

## Predictive Mathematical Modelling of the Total Number of COVID-19 Cases for Brazil

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### ABSTRACT

In this paper, we present different growth models such as Von Bertalanffy, Baranyi-Roberts, Morgan-Mercer-Flodin (MMF), modified Richards, modified Gompertz, modified Logistics and Huang in fitting and analyzing the epidemic trend of COVID-19 in the form of total number of infection cases of SARS-CoV-2 in Brazil as of 15<sup>th</sup> of July 2020. The MMF model was found to be the best model with the highest adjusted  $R^2$  value with the lowest RMSE value. The Accuracy and Bias Factors values were close to unity (1.0). The parameters obtained from the MMF model include maximum growth rate (log) of 0.026 (95% CI from 0.024 to 0.028), curve constant ( $\delta$ ) that affects the inflection point of 1.094 (95% CI from 1.024 to 1.165) and maximal total number of cases ( $y_{max}$ ) of 66,527,316 (95% CI from 35,156,044 to 143,548,943). The MMF predicted that the total number of cases for Brazil on the coming 15<sup>th</sup> of August and 15<sup>th</sup> of September 2020 will be 2,993,850 (95% CI of 3,407,196 to 2,630,649) and 4,676,829 (95% CI of 5,553,936 to 3,938,240), respectively. The predictive ability of the model utilized in this study is a powerful tool for epidemiologist to monitor and assess the severity of COVID-19 in Brazil in months to come. However, as with any other model, these values need to be taken with caution due to the unpredictability of the COVID-19 situation locally and globally.

### INTRODUCTION

A novel corona virus, 2019-nCoV, was identified as the cause of a respiratory disease outbreak that originated in Wuhan, China, and spread to several other countries around the world [1]. As the global death toll of COVID-19 continues to rise, there is an increasing awareness that SARS-COV-2 mortality is inequitably distributed among vulnerable populations. Such vulnerable groups include the elderly, densely populated, low socio-economic status, refugees and minorities. All groups are vulnerable. The rates of these classes are higher, placing them at a high risk of infection and for serious disease consequences [2,3]. At first, modelling studies centered on the Chinese

epidemic, and in particular on the epidemic dynamics in Wuhan City and in Hubei Province[4]. Much effort has been made at this early stage to evaluate surveillance data from China in order to obtain parameter estimates such as basic reproduction number ( $R_0$ ), case fatality rate and incubation period [5]. For the first attempts at Susceptible-Exposed-Infectious-Recovered (SEIR) style dynamic models, parameter estimates were 'borrowed' from what was known about other corona viruses (SARS-CoV and MERS-CoV) and/or obtained by fitting the models to the monitoring data [6].

The growth curve of viruses and microorganisms on substrates such as nutrients or other organisms, including

humans, typically followed a sigmoidal pattern, beginning with the lag section just after  $t = 0$ , preceded by the logarithmic segment and then entering the stationary period and eventually heading into the period of death or decreasing growth. There are various sigmoidal functions to define the growth curve of organisms, such as Von Bertalanffy, Baranyi-Roberts, modified Richards, modified Gompertz and modified Logistics [7] including Morgan-Mercer-Flodin (MMF) [8]. The growth curve valuable parameters include the maximum specific growth rate ( $\mu_m$ ), the lag period and the asymptotic values.

COVID-19 pandemic analyzes can be performed using statistical models, including theoretical, quantitative and simulation. For the analysis of the COVID-19 pandemic [9], strong predictive ability was employed models, such as updated Gompertz and Bertalanffy and logistics. The purpose of the paper is to test many available models like: "Logistic" (Ricker 1979[10]; [10], "Baranyi-Roberts" [11], "Von Bertalanffy"[12] ; [13] and "Margan" [14], "Baranyi Robert" [11]; [15] [11]" Morgan et al. and three-phase (Buchan 1993) recently Huang model [16] in fitting and analyzing the epidemic trend of COVID-19 in the form of total infection case of SARS-CoV-2 in Brazil as of 15<sup>th</sup> of July 2020.

**MATERIALS AND METHODS**

Data were collected from Worldometer (Worldometer 2020) for the total number of cases infected from Brazil as of 15 July 2020. Data were first converted to logarithmic values, and the time after first infected was utilized for time zero.

**Statistical analysis**

Statistical significant **difference** between the models was calculated through various methods including the adjusted coefficient of determination ( $R^2$ ), accuracy factor (AF), bias factor (BF), Root-Mean-Square Error (RMSE) and corrected AICc (Akaike Information Criterion) as before [17]. The RMSE was calculated according to Eq. (1), where  $Pd_i$  are the values predicted by the model and  $Ob_i$  are the experimental data,  $n$  is the number of experimental data, and  $p$  is the number of parameters of the assessed model.

$$RMSE = \sqrt{\frac{\sum_{i=1}^n (Pd_i - Ob_i)^2}{n - p}} \tag{eqn.1}$$

The adjusted  $R^2$  is used to calculate the quality of nonlinear models according to the formula where RMS is Residual Mean Square and  $S_y^2$  is the total variance of the y-variable ad calculated as follows;

$$Adjusted (R^2) = 1 - \frac{RMS}{S_y^2} \tag{eqn. 2}$$

$$Adjusted (R^2) = 1 - \frac{(1 - R^2)(n - 1)}{(n - p - 1)} \tag{eqn.3}$$

The Akaike information criterion (AIC) [18] was calculated as follows;

$$AICc = 2p + n \ln \left( \frac{RSS}{n} \right) + 2(p+1) + \frac{2(p+1)(p+2)}{n-p-2} \tag{eqn. 4}$$

Where  $n$  is the number of data points and  $p$  is the number of parameters of the model. The model with the smallest AICc value is highly likely correct [19].

Accuracy Factor (AF) and Bias Factor (BF) as suggested by Ross [20] were calculated as follows;

$$\text{Bias factor} = 10^{\left( \frac{\sum_{i=1}^n \log \left( \frac{Pd_i / Ob_i}{n} \right)}{n} \right)} \tag{eqn. 5}$$

$$\text{Accuracy factor} = 10^{\left( \frac{\sum_{i=1}^n \log \left( \frac{|Pd_i / Ob_i|}{n} \right)}{n} \right)} \tag{eqn. 6}$$

**FITTING OF THE DATA**

Fitting of the bacterial growth curve using various growth models (**Table 1**) was carried out using GraphPad Prism (v 8.0 trial version).

**Table 1.** Models used in this study.

Model	p	Equation
Modified Logistic	3	$y = \frac{A}{1 + \exp \left[ \frac{4\mu_m}{A} (\lambda - t) + 2 \right]}$
Modified Gompertz	3	$y = A \exp \left\{ -\exp \left[ \frac{\mu_m e}{A} (\lambda - t) + 1 \right] \right\}$
Modified Richards	4	$y = A \left\{ 1 + v \exp(1+v) \exp \left[ \frac{\mu_m}{A} (1+v) \left( 1 + \frac{1}{v} \right) (\lambda - t) \right] \right\}^{-1}$
Morgan-Mercer-Flodin (MMF)	4	$y = y_{max} - \frac{(y_{max} - \beta)}{1 + (\mu_m t)^\delta}$
Baranyi-Roberts	4	$y = A + \mu_m x + \frac{1}{\mu_m} \ln \left( e^{-\mu_m x} + e^{-h_0} - e^{-\left[ \frac{\mu_m x + \frac{1}{\mu_m} \ln \left( \frac{e^{-\mu_m x} + e^{-h_0} - e^{-\mu_m x - h_0}}{e^{(y_{max} - A)} - 1} \right)} \right]} \right)$
Von Bertalanffy	3	$y = K \left[ 1 - \left( \frac{A}{K} \right)^3 \exp \left( -\frac{\mu_m x}{3K} \right) \right]^{\frac{1}{3}}$
Huang	4	$y = A + y_{max} - \ln \left( e^A + (e^{y_{max} - A}) e^{-\mu_m B(x)} \right)$ $B(x) = x + \frac{1}{\alpha} \ln \frac{1 + e^{-\alpha(x-\lambda)}}{1 + e^{\alpha\lambda}}$
Buchanan Three-phase linear model	3	Y = A, IF X < LAG Y = A + K(X-λ), IF λ ≤ X ≤ X <sub>MAX</sub> Y = Y <sub>MAX</sub> , IF X ≥ X <sub>MAX</sub>

Note:  
 A= maximum no of cases lower asymptote;  
 y<sub>max</sub>= maximum no of cases upper asymptote;  
 μ<sub>m</sub>= maximum specific growth rate;  
 v= affects near which asymptote maximum no of cases occurs.  
 λ=lag time  
 e = exponent (2.718281828)  
 t = time after first case is reported  
 α,β,δ and k = curve fitting parameters  
 h<sub>0</sub> = a dimensionless parameter quantifying the initial physiological state of the reduction process. The lag time (h<sup>-1</sup>) or (d<sup>-1</sup>) can be calculated as h<sub>0</sub>=μ<sub>m</sub>  
 When data at time zero is 0 (Day after 1<sup>st</sup> case log 1=0 for COVID-19) the MMF is reduced to a 3-parameter model

**RESULTS AND DISCUSSION**

The coefficient of these models are displayed in (Fig 1 to 8), their values determine the shape of the curves. All the curves tested show visually acceptable fitting with the exception of the Buchanan-3-phase model (Figs 3). The best performance was the MMF model (fig 2) with the lowest value for RMSE, AICc and the highest value for adjusted  $R^2$ . The AF and BF values were also excellent for the model with their values which were the closest to 1.0. The worst performance among them was seen in fig 3 Buchanan-3- phase model (Table 2). This model proved to be unfit for data due to high residual variance and high nonlinearity values[21]. The coefficients for the MMF model are shown in Table 3, a good agreement between experimental data and predicted values was obtained.

The fitness order to actual data according to  $R^2$  values as well as MSE of all models explained the Total Number of COVID-19 Cases for Brazil in the following order; MMF> von Bertalanffy > modified Gompertz > modified Richards > Baranyi-Roberts > Huang > modified Logistics. Thus, MMF was determined to be the best fitted models and Buchanan-3-phase the worst[22].

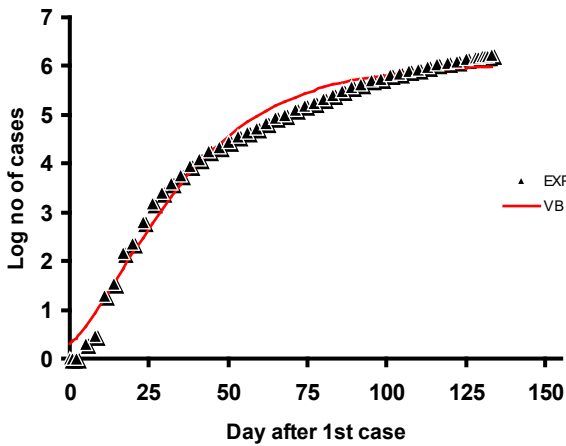


Fig. 1. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the von Bertalanffy model.

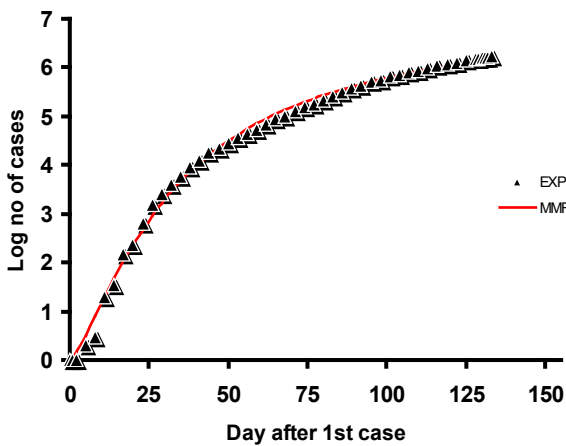


Fig. 2. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the MMF model.

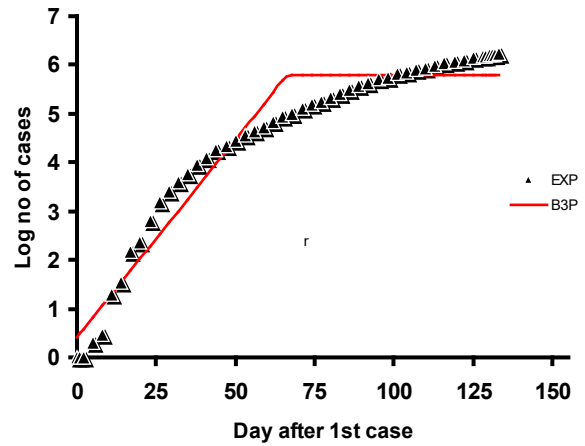


Fig. 3. Total no of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the Buchanan-3-phase model.

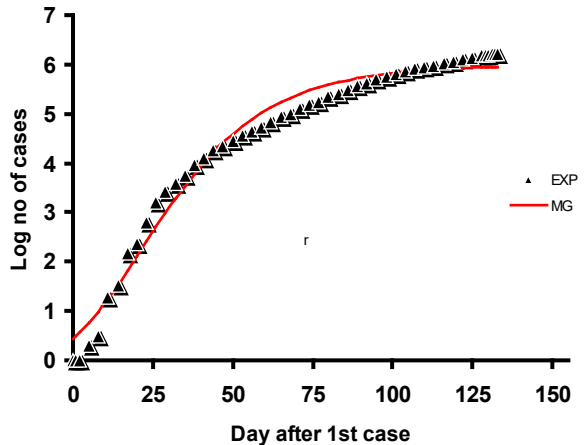


Fig.4. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the modified Gompertz model.

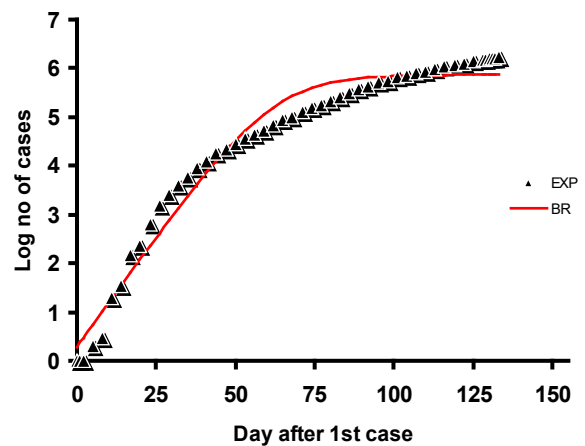


Fig. 5. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the Baranyi-Roberts model.

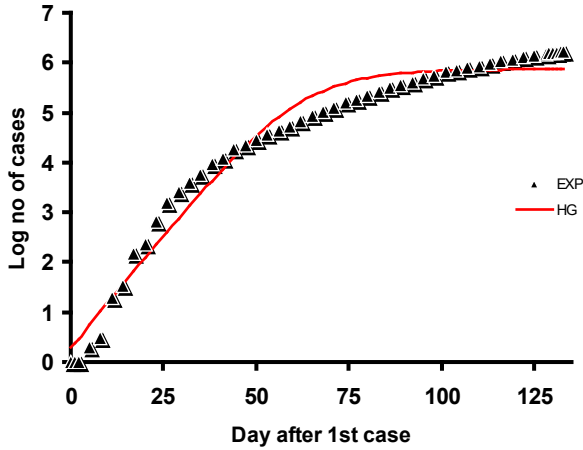


Fig. 6. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the Huang model.

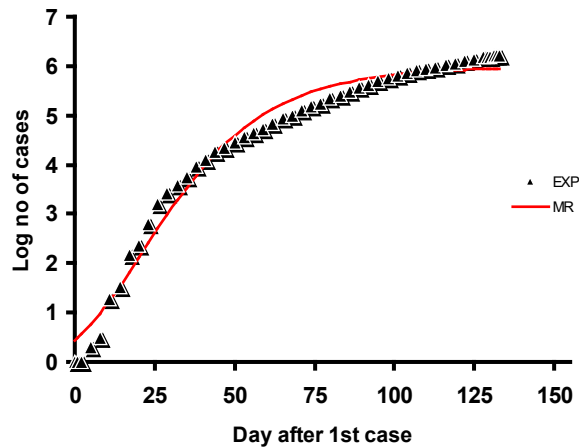


Fig. 7. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the modified Richard model.

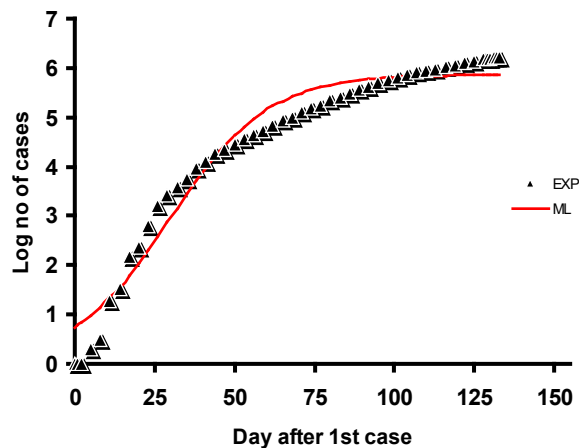


Fig. 8. Total number of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020 as modelled using the modified logistics model.

Table 2. Statistical tests for the various models utilized in modelling the total no of SARS-CoV-2 cases in Brazil as of 15<sup>th</sup> of July 2020.

Model	$p$	RMSE	$R^2$	$adR^2$	AF	BF	AICc
Huang	4	0.330	0.970	0.968	1.046	0.99	-95.73
Baranyi-Roberts	4	0.330	0.971	0.968	1.046	0.99	-95.74
modified Gompertz	3	0.279	0.978	0.976	1.062	1.00	-115.73
Buchanan-3-phase	3	0.433	0.948	0.945	1.062	0.99	-72.01
modified Richards	4	0.282	0.978	0.976	1.034	1.00	-111.25
MMF	3	0.116	0.996	0.996	1.013	1.00	-203.74
modified Logistics	3	0.382	0.956	0.953	1.045	0.99	-84.45
von Bertalanffy	3	0.232	0.985	0.984	1.029	1.00	-134.16

Note:  $p$  is no of parameter

Table 3. Coefficients as modelled using the MMF model.

Parameters	Value	95% Confidence interval
$\mu_m$	0.026	0.024 to 0.028
$\delta$	1.094	1.024 to 1.165
$y_{max}$	66,527,316	35,156,044 to 143,548,943

Table 4. Predictions of COVID-19 pandemic for Brazil based on the MMF model.

Prediction	Mean	95% Confidence interval
Maximum number of total cases by the end of COVID-19	66,527,316	35,156,044 to 143,548,943
Maximum number of total cases by 15 <sup>th</sup> of August 2020	2,993,850	3,407,196 to 2,630,649
Maximum number of total cases by 15 <sup>th</sup> of September 2020	4,676,829	5,553,936 to 3,938,240

The parameters obtained from the MMF model include maximum growth rate (log) of 0.026 (95% CI from 0.024 to 0.028), curve constant ( $\delta$ ) that affects the inflection point of 1.094 (95% CI from 1.024 to 1.165) and maximal total number of cases ( $y_{max}$ ) of 66,527,316 (95% CI from 35,156,044 to 143,548,943). The MMF model predicted that COVID-19 will end about 720 days (95% CI of 498 to 942) days from 15<sup>th</sup> of July 2020 based on the lower bound of the 95% CI from the calculated maximum number of total cases ( $y_{max}$ ) while the mean and upper 95% CI bound values failed to be predicted by the software for their number of days. The MMF predicted that the total number of cases for Brazil on the coming 15<sup>th</sup> of August and 15<sup>th</sup> of September 2020 will be 2,993,850 (95% CI of 3,407,196 to 2,630,649) and 4,676,829 (95% CI of 5,553,936 to 3,938,240), respectively. This prediction has to be taken with caution since the model failed to predict the number of days for the mean and upper 95% CI values and the number of days for COVID-19 to end may be much larger.

The MMF was initially developed as a model to describe nutrient-response relationships in a variety of organisms especially higher ones [8]. It has been utilized to model a variety of growth rates of animals including sheep, rabbit, horse and even microorganisms [23–27], palm oil’s yield [28], ethanol [29] and even in economics [30]. If the predicted data is right, this will depend on case by case and includes lockdown effectiveness. Viral mutations that increase the infectivity rate of the virus will change the course of the model outcome. Probably, the models should be revisited every few months to restructure the data and have a stronger forecasting ability.

## CONCLUSION

In conclusion, the MMF model was the best model and first in modelling the number of total infected case of Covid-19 based on statistical tests such as corrected AICc (Akaike Information Criterion), bias factor (BF), adjusted coefficient of determination ( $R^2$ ) and root-mean-square error (RMSE). Parameters obtained from the fitting exercise were maximum growth rate ( $\mu_m$ ), the curve constants ( $\delta$ ) and maximal total number of cases ( $Y_{max}$ ). The model predicts the total number of cases in Brazil and varies according to different numbers of factors. Notwithstanding this, an important method for epidemiologists in this study to track and determine the extent of COVID-19 in Brazil in the upcoming months is the predictive ability of a sample.

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