; and 4) maintaining essential services. Actions across the four pillars complement and support one another. In principle if the virus is left to infect people without any containment measure, the population may acquire immunity in one semester or less. However, hospital intensive care services would lack the capacity to deal with the sudden, large inflow of severely ill people resulting in a very large number of deaths. Mitigation strategies aim to slow the disease, and to reduce the peak in health care demand. This includes policy actions such as social distancing, lock-down determination, and improved personal and environmental hygiene. Studies as this presented here, consistently conclude that packages of containment and mitigation measures are now days an effective approach to reduce the impact of the Covid-19 epidemic. Epidemiological models are commonly stochastic, diffusive-spatial, network based, with heterogeneous sub-populations (meta-population approaches) [2, 3, 4]. However, the parameters of dynamical and deterministic models, such as SIR and SEIR, are more directly related to and interpretable as physical processes [5]. On the other hand, deterministic models impose restrictive analysis, once the dynamics of the host population and the virus are not deterministic. The population has free will, and the virus undergoes “random” mutations [6].

The intent of this work was to build a simple epidemiological tool to estimate the main results for the basic dynamics of the Covid-19 epidemic breakout. The methodology employed is the application of the

deterministic and discrete SEIRD Model to characterize the Covid-19 outbreak in São Paulo – Brazil. The model accounts for the following 5 groups: Susceptible, Exposed, Infected, Recovered, and Deaths. A time-dependent incidence weight on the SEIRD basic reproductive number $R(t)$ was used to account for dynamical transmission behavior, to model and quantify local Mitigation Policies (MP). The insights gained from analysis of these successful interventions can be used to predict results for the MP of other Brazilian regions.

II. MODELING THE COVID-19 EPIDEMIC

Recent published data [7, 8] from January, 1st, 2020 to July 7, 2020 was used to adjust all the model parameters aiming to forecast the evolution of the COVID-19 epidemic in the city of Sao Paulo - Brazil. The model provides predictions of the time series of infected individuals and fatalities in the area studied. Simulations of mid-term scenarios of the epidemic outbreak were done dependent on the level of confinement policies. Forecasts show how confinement policy alters the pattern of contamination, and suggest the existence of post epidemic periods. Application of mathematical models to disease surveillance data can be used to address both scientific hypotheses and disease-control policy questions [9]. Such models have been used to estimate the demand for hospital beds, ICU days, number of critical equipments, and the need and required extent of governmental intervention. A model is only as good as the assumptions put into it. Clearly, there are phenomena of the COVID-19 epidemic that are not yet understood. Models are constantly being updated and improved.

The SARS-CoV-2 is a membrane protected, single-stranded RNA virus. It is commonly referred to by the name of the disease it causes, which is COVID-19. The incubation period is defined as the time between infection and onset of symptoms. It is estimated as the time between exposure and report of noticeable symptoms. Currently, the incubation period for COVID-19 is somewhere between 2 to 5 days after exposure. More than 97 percent of people who contract SARS-CoV-2 show symptoms within 12 days of exposure. For many people, COVID-19 symptoms start as mild symptoms and gradually get worse over a few days. Other large and unknown fraction of exposed people is asymptomatic. The story of the COVID-19 outbreak is ongoing. Our knowledge of this novel virus is in a state of flow. Every week seems to bring additional important medical and epidemiological information. COVID-19 has a latent or incubation period, during which the individual is said to be infected but not infectious. Members of this population in this latent stage are labeled as Exposed (but not infectious) here on. The model with the Susceptible, Exposed, Symptomatic, Asymptomatic, and Removed groups is known as SEIR Model. During the initial 20 days exponential phase of growth, the SP data showed that in 2.3 ± 0.1 days the number of symptomatic infected people doubled. During this initial exponential period, the model confirms the predicted values for the incubation, and immunization periods of time.

SIR, SEIR, and SEIRD Models

The SIR model [5] is one of the simplest compartmental models, and many other models are created from this basic formulation. The model consists of three compartments: S for the number of susceptible, I = $I_s + I_a$ for the number of infectious (symptomatic, I_s and asymptomatic, I_a) or active cases, and R for the number of recovered, deceased (or immune) individuals. This model is reasonably predictive for infectious diseases that are transmitted from human to human. In epidemics or pandemic outbreaks, the numbers of susceptible, infected and recovered individuals varies with time (even if the total population size remains constant). For a specific disease in a specific population, these functions may be worked out in order to predict possible outbreaks and bring them under control. For many important viral infections, there is a significant incubation period during which individuals have been infected but are not yet infectious themselves. During this period the individual is in a new compartment E (for exposed) as in the SEIR model [5].

At the initial exponential outbreak, let β_0 be the average number of contacts (per unit time) multiplied by the probability of transmission from an infected person. Let $\beta(t) = f[\beta_0, \psi(t)]$ be a selected function that models temporal Mitigation Policies (MP). Let $\psi(t)$ quantify the MP in terms of social distancing plus the use of personal protective equipment (PPE). Let σ be the rate that exposed individuals get infected. Let γ be the removed rate that infected individuals (symptomatic and asymptomatic) recover, leaving the infected groups, at constant per capita probability per unit of time. Let μ be the removed rate that infected individuals die. Let ξ being the fraction of symptomatic individuals. Let N be the susceptible population considered in the study. Worth to mention that by the time this study was conducted, estimative on the number of sub notifications and information on the number of asymptomatic infected individuals in the city of S. Paulo were scarce. Under those assumptions, the set of ordinary differential equations leads to our SEIRD model;

$$\frac{dS(t)}{dt} = -\beta(t).I(t).\frac{S(t)}{N} \tag{1}$$

$$\frac{dE(t)}{dt} = \beta(t).I(t).\frac{S(t)}{N} - \sigma_o.E(t) \tag{2}$$

$$\frac{dI(t)}{dt} = (\gamma + \mu).I(t) + \sigma_o.E(t) \tag{3}$$

$$\frac{dR(t)}{dt} = \gamma.I(t) \tag{4}$$

$$\frac{dF(t)}{dt} = \mu.I(t) \tag{5}$$

where $S(t)$, $E(t)$, $I(t)$, $R(t)$, and $F(t)$ are respectively daily numbers of Susceptible, Exposed, Infected or active cases, Removed or recovered individuals, and Fatalities (deaths). $S(t)+E(t)+I(t)+R(t)+F(t)=N=Constant$. The constant N assumption is very restrictive, and limits model’s coverage. Releasing this assumption goes beyond the scope of this study. $R(t)=\beta(t)/\gamma$. $R(t=0)=R_o$, is defined as the Basic Reproductive Number. This number quantifies the expected number of new infections (these new infections are sometimes called secondary infections) that arise from a typical primary case in a completely susceptible population N , where all individuals are susceptible.

Methodology

Official data, from March 1st to Jul 7th, 2020, provided by the Ministry of Health of Brazil [7], and Sao Paulo government [8] was considered to estimate part of the epidemiological parameters that govern the dynamics established by Eqns. (1) to (5). By the lack of information about the asymptomatic individuals, the mortality rate in the model is evaluated over the symptomatic ones. All model parameters were estimated by minimizing the mean squared quadratic errors. A key parameter in deterministic transmission models is the reproductive number R , which is quantified by both, the pathogen and the particular population in which it circulates. Thus, a single pathogen, like the SARS-CoV-2, will have different R values depending on the characteristics and transmission dynamics of the population experiencing the outbreak. When infection is spreading through a population that may be partially immune, it has been suggested to use an effective reproductive number R , defined as the number of secondary infections from a typical primary case. Accurate estimation of the R value is crucial to plan and control an infection [10]. The methodology to estimate R follows: The exponential growth rate of the epidemic, r was obtained from the early stages of the epidemic in Sao Paulo, such that the effect of MP discussed later will be relative to post stages of this outbreak. This assumption is implicit in many estimative of R_o . The growth rate $r = 0.31 \pm 0.02$ of infected people was estimated applying the Levenberg-Marquardt method [11], to data of symptomatic infected people (Fig. 1), during the first 15 days of exponential growth according to the expression $I(t) = I_o.exp(-r.t)$.

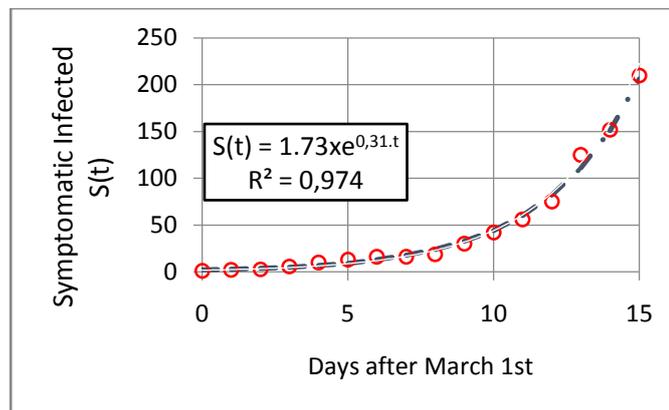


Figure 1: Exponential fitting of the initial growth of symptomatic infect people. The growth rate $r = 0.31 \pm 0.02$ was estimated applying the Levenberg-Marquardt method.

The time period required to double the number of symptomatic cases is straightforward given by $\ln(2)/r = 2.3 \pm 0.1$ days. The initial basic Reproductive Number $R_o = 2.53 \pm 0.09$ (restricted to symptomatic cases) was estimated according to; “In an epidemic, driven by human-to-human transmission, whereas growing exponentially, in a deterministic manner, the incidence $I(t)$ can be described by the Renewal Equation”, or the Lotka–Euler Equation [12, 13]:

$$I(t) = \int_0^\infty I(t - \tau)\beta(\tau)d\tau \tag{6}$$

Where $\beta(\tau)$ is the mean rate at which an individual infects others a time after being infected itself. Substituting into Eqn. (6) an exponentially growing incidence, $I(t) = I_0 \cdot \exp(r \cdot t)$, $I_0 = 1$, gives the condition,

$$1 = \int_0^{\infty} e^{-\tau \cdot r} \beta(\tau) d\tau \tag{7}$$

where

$$\beta(\tau) = R_0 \cdot \omega(\tau) \tag{8}$$

$\omega(\tau)$ is the generation time distribution, i.e. the probability density function for the time between an individual becoming infected and their subsequent onward transmission events. R_0 is the basic reproduction number. If the exponential growth rate r and the generation time distribution $\omega(\tau)$ have been estimated, R_0 is readily determined from Eqn. (7), as

$$R_0^{-1} = \int_0^{\infty} e^{-r \cdot \tau} \cdot \omega_n(\tau, \lambda, \kappa) \cdot d\tau \tag{9}$$

The term that appears in the right-hand side of Eqn. (9) is the Laplace transform of the integrand function. More specifically, it is known as the Momentum Generating Function of this distribution. As in [12], a normalized Weibull generation distribution is adopted (Eqns. 10, 11 and 12), with mean = 0.89, median = 3.1, and peak value at 2.3 days (Fig. 2).

$$\omega_n(\tau, \lambda, \kappa) = \frac{\omega(\tau, \lambda, \kappa)}{I_\omega} \tag{10}$$

$$\omega_n(\tau, \lambda, \kappa) = \left(\frac{\kappa}{\lambda}\right) \cdot \left(\frac{\tau}{\lambda}\right)^{(\kappa-1)} \cdot e^{-\frac{\tau \cdot \kappa}{\lambda}} \tag{11}$$

$$I_\omega = \int_0^{\infty} \omega(\tau, \lambda, \kappa) \cdot d\tau \tag{12}$$

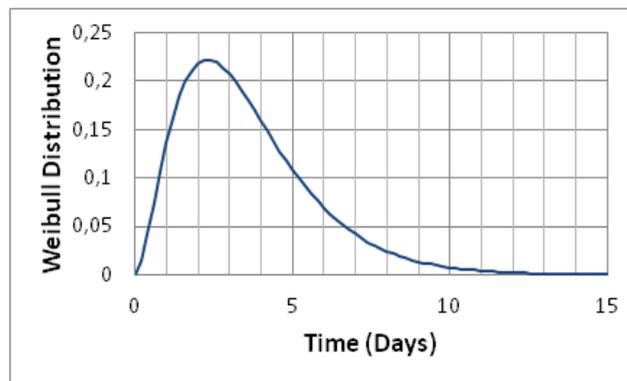


Figure 2: Distribution of generation times. Our data was described by the Weibull distribution with mean = 0.89, median = 3.1 days, and peak value 2.3 days. Dates of symptom onset with intervals of exposure for both source and recipient (when available) were collected in [13] in order to select the best distribution.

In short, the values of the parameters governing SEIRD model are: $\beta(t), \gamma, \sigma, \xi, \mu$, and N . The SEIRD dynamics is constrained to the following initial conditions $S(0) = N, E(0) > 0, I(0) = 0, R(0) = 0$. March 1st 2020 was considered the first day (day zero) to model the epidemic outbreak at the city of Sao Paulo. The temporal impact of the confinement policy was considered weighting the initial transmission factor β_0 by $\psi(t)$ fitted to data, and adjustable to allow releases of the MP (Fig. 3a) [7,8]. This leads to β factor as a temporal function $\beta(\psi_o, t_o, t_1, t_2, \varphi, t)$, where ψ_o, t_o, t_1, t_2 , and φ are values set by the MPs considered in this report.

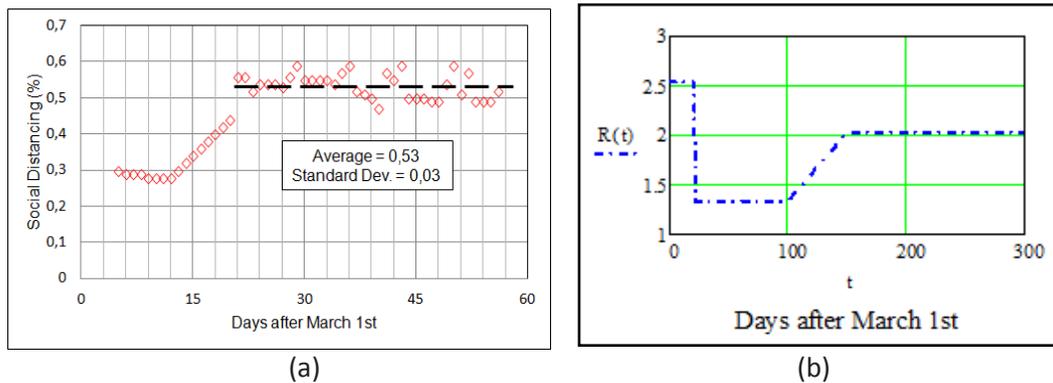


Figure 3: Quantification of the MP application on the city of S. Paulo by Social Distancing (SD) measures (a), Reproductive Basic Number $R(t)$ modeling the Social Distancing effective change after March 22nd (b). Blue point and dashed line represent the progressive return to R_0 after day 100.

Accordingly, $R(t) = \psi(t) \cdot R_0$ becomes dependent on the confinement policy (Fig. 3b). The effectiveness of this policy may be quantified by a social distancing factor defined as $SD = 1 - \psi(t)$. Furthermore, by lack of information about asymptomatic hosts, a value of 50% for the asymptomatic infected individuals ($\xi = 0.5$) was assumed [14].

III. RESULTS AND DISCUSSION

The fittings to data on symptomatic infected individuals, and fatalities to the SEIRD model are shown in Fig. 4. The fitting values are: $N = 5.10^5$ (4.1% of S. Paulo city population), $R_0 = 2.53$, $\beta_0 = 0.9$, $\gamma_0 = 0.3875$, $\psi_0 = 0.525$, $\mu = 0.017$, $\sigma_0 = 0.5$, and $t_1 = 22$. The fitting to the infected individuals (logarithmic scale) presents standard deviation $SD = 0.05$ and root mean square $RMS = 0.07$. The exponential rate of incubation $1/\sigma_0$ was assumed here as 2 days. Important to mention that the model was also applied assuming $1/\sigma_0 = 4$ days [12], and fixed values for $R_0 = 2.53$, $\psi_0 = 0.525$, $\mu = 0.017$, $\sigma_0 = 0.5$, and $t_1 = 22$. The new fittings to data, preserving R_0 ($\beta_0 = 2.445$, and $\gamma = 0.923$), led essentially to the same results.

Figure 4 shows reductions in exponential growths of infected and death rates after the third week. This reduction clarifies the effectiveness of the confinement policy, when social distancing took place. In fact, after day 22, the data shows reduction in the tax of transmission. So, day 22 was defined as the initial date to estimate the parameter ψ_0 , keeping all model parameters as previously estimated. The transmission rate $\beta(t)$ is reduced to approximately 52% of its original value β_0 , leading to a SD of 48% in accordance to the data shown in Fig. 3a. Keeping SD constant, the final average values for asymptomatic infected individuals and fatalities, at day 240, are respectively 110,000, and 9,800. Taking into consideration all the hosts, the average rate of lethality is $4.5 \pm 0.5\%$, in agreement to recent published data [15]. Maintaining a constant SD of 48% in the city of S. Paulo after two months was proved to be unfeasible. Shortening SD increases the initial 4.1% fraction of susceptible individuals to be exposed. The SD reduction in time was modeled (Fig.4 Inset), and taken into account in that figure as error bars.

Figure 5a presents data on the daily number of new symptomatic infected, compared to as predicted from modeling. The large data scattering on day to day-plus differences comes from the available reported cases. The number of Intensive Care Units (ICU) was estimated to be proportional to the weekly number of symptomatic infected.

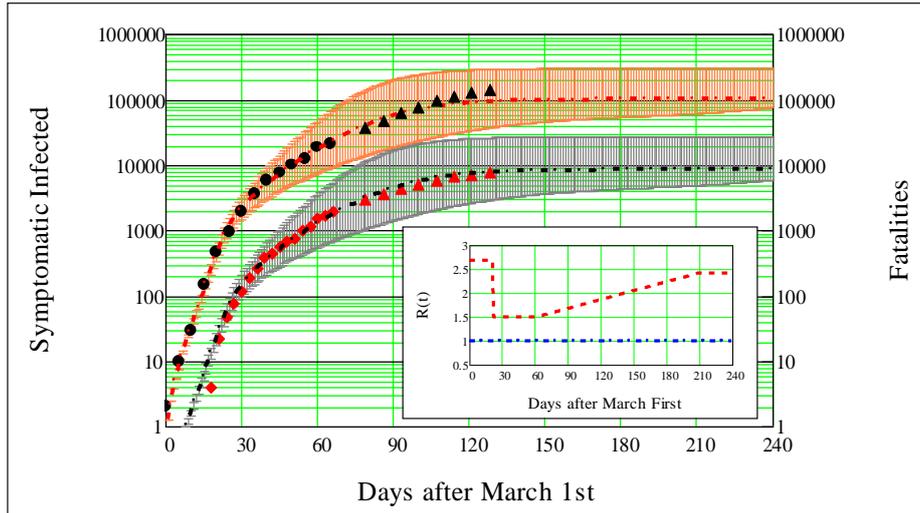


Figure 4: Primary official data on symptomatic infected individuals and fatalities are shown by black circles, and red diamonds respectively. Error bars account for the RMS values on the SEIRD parameters. The fitting values to SEIRD model are: $N = 5.10^5$, $R_0 = 2.53 \pm 0.09$, $\beta_0 = 0.980 \pm 0.012$, $\gamma = 0.387 \pm 0.0185$, $\mu = 0.017$, $\psi_0 = 0.525 \pm 0.025$, $\sigma_0 = 0.50 \pm 0.02$, and $t = 22$ days. Average values and error bars for infected individuals, and fatalities are shown in red, and black respectively. The mean fitting to the infected individuals (logarithmic scale) presents standard deviation $SD = 0.05$ and root mean square $RMS = 0.07$. Data points for the last seven weeks (solid triangles) were added after the model was complete, granting confidence on the outcomes. Inset shows the variation $R(t)$ as the result of the SD progressive reduction.

The proportionality constant in Eqn. (13) was determined as the ratio of occupied ICU beds (SARG - COVID-19) to the number of new infected during the 8th week post March 1st, where SARG = 8469 [7].

$$ICU(t) = \frac{SARG}{\int_{50}^{57} I(t).dt} \cdot \int_{t-7}^t I(t).dt \quad (13)$$

Prediction on the average weekly number of intensive care units (ICU) is also included in Fig. 5b by solid blue circles. Error bars account for the RMS values on the SEIRD parameters. The maximum weekly number of ICU beds for COVID-19 is predicted to reach $(1.6 \pm 0.8) \cdot 10^4$ by the third week of June, 2020.

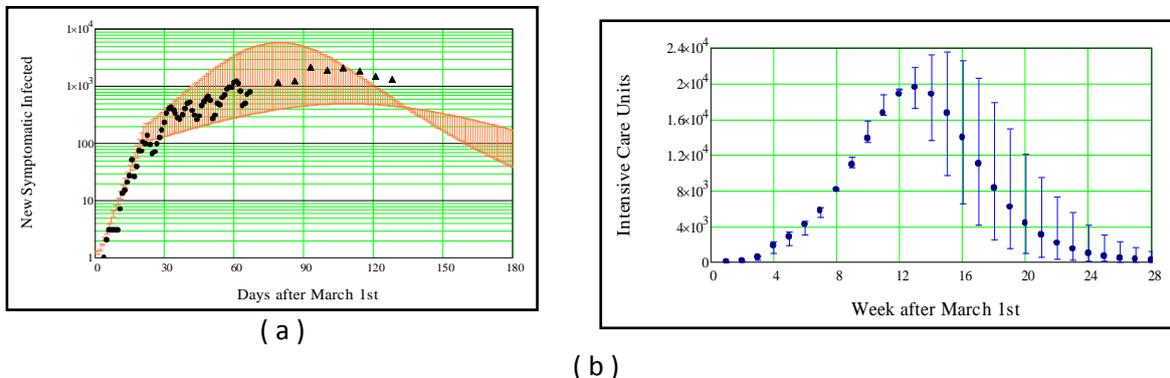
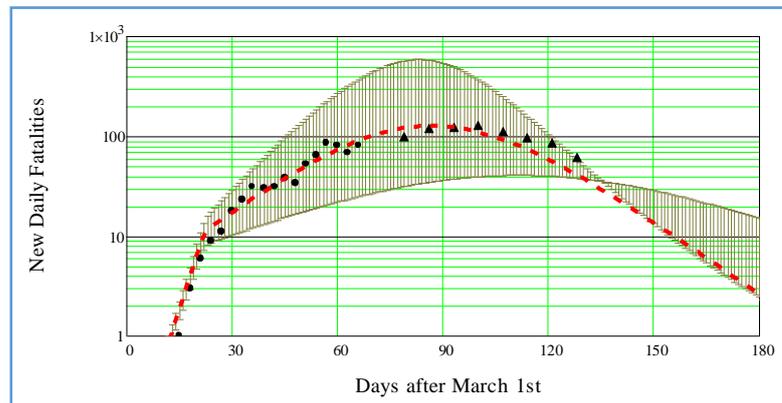


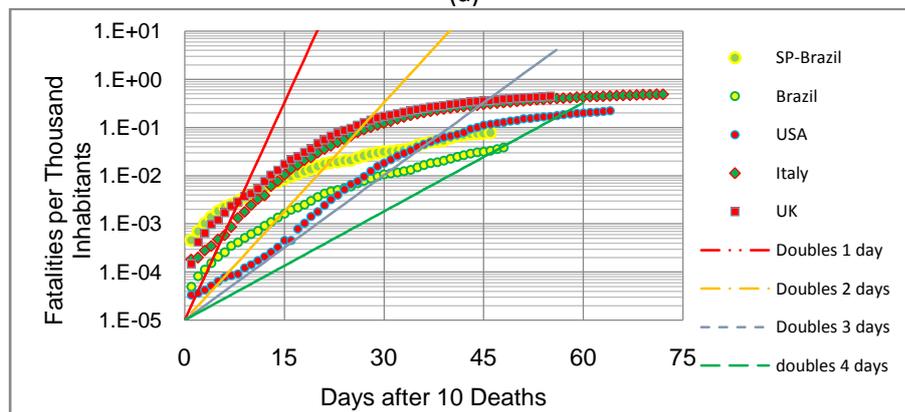
Figure 5: (a) Data (solid black circles) on the daily number of new symptomatic infected compared to predictions of the SEIRD model. Data points for the last seven weeks (solid triangles) were added after the model was complete. The large data scattering on day to day-plus differences comes from the available reported cases. (b) The number of Intensive Care Units (ICU) was estimated to be proportional to the weekly number of symptomatic infected. Permanence of SD after day 22nd was assumed. Error bars account for the RMS values on the SEIRD parameters.

Regarding fatalities, average data on daily new casualties is compare to model prediction on Fig.7a. Do to large data scattering; a seven day moving average was applied to data on Fig. 6a. Assuming the variation of $R(t)$ as already modeled for the SD progressive reduction (Fig. 4 Inset), the maximum number of new daily

fatalities is predicted to happen somewhere in between the last week of May, and the first week of June 2020. Data points for the last seven weeks (solid triangles), added after the model was complete, confirm this prediction. In order to compare this outbreak regarding other countries, we present in Fig 6b the number of fatalities per thousand inhabitants recently published to a few other countries [16].



(a)



(b)

Figure 6: (a) Data on daily new fatalities (solid black circles) is compare to model prediction. Keeping the MP as modeled after March 22nd. Dashed red line shows the average value. The RMS values to SEIRD model are shown by black error bars. The maximum number of new daily fatalities was predicted to happen between the last week of May, and first week of June, 2020. Data points for the last seven weeks (solid triangles) were added after the model was complete (b) Number of fatalities per thousand inhabitants reported from other countries. The fatality raise in the city of S. Paulo is shown for comparison.

The number of susceptible individuals $N = 500,000$, which represents 4.1% the population of Sao Paulo city, was considered to be the minimum number to fit preliminary available data on new symptomatic hosts. As early mentioned, the constant N assumption restricts our analysis, and forecasts. Since the number of sub notifications is a reality well accepted, we present in Table 1 suggested results folding N by a factor of three, but keeping fixed SD after March 22nd.

Table 1. Folding N by a factor of 3, but keeping fixed SD after March 22nd, peaks the active cases 2.5 weeks later, and raises the number of immune hosts, ITU units, and fatalities by the same factor. Peak Infection shown by the number of days after March 1st.

% Population	# Immunes ($\times 10^5$)	Peak Infection (day)	Max # ICU ($\times 10^4$)	# Fatalities ($\times 10^4$)
4.1	2.1	89	2.2	1.1
6.3	4.7	95	2.7	2.2
8.2	6.5	138	4.3	2.9
10.3	8.2	137	6.1	3.7
12.3	9.9	136	8.0	4.5

Quantifying the Impact of Social Distancing

In order to quantify the MP imposed at the city of Sao Paulo, a sequence of 3 plots presented by Fig. 7 demonstrate the effects of progressive releases on SD achieved by the city regulations. The sequence shows the daily numbers of additional symptomatic infected, and deaths. Social distancing of 48% was set as a constant from day 22nd up to day t_2 . Selected values for beginning a linear progressive SD release (t_2), ending it (t_{end}) to a final value of 24% are shown in Figs. 7a, 7b, and 9c. The sequence forecasts additional daily numbers of symptomatic infected (red line), and additional deaths (black line). The prospected accumulated additional fatalities for $t_2 = \{90, 100, 110\}$, and $t_{end} = t_2 + 30$ days are respectively 48, 36, and 32%.

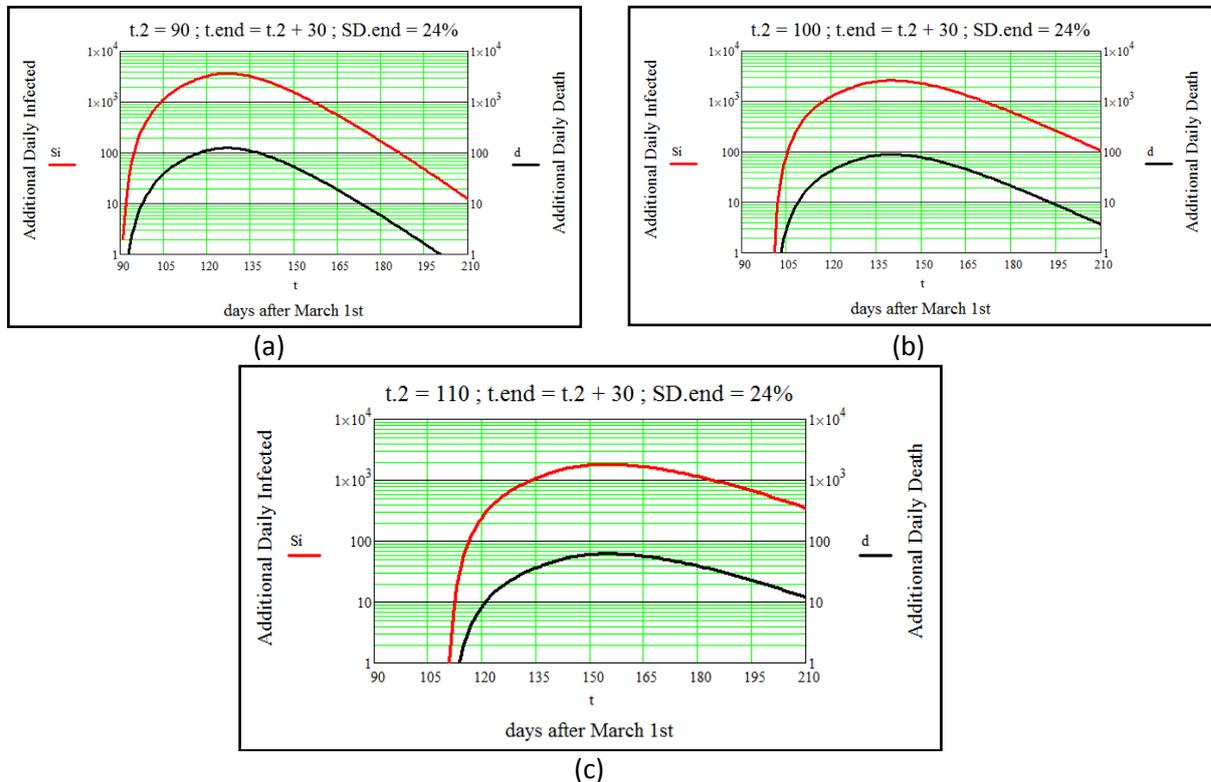


Figure 7: The sequence of figures demonstrates effects of releasing the 48% social distancing set constant from day - 22nd. Selected values: t_2 - beginning progressive SD release; t_{end} - end SD release, final SD = 24%. The sequence forecasts additional daily numbers of symptomatic infected (red line), and additional deaths (black line).

The analysis can also be applied to forecast the consequences of releasing SD over a longer period of time. Figure 8 illustrates this simulation by a linear and progressive SD release starting by the end of the third month, and ending ten months later, when the reproductive number returns to its initial value R_0 . Surprisingly or not, the model suggests an “endemic” outbreak of Covid-19 as shown in Fig. 8 by the presence of the second peak ~11 months after the first one. The number of infected individuals is estimated to be 55% lower compared to the first outbreak. This is a result of a partial reduction of 58% on the initial number of susceptible individuals. As recently published by US National Library of Medicine, National Institutes of Health, agencies worldwide prepare for the seemingly inevitability regarding the COVID-19, to become endemic [17].

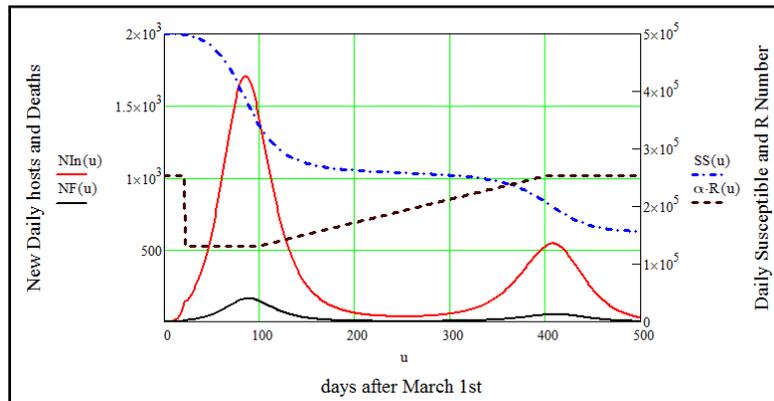


Figure 8: Result of the MP release over a longer period of time. Simulation by a progressive SD release, starting by the end of the third month, and ending ten months later, as shown by R(t).

Herd Immunity Analysis

Acquired immunity is conquered at the level of the individual, either through natural infection with a pathogen or through immunization with a vaccine. Herd immunity stems from the effects on individual immunity scaled to the level of the whole population. It is referred to the indirect protection from infection conferred to susceptible individuals when a sufficiently large proportion of immune individuals exist in a population. Depending on the prevalence of existing immunity to a pathogen in a population, an infected individual propagates the disease through susceptible hosts, following effective exposure to infected individuals as described by the SEIR model. However, if a percentage of the population has acquired some immunity level, the likelihood of an effective contact between infected and susceptible hosts is reduced, and the infection will not be transmitted by this path. The threshold proportion of susceptible persons required for transmission is known as the critical proportion P_c [18].

A relevant measure to evaluate the social cost of achieving global SARS-CoV2 herd immunity is the use of the Causality Rate (CR), defined as the proportion of deaths caused by a certain disease among all infected individuals. Now days in Brazil many Covid-19 cases are not reported, especially among asymptomatic hosts or individuals with mild symptoms, the CR will inherently be lower due to sub-notifications. It is important to remember that was established 50% asymptomatic hosts to the SEIR Model in this study. Massive serological testing will be required to better determine how many individuals have been infected, how many are immune, and how far we are from reaching the herd immunity threshold.

Within all those limitations, we can estimate a value for the herd immunity threshold P_c . Under the deterministic SEIR model, $P_c = 1 - 1/R_{eff}$, i.e., herd immunity threshold depends on a single parameter, the effective basic reproduction number R_{eff} [13]. Where $R_{eff} = (1 + r/\sigma_o).(1 + r/\gamma_o)$, r is the rate of the initial exponential growth, σ_o the exposed rate and γ_o the rate to be removed from the symptomatic infected group [13]. Since the onset of SARS-CoV-2 spread, studies have estimated the value of R_{eff} in the range of $1.1 < R_{eff} < 6.6$. From the previous values determined for r , σ_o , and γ_o , we obtained $R_{eff} = 3.0 \pm 0.3$, and $P_c = 0.67 \pm 0.03$. Again, in this study, R_{eff} is restricted to symptomatic hosts only, i.e., 50% of the exposed ones. As a result, the herd immunity threshold will be $\xi_{\sigma} \cdot P_c = 0.34 \pm 0.03$, and at least 35% of the population considered here remains to be immunized. As commented before, the number of COVID-19 notifications does not include the asymptomatic hosts or individuals with mild symptoms. As a result the number N of susceptible was fold by a factor of three, representing 12.3% of the Sao Paulo population. An extra fraction of 35% or, in the total, over 2 Million of exposed individuals to SARS-CoV-2 are required to cross herd immunity threshold at the city of S. Paulo.

Finally, given that the CR of COVID-19 estimated here is 0.23%, and preserving a factor of three fold in the number N of susceptible hosts, 44,000 is estimated as the number of people who could potentially die from COVID-19, whilst the population naturally reaches herd immunity. This number is difficult to be accepted, so, before a new vaccine becomes available reinforcements of mitigation policy become imperative.

ACKNOWLEDGEMENTS

The author acknowledges the financial support from Project Plat IoT, granted by FUNTTEL (“*Fundo para o Desenvolvimento Tecnológico das Telecomunicações*”). Financial grant # 01.16.0053.01 FINEP/MCTI from Brazilian Ministry of Science, Communications, and Innovation. This paper reflects only the author’s views and the Agencies are not responsible for any use that may be made of the information contained therein.

IV. CONCLUSION

A simple epidemiological model was presented to estimate the main results for the dynamics of the Covid-19 epidemic breakout in the city of São Paulo – Brazil. A time-dependent incidence weight on the reproductive basic number accounts for modeling social distancing, and local Mitigation Policies (MP). Published data from January 1st to July 7th, 2020 was used to adjust all the model parameters aiming to forecast the evolution of the COVID-19 epidemic outbreak. Data points for the last seven weeks, prior July 7th, were added after the model was complete, granting confidence on forecasts. By the last quarter of 2020, the predicted numbers of exposed individual in the city of S. Paulo may reach 1.9 million, symptomatic infected would sum 200 thousands and fatalities would accounts for 18 thousands. A value for the herd immunity threshold was estimated that at least 35% of the population considered here remains to be immunized. Therefore, over 2 Million of additional exposed individuals to SARS-CoV-2 are required to cross herd immunity threshold at the city of S. Paulo. This number is difficult to be accepted, so, before a new vaccine becomes available reinforcements of mitigation policy become imperative.

REFERENCES

- [1]. <https://ourworldindata.org/covid-testing>. Page visited on May 1st 2020.
- [2]. Zlojutro, A., Rey, D. & Gardner, L., “A decision-support framework to optimize border control for global outbreak mitigation”, *Nature Sci Rep* 9, 2216 (2019). doi.org/10.1038/s41598-019-38665-w.
- [3]. Wang, L., Wu, J.T., “Characterizing the dynamics underlying global spread of epidemics”. *Nature Commun* 9, 218 (2018). doi: 10.1038/s41467-017-02344-z
- [4]. Ruiyun Li, Sen Pei et. Al., “Substantial undocumented infection facilitates the rapid dissemination of novel coronavirus (SARS-CoV-2)”, *Science* 01: Vol. 368, Issue 6490, pp. 489-493, May 2020. doi: 10.1126/science.abb3221
- [5]. Kissler, S. M., Tedijanto, C., Goldstein, E., ET. Al., “Projecting the transmission dynamics of SARS-CoV-2 through the postpandemic period”, *Science* 14 Apr 2020. doi: 10.1126/science.abb5793
- [6]. B. Korber, B. et al., “Tracking changes in SARS-CoV-2 Spike: evidence that D614G increases infectivity of the COVID-19 virus”, *Journal Pre-proof*, Reference: CELL 11502, <https://doi.org/10.1016/j.cell.2020.06.043>
- [7]. “Painel de casos de doença pelo coronavírus 2019 (COVID-19) no Brasil pelo Ministério da Saúde”, <https://covid.saude.gov.br/>. Page visited on July 7, 2020.
- [8]. Coronavirus casos em SP <https://www.seade.gov.br/coronavirus/>. Visited on July 7.
- [9]. Grassly, N., Fraser, C. “Mathematical models of infectious disease transmission”. *Nature Rev Microbiol* 6, 477–487 (2008). doi: 10.1038/nrmicro1845.
- [10]. Anderson R.M., May R.M. Oxford Science Publications; Oxford, UK: 1992. *Infectious Diseases of Humans: Dynamics and Control*; p. 768. ISBN-10: 019854040X. ISBN-13: 978-0198540403.
- [11]. Gill, P. R.; Murray, W.; and Wright, M. H. “The Levenberg-Marquardt Method” §4.7.3 in *Practical Optimization*. London: Academic Press, pp. 136-137, 1981.
- [12]. Ferretti, L., Wyman, C., “Quantifying SARS-CoV-2 transmission suggests epidemic control with digital contact tracing”, *Science* 31 Mar 2020. doi: 10.1126/science.abb6936
- [13]. Wallinga, J., Lipsitch, M., “How generation intervals shape the relationship between growth rates and reproductive numbers”, *Proc Biol Sci.* 2007 Feb 22; 274(1609): 599–604. doi: 10.1098/rspb.2006.3754
- [14]. Oran, D. P., Topol, E. J., “Prevalence of Asymptomatic SARS-CoV-2 Infection”, *Annals of Internal Medicine*, June 2020, <https://doi.org/10.7326/M20-3012>
- [15]. Li, L. et. al., “COVID-19 patients' clinical characteristics, discharge rate, and fatality rate of meta-analysis”, *J. Med. Virol.*, 12 March 2020. <https://doi.org/10.1002/jmv.25757>.
- [16]. Hasell, J., et. al., “Research and data to make progress against the world’s largest problems” <https://ourworldindata.org/>. Visited on May, 1st.
- [17]. Hunter, P., “The spread of the COVID-19 coronavirus: Health agencies worldwide prepare for the seemingly inevitability of the COVID-19 coronavirus becoming endemic”, *EMBO Rep.* 2020 Apr 3;21(4): e50334. doi: 10.15252/embr.202050334
- [18]. Kwok, K., Lai, F., “Herd immunity – estimating the level required to halt the COVID-19 epidemics in affected countries”, *J Infect.* 2020 Jun; 80(6): e32–e33. doi: 10.1016/j.jinf.2020.03.027

Sergio Celaschi. "Quantifying Effects, Forecasting Releases, and Herd Immunity of the Covid-19 Epidemic in S. Paulo – Brazil." *The International Journal of Engineering and Science (IJES)*, 9(7), (2020): pp. 33-42.