

APPLICATION OF DIFFERENTIAL TRANSFORMATION METHOD FOR SOLVING HIV MODEL WITH ANTI-VIRAL TREATMENT

Aplikasi Metode Tranformasi Diferensial Dalam Penyelesaian Model HIV Dengan Antiviral

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Abstract

Mathematical models have been widely used to understand complex phenomena. Generally, the model is in the form of system of differential equations. However, when the model becomes complex, analytical solutions are not easily used and hence a numerical approach has been used. A number of numerical schemes such as Euler, Runge-Kutta, and Finite Difference Scheme are generally used. There are also alternative numerical methods that can be used to solve system of differential equations such as the nonstandard finite difference scheme (NSFDS), the Adomian decomposition method (ADM), Variation iteration method (VIM), and the differential transformation method (DTM). In this paper, we apply the differential transformation method (DTM) to solve system of differential equations. The DTM is semi-analytical numerical technique to solve the system of differential equations and provides an iterative procedure to obtain the power series of the solution in terms of initial value parameters. In this paper, we present a mathematical model of HIV with antiviral treatment and construct a numerical scheme based on the differential transformation method (DTM) for solving the model. The results are compared to that of Runge-Kutta method. We find a good agreement of the DTM and the Runge-Kutta method for smaller time step but it fails in the large time step.

Keywords: DTM, Numerical methods, HIV, Model

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1. INTRODUCTION

Mathematical models have been widely used to understand complex phenomena and have been applied in many fields such as economics, epidemiology, biology, and social [1]–[4]. Many mathematical models have been formulated to understand complex phenomena such as disease transmission dynamics or rumor spread [4]–[8]. These models are in the form of system of differential equations and have been solved using the available software.

Biological problems are often complex and require a lot of resources to be investigated and understood clearly. Human Immunodeficiency Virus (HIV) is a major problem in the world [9] and thus its dynamics needs to be understood clearly. A mathematical model can be formulated to gain nonintuitive insights of the HIV transmission dynamics. Many mathematical models for HIV/AIDS transmission dynamics have been formulated and their corresponding numerical solutions have been presented [10]–[13]. They applied the nonstandard finite difference scheme and the differential transformation method for solving HIV mathematical models. Generally, they found that the solutions generated by these numerical schemes are in agreement with that use the traditional method such as Euler and Runge-Kutta.

In this paper, we implement the differential transformation method for solving HIV model with antiviral treatment. Mathematical models of HIV with anti viral treatment have also been developed [9], [14]–[16]. This paper extends the mathematical model by Priti [15] by including the logistic growth of T cells that are produced by proliferation of existing T cells.

A mathematical model is often complex since it includes many variables and parameters. Although several analytical analyses such as steady states can be derived, in general, the analytical solution of the models is not easily derived. Thus, a numerical approach is implemented. One of the numerical approaches to be used is the Differential Transformation Method (DTM). This method allows us to write a semi-analytical solution of the model which depends on the Taylor series [17]–[21]. The DTM has been successfully applied to solve various mathematical models including disease transmission models [18], [22]. Arenas et al. [17] used the differential transformation method for solving non autonomous system of differential equations. They found that the DTM is easy to implement, accurate, and satisfy the properties of its corresponding continuous model. Research by Jodar et al. also showed that the DTM is in good agreement with the Runge-Kutta method when it is applied to linear and non-linear stiff systems of differential equations. Allahviranloo et al. (2009) implement the DTM to solve fuzzy differential equations and found the identical solution between the DTM and differential inclusion. The DTM can be applied to solve the traveling wave solution [24].

Although the DTM has been applied to solve many mathematical models and shows to be accurate, several research shows that the DTM may fail in generating solution. Ndi et al. [20] compared the performance of the DTM and the nonstandard finite difference scheme and showed that the DTM may fail in generating the solution of system of differential equations. Ghafoori et al., (2011) compared the performance of the DTM and variation iteration method (VIM) and the homotopy perturbation method (HPM) and found that the DTM is more accurate than the other two methods. Furthermore, the DTM performance is similar to that of Adomian decomposition method (ADM). In this paper, we employ the DTM to solve the HIV model with antiviral treatment. We compare the results to that generated by Runge-Kutta method. The focus of this paper is the construction of the DTM method for solving the formulated mathematical model. We examine the convergence of the method by using different time step. The discussion of the results is presented.

The remainder of the paper is organised as follows. Section 2 presents methods consisting of formulation of mathematical model of HIV with antiviral treatment and the differential transformation method (DTM). Results and discussions are then presented. Finally, the conclusions are presented.

2. RESEARCH METHODS

2.1. Formulation of Mathematical Model

We implement the differential transformation method to solve the HIV model in the form of system of differential equations. A mathematical model for HIV with anti-viral treatment is proposed. The model is compartment-based model in the form of system of differential equations where the population is divided into different groups depending on their status.

We modified the model of Priti [15] by including the logistic growth of T cells as given in [16]. Let $x(t)$ be the uninfected CD4⁺ T cell population, $y_i(t)$ be the infected CD4⁺T cell population, $y_l(t)$ be latently infected CD4⁺T cell population and $z(t)$ be the CTL responses. We assume that the the new T cells are created by the proliferation of existing T cells, which is governed by a logistic function as given in [16]. Therefore, the growth of T cell is determined by natural growth λ and that created by the proliferation of existing T cells $ax\left(1-\frac{x}{x_{max}}\right)$. The cytotoxic T lymphocyte (CTL) responses at a rate s where the response depends on the population of infected T cell. This response will kill the infected T cell at a rate β_2 . Some CD4⁺T cells remain in the latent class but they are activated and move to the infected class at a rate δ . The RTI reduces the infection rate by $(1-\eta_1u_1)$ where η_1 is drug efficacy and u_1 is the control input doses of the drug RTI [15]. The detail of the explanation of the model is given in [15].

The model is governed by the following system of differential equations.

$$\begin{aligned} \frac{dx}{dt} &= \lambda - dx + ax\left(1 - \frac{x}{x_{max}}\right) - (1 - \eta_1u_1)\beta_1xy_i + \eta_2u_2x, \\ \frac{dy_i}{dt} &= (1 - \eta_1u_1)\beta_1xy_i - d_2y_i - \beta_2y_iz + \delta y_l, \\ \frac{dy_l}{dt} &= vy_i - d_3y_l - \delta y_l, \\ \frac{dz}{dt} &= sy_i - d_4z + \eta_3u_2z. \end{aligned} \tag{1}$$

Table 1. Parameter descriptions and values. The parameter values is given in [15], [16]

Parameters	Description	Values
a	Proliferation rate	0.5
x_{max}	Maximal population level of CD4 ⁺ T cells	1200
λ	Source term for uninfected CD4 ⁺ T cells	10
d	Death rate of uninfected CD4 ⁺ T cells	0.03
d_2	Death rate of infected CD4 ⁺ T cell population	0.3
d_3	Natural death rate of latently infected T cells	0.03
d_4	Natural death rate of CTL responses	0.2
η_1	Drug efficacy	0.9
η_2	Drug efficacy for IL-2 for uninfected CD4 ⁺ T cells	0.2
η_3	Drug efficacy for IL-2 for CTL responses	0.9
u_1	Control input doses of the drug RTI	0.5
β_1	Infection rate of uninfected cells	6×10^{-4}
β_2	Death of infected T cells due to CTL responses	0.001
δ	Progression from latent to infected	0.1
v	Rate of infected cells undergone latent stage	0.5
s	CTL response rate	0.2

2.2. The Differential Transformation Method (DTM)

The differential transformation method is a semi-analytic numerical technique which depends on the Taylor series. The method provides an iterative procedure to obtain the power series of the solution in terms of initial value parameters. In general, the procedure of the method is follows. First, the original function is transformed using the definition of the DTM, and then the solution of the model is obtained by iterative process using the Taylor series.

In this part, we develop a numerical scheme based on the concept of the Differential Transformation Method (DTM). The definition and the procedure of obtaining the solutions of the model are presented.

Definition 2.2.1. [16], [20], [25] Let $f(t)$ be analytic in a domain D and let $t = t_0$ represent any point in D . The Differential Transformation Method (DTM) for k^{th} derivative of function $f(t)$ is defined as

$$F(k) = \frac{1}{k!} \left[\frac{d^k f(t)}{dt^k} \right]_{t=t_0} \quad \text{for } t \in D, \quad (2)$$

where $f(t)$ is the original function and $F(k)$ is the transformed function.

Definition 2.2.2. [16], [20], [25] The inverse of the transformed function is defined as,

$$f(t) = \sum_{k=0}^{\infty} F(k)(t-t_0)^k, \quad \text{for } t \in D \quad (3)$$

From Equations (2) and (3), we obtain

$$f(t) = \sum_{k=0}^{\infty} \frac{(t-t_0)^k}{k!} \left[\frac{d^k f(t)}{dt^k} \right]_{t=t_0}, \quad t \in D. \quad (4)$$

Equation 4 implies that the concept of differential transformation method is derived from Taylor series.

We can use the Definition 2.2.1. and 2.2.2. to derive the transformed functions given in Table 2. In real application, we only use finite series for $f(t)$ as

$$f(t) = \sum_{k=0}^n F(k)(t-t_0)^k. \quad (5)$$

Using the above definition, we can derive the transformed function for functions. The results are given in Table 2, and details of the proofs of the transformed function in Table 2 are given in [20].

Table 2. Operation of differential transform	
Original function	Transformed function
$f(t)=u(t)+v(t)$	$F(k)=U(k)+V(k)$
$f(t)=au(t)$	$F(k)=aU(k)$
$f(t) = \frac{du(t)}{dt}$	$F(k)=(k+1)F(k+1)$
$f(t)=u(t)v(t)$	$F(K) = \sum_{l=0}^k V(l)U(k-l)$

The procedure for deriving the solutions of system of differential equations using the DTM is the following.

Let $[0, T]$ be the time interval of interest. We divide the time of interests into M subinterval $[t_{m-1}, t_m]$, $m = 1, 2, \dots, M$ of equal step size. We then find the solution of each time step using the concept of differential transformation method. The process of obtaining the solutions is the following. First, we obtain the solution of the first interval $[0, t_1]$ using

$$f_1(t) = \sum_{k=0}^N f_1(k)t^k, \quad t \in [0, t_1] \quad (6)$$

In the next time step, we use the value at $t=t_1$ as initial condition. In general, the value at the end time of the previous time step is used as an initial condition for the next time step. This is written as

$$f_m(t_m - 1) = f_{m-1}(t_m - 1) \quad (7)$$

and the solution of each time step is approximated using

$$f_m(t) = \sum_{k=0}^n F_m(k)(t-t_{m-1})^k, t \in [t_{m-1}, t_m]. \quad (8)$$

Let $m=M$ the multi-step DTM gives the following solution

$$f(t) = \begin{cases} \sum_{k=0}^n F_1(k)t^k, & t \in [0, t_1] \\ \sum_{k=0}^N F_2(k)(t-t_1)^k, & t \in [t_1, t_2] \\ \vdots \\ \sum_{k=0}^N F_m(k)(t-t_{m-1})^k, & t \in [t_{m-1}, t_m] \end{cases}$$

We then obtain the numerical solutions of the model.

We illustrate the use of the operation of differential transform.

$$\frac{dx}{dt} = 2xy.$$

The transformation of the above equation is

$$(k+1)X(k+1) = 2 \sum_{l=0}^k Y(l)X(k-l).$$

The results and discussion are presented in the next section.

3. RESULTS AND DISCUSSION

In this section, we present the results of the transformation of the Model 1 using the concept of the differential transformation method and the solutions of the model using the DTM. The operations of differential transform given in Table 2 are used. The transformed model is the following:

$$\begin{aligned} X(K+1) &= \frac{1}{k+1} (\lambda - dX(K) + aX(K) \\ &\quad - \frac{a}{x_{max}} \sum_{l=0}^K X(l)X(K-l) - \\ &\quad (1-\eta_1 u_1) \beta_1 \sum_{l=0}^K X(l)Y_i(K-l) \\ &\quad + \eta_2 u_2 X(K)), \\ Y_i(K+1) &= \frac{1}{K+1} ((1-\eta_1 u_1) \beta_1 \sum_{l=0}^K X(l)y_i(K-l) \\ &\quad - d_2 Y_i(K) - \beta_2 \sum_{l=0}^K Y_i(l)Z(K-l) \\ &\quad + \delta Y_i(K)), \\ Y_l(K+1) &= \frac{1}{K+1} (vY_l(K) - (d_3 + \delta)Y_l(K), \\ Z(K+1) &= \frac{1}{K+1} (sY_i(K) - d_4 Z(K) + \eta_3 u_2 Z(K)). \end{aligned} \quad (9)$$

The invers of this function is the solution of the model above. Let divide t into $t_1, t_2, t_3, \dots, t_n$

$$x(t) = \begin{cases} \sum_{K=0}^N X(K)(t-t_0)^K, & t \in [t_0 \quad t_1] \\ \sum_{K=0}^N X(K)(t-t_1)^K, & t \in [t_1 \quad t_2] \\ \sum_{K=0}^N X(K)(t-t_2)^K, & t \in [t_2 \quad t_3] \\ \vdots \\ \sum_{K=0}^N X(K)(t-t_{m-1})^K, & t \in [t_{m-1} \quad t_m] \end{cases}$$

$$y_i(t) = \begin{cases} \sum_{K=0}^N Y_i(K)(t-t_0)^K, & t \in [t_0 \quad t_1] \\ \sum_{K=0}^N Y_i(K)(t-t_1)^K, & t \in [t_1 \quad t_2] \\ \sum_{K=0}^N Y_i(K)(t-t_2)^K, & t \in [t_2 \quad t_3] \\ \vdots \\ \sum_{K=0}^N Y_i(K)(t-t_{m-1})^K, & t \in [t_{m-1} \quad t_m] \end{cases}$$

$$y_l(t) = \begin{cases} \sum_{K=0}^N Y_l(K)(t-t_0)^K, & t \in [t_0 \quad t_1] \\ \sum_{K=0}^N Y_l(K)(t-t_1)^K, & t \in [t_1 \quad t_2] \\ \sum_{K=0}^N Y_l(K)(t-t_2)^K, & t \in [t_2 \quad t_3] \\ \vdots \\ \sum_{K=0}^N Y_l(K)(t-t_{m-1})^K, & t \in [t_{m-1} \quad t_m] \end{cases}$$

$$z(t) = \begin{cases} \sum_{K=0}^N Z(K)(t-t_0)^K, & t \in [t_0 \quad t_1] \\ \sum_{K=0}^N Z(K)(t-t_1)^K, & t \in [t_1 \quad t_2] \\ \sum_{K=0}^N Z(K)(t-t_2)^K, & t \in [t_2 \quad t_3] \\ \vdots \\ \sum_{K=0}^N Z(K)(t-t_{m-1})^K, & t \in [t_{m-1} \quad t_m] \end{cases}$$

In this section, we present a numerical experiments of the DTM for the HIV model. The parameter values are given in Table 1. Here we compare the results using DTM and the Runge Kutta. The results are given

in Figure 1. Numerical comparison between Runge-Kutta and Differential Transformation Method for $h=0.001$ and Table 3.

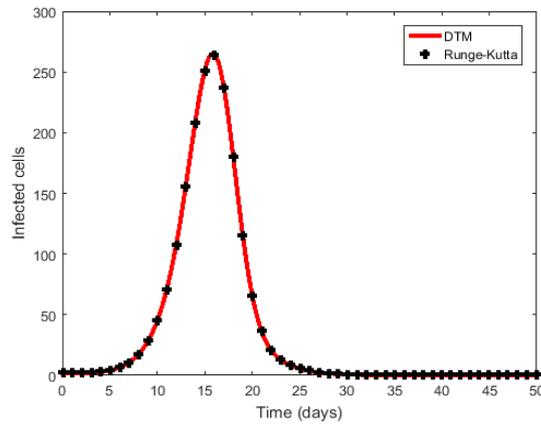


Figure 1. Numerical comparison between Runge-Kutta and Differential Transformation Method for $h=0.001$

Figure 1 shows that the DTM is in agreement with the Runge-Kutta method. The result differences between the DTM and Runge-Kutta are given in Table 3. The model is for the time step of 0.01. However, when we increase the time step h the solution cannot be generated as given in Figure 2.

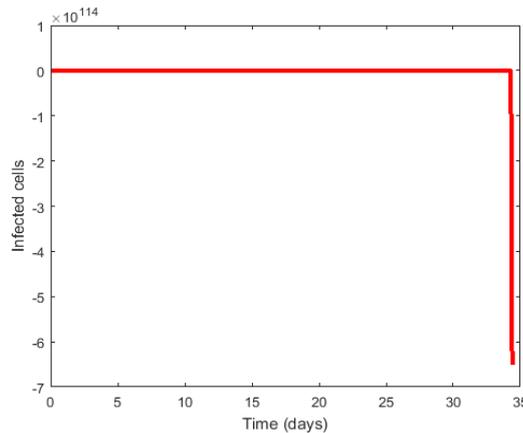


Figure 2. The solution using the Differential Transformation Method for $h=0.1$.

Table 3. The error of the numerical results of the model using the differential transformation method and Runge-Kutta method. The time step is 0.01 and error tolerance for Runge-Kutta is 10^{-6}

t (days)	Differences
5	0.000115545
10	0.001754874
15	0.005111424
20	0.002726564
25	0.000149172
30	1.82729×10^{-5}
35	2.73116×10^{-6}
40	6.68361×10^{-7}
45	6.79724×10^{-8}
50	1.3401×10^{-9}

This paper aims to construct the DTM to solve anti-viral treatment. This shows that the DTM can generate the solutions of the model. However, it shows that the DTM cannot generate the solution if the time step h is not appropriate. We found that although the time step is 0.1, the DTM cannot generate the solution. The same results are found by Ndi et al. [25], and Odibat (2008). The DTM has been applied to many mathematical models and found a good agreement between the DTM and the traditional methods

such as Runge-Kutta and Euler methods. However, in this paper, our results suggest that the DTM cannot generate the solution of the model if the time step is high.

The results imply that the DTM may not appropriate for the use in solving system of differential equations when the traditional methods fail in generating solution. Given that the traditional numerical schemes such as Runge-Kutta, Euler fail in generating the solution of differential equations, an alternative numerical schemes are required. A number of numerical schemes has been proposed such as the nonstandard finite difference scheme (NSFDS), Adomian Decomposition method (ADM), Variation iteration method (VIM), homotopy perturbation method (HPM) [26]–[31]. A little research has been done to compare the performance between the NSFDS and the DTM. Ndi et al. [25] compared the performance between these methods and showed that the similar performance of the NSFDS and DTM in small time step. When the time step is larger, the DTM fails in generating the solution. The performance of the DTM is similar to Adomian decomposition method (ADM) [27] and is more accurate than the Variational iteration method (VIM) and Homotopy perturbation method (HPM) [26]. Furthermore, the DTM can be modified to enhance its performance to generate the solutions of the system of differential equations.

The results suggest that the use of the numerical schemes depends on the problems to be solved. The DTM performs better than the VIM and HPM but is not better than the ADM and the NSFDS. Therefore, the use of the DTM can be applied for the smaller time step.

4. CONCLUSIONS

Research showed that the DTM is only effective for a small time step. When the time step is higher, the DTM fails in generating the solutions of the model. Some modifications may be required to enhance the performance of the method but it is not the focus of the paper.

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