ANALYSIS OF THE SIRS MODEL OF THE SPREAD OF DENGUE HEMORRHAGIC FEVER USING RUNGE-KUTTA METHOD AND GENETIC ALGORITHM

Melyssa Mentari Tjioenata¹⁾, Samuel Lukas^{2*)}, Dina Stefani³⁾, Petrus Widjaja⁴⁾

¹Mathematics Department, Faculty of Science and Technology, Universitas Pelita Harapan, Indonesia ²Mathematics Department, Faculty of Science and Technology, Universitas Pelita Harapan, Indonesia ³Informatics Department, Faculty of Computer Science, Universitas Pelita Harapan, Indonesia ⁴Informatics Department, Faculty of Computer Science, Universitas Pelita Harapan, Indonesia Email : ¹melyssa010698@gmail.com, ²samuel.lukas@uph.edu, ³dinastefanilukas@gmail.com, ⁴petrus.widjaja@uph.edu

*Penulis Korespondensi

Abstract –Dengue Hemorrhagic Fever is a contagious disease that often occurs in Indonesia. Dengue Hemorrhagic Fever, caused by the Dengue Virus, has four serotypes : DEN-1, DEN-2, DEN-3 and DEN-4. Someone can be infected four times, once for each serotype. After recovering from one serotype, a person gets a lifetime of immunity against that serotype. In this paper a model will be created for modeling the spread of Dengue Hemorrhagic Fever with assumption there are only two serotypes, namely DEN-1 and DEN-2. Model formed based on SIR Model (Susceptible, Infected, Recovered) and SIRS Model (Susceptible, Infected, Recovered, Susceptible). The changes in populations over time are written into a system of differential equations. The system of differential equations is then used to do an equilibrium point analysis. The numerical solution for the system of differential equations can be found using the Runge-Kutta Method. Genetic Algorithm are used to find the values of parameters in the model that are unknown. A series of simulations are performed to get the combination that produces the Genetic Algorithm system that will produce the best approximation solution. The combination includes population size, selection value, crossover value and mutation value. This best combination then will produce an approximation solution with the smallest error.

Keywords: Epidemiological model, Equilibrium point, Runge-Kutta, Genetic Algorithm

Abstrak – Demam Berdarah Dengue merupakan penyakit menular yang sering terjadi di Indonesia. Demam Berdarah Dengue, yang disebabkan oleh Virus Dengue, memiliki empat serotipe: DEN-1, DEN-2, DEN-3 dan DEN-4. Seseorang dapat terinfeksi empat kali, sekali untuk setiap serotipe. Setelah sembuh dari satu serotipe, seseorang mendapat kekebalan seumur hidup terhadap serotipe tersebut. Pada makalah ini akan dibuat suatu model untuk pemodelan penyebaran Demam Berdarah Dengue dengan asumsi hanya ada dua serotipe yaitu DEN-1 dan DEN-2. Model dibentuk berdasarkan Model SIR (Susceptible, Infected, Recovered) dan Model SIRS (Susceptible, Infected, Recovered, Susceptible). Perubahan populasi dari waktu ke waktu ditulis ke dalam sistem persamaan diferensial. Sistem persamaan diferensial ini kemudian digunakan untuk melakukan analisis titik setimbang. Solusi numerik untuk sistem persamaan diferensial dapat ditemukan dengan menggunakan Metode Runge-Kutta. Algoritma Genetika digunakan untuk mencari nilai parameter dalam model yang tidak diketahui. Serangkaian simulasi dilakukan untuk mendapatkan kombinasi yang menghasilkan sistem Algoritma Genetika yang akan menghasilkan solusi aproksimasi terbaik. Kombinasi tersebut meliputi ukuran populasi, nilai seleksi, nilai crossover dan nilai mutasi. Kombinasi terbaik ini kemudian akan menghasilkan solusi aproksimasi dengan error terkecil.

Kata Kunci: Model epidemiologi, Titik kesetimbangan, Runge-Kutta, Algoritma Genetika

INTRODUCTION

Dengue Hemorrhagic Fever is still one of the major health problems in Indonesia [1]. Dengue fever is mosquito-borne tropical disease with severe flu-like symptoms, caused by the Dengue Virus spreads through bites of infected mosquitoes. That mosquito infected can transmit dengue virus for the rest of its life [2]. There are four serotypes of Dengue Fever, namely: Dengue Virus 1 (DEN-1 or DENV-1), Dengue Virus 2 (DEN-2 or DENV-2), Dengue Virus 3 (DEN-3 or DENV-3) and Dengue Virus 4 (DEN-4 or DENV-4). After recovering from certain serotype Dengue Fever, individuals get lifelong immunity life against the serotype dan temporary immunity to three other serotypes [2]. The spread of Dengue Fever can be made in the form of mathematical models. There are various mathematical models that can be used for epidemics, the simplest was discovered by Kernack and McKendrick in 1927, SIR model (Susceptible, Infected, Recovered). Finding certain parameter values from a model can be done in many ways. In this paper, Genetic Algorithms will be used. Genetic Algorithm (Genetic Algorithms) was first introduced by John Holland in years 1960s and later developed by Holland and colleagues as well his students at the University of Michigan in the 1960s and 1970s. Algorithm Genetics is a method that uses a set of "chromosomes" to create new populations using "natural selection". Each chromosome consists of "genes". Selected chromosomes can produce offspring, where chromosomes with a higher fitness value tend to produce better offspring [3].

EPIDEMIOLOGICAL MODELS

The model used is the epidemiological model, namely the SIR model and SIRS model. This research assumes that there are only two serotypes, which are means when an individual suffers from Dengue Fever for the first time, a model the SIRS model is used. When individuals suffer from Dengue Fever for the second time, the SIR model was used, which means the individual will never has Dengue Fever again all his/her life because she/he has gained immunity lifetime for both serotypes. The population is divided into four groups [4]:

- 1. Susceptible Individuals are not infected with pathogens, but have low immunity against the disease.
- 2. Exposed Individuals who interact with other individuals who have contracted with the pathogen.
- 3. Infected Individuals from the susceptible group who have high levels of parasites in the body as well as being able to transmit diseases to other individuals within susceptible group.
- 4. Recovered Individuals whose immune system has eliminated all parasites in the body and are no longer able to transmit disease to other individuals within susceptible group.

Based on the group, the SEIR model was formed. In order to make the simpler model, SIR model, where the exposed groups are ignored. Besides SIR, there are also SI model that are commonly found on plants, SIS model that are commonly found in sexually transmitted diseases, and SIRS model where immunity is temporary.

The equation for the SIR model in general is as (1) [4]:

$$\begin{cases} \frac{ds}{dt} = \mu - \beta IS - \mu S\\ \frac{dI}{dt} = \beta IS - \gamma I - \mu I\\ \frac{dR}{dt} = \gamma I - \mu R \end{cases}$$
(1)

The parameter β is the transmission rate, γ is the removal or recovery rate and μ is the rate individuals suffer natural mortality and also the birth rate.

The equation for the SIRS model in general is as (2) [4]:

$$\begin{cases} \frac{ds}{dt} = \mu - \beta IS - \mu S + \omega R\\ \frac{dI}{dt} = \beta IS - \gamma I - \mu I\\ \frac{dR}{dt} = \gamma I - \mu R - \omega R \end{cases}$$
(2)

The parameter ω is the rate at which immunity is lost. The equation for the SIR and SIRS model for dengue fever is as (3). Mosquito vector and human vector model are shown in Fig.1 and Fig.2. Red line represents output of each process. The Description and value of each variabel is tabulated in Table 1.

$$\begin{cases} \frac{ds}{dt} = v - \Omega \left(\frac{\beta_h b}{N_h} I_v S \right) - (1 - \Omega) \left(\frac{\beta_h b}{N_h} I_v S \right) - \mu_s S \\ \frac{dI_1}{dt} = \Omega \left(\frac{\beta_h b}{N_h} I_v S \right) - \gamma_1 I_1 - \mu_{I_1} I_1 \\ \frac{dR_1}{dt} = \gamma_1 I_1 - \omega_1 R_1 - \mu_{R_1} R_1 \\ \frac{dR_2}{dt} = \gamma_2 I_2 - \omega_2 R_2 - \mu_{R_2} R_2 \\ \frac{dS_1}{dt} = \omega_1 R_1 - \left(\frac{\beta_h b}{N_h} I_v S_1 \right) - \mu_{S_1} S_1 \qquad \dots (3) \\ \frac{dS_2}{dt} = \omega_2 R_2 - \left(\frac{\beta_h b}{N_h} I_v S_2 \right) - \mu_{S_2} S_2 \\ \frac{dI_{12}}{dt} = \left(\frac{\beta_h b}{N_h} I_v S_1 \right) + \left(\frac{\beta_h b}{N_h} I_v S_2 \right) - \gamma_{12} I_{12} - \mu_{I_{12}} I_{12} \\ \frac{dR_{12}}{dt} = \gamma_{12} I_{12} - \mu_{R_{12}} R_{12} \\ \frac{dS_v}{dt} = \mu - \frac{\beta_v b}{N_h} (I_1 + I_2 + I_3) S_v - \mu_{S_v} S_v \\ \frac{dI_v}{dt} = \frac{\beta_v b}{N_h} (I_1 + I_2 + I_3) S_v - \mu_{I_v} I_v \end{cases}$$

Fig.1 Mosquito Vector Model



EQUILIBRIUM POINT

Equilibrium Point will occur when all values in left side of equation [3] are zero [5]. The equation is solved using Maple software progam. It is tabulated in Table 2.

X _i	X_1	X_2	X_3	X_4
S	110.80	23.16	-454.96	-290.28
I_1	0	0.24	1.15	1.10
I_2	0	0,15	1	0.71
R_1	0	2.71	17.57	12.45
R_2	0	1.73	11.17	7.92
S_1	0	0.96	352.18	-463.54
S_2	0	0.60	128.54	778.98
I_{12}	0	0.03	-2.68	-1.96
<i>R</i> ₁₂	0	3.80	-385.19	-280.84
S_v	55.00	53.86	55.37	55.41
I_v	0	1.14	-0.37	-0.41

There are four equilibrium points from table 2. The first equilibrium point, X_1 is the equilibrium point when there is no Dengue Fever in the entire population. X_1 is the equilibrium point without disease. The second equilibrium point, X_2 is the equilibrium point when both serotypes Dengue fever exists in the population. The third equilibrium point, X_3 is the "unnatural" equilibrium point, because population values cannot be negative. The fourth equilibrium point, X_4 is the "unnatural" equilibrium point, because population values cannot be negative.

Table 2. Equilibrium Point

Table 1. Description and value of each variable

Var	Description	Unit	Value
υ	Per capita birth rate	Per day	0.01662
Ω	Percentage of individuals infected Serotype 1 Dengue Fever (DENV-1)	%	0.61
$\frac{\beta_h b}{N_h}$	Transmission rate of Dengue Fever for humans	Per day	0.0005
γ ₁	Recovery rate of Serotype 1 Dengue Fever (DENV-1)	Per day	0.033
γ ₂	Recovery rate of Serotype 2 Dengue Fever (DENV-2)	Per day	0.033
γ ₁₂	Recovery individual suffers rate from Dengue Fever for the second time	Per day	0.033
ω_1	Immunity lost rate against Serotype 1 Dengue Fever (DENV-1)	Per day	0.0027
ω_2	Immunity lost rate against Serotype 2 Dengue Fever (DENV-2)	Per day	0.0027
μ	Per capita birth rate for mosquitos vector	Per day	0.055
$\frac{\beta_v b}{N_h}$	Transmission rate of Dengue Fever for vectors	Per day	0.00005
$\frac{N_h}{S}$	Population who have never suffered from Dengue Fever and susceptible to both serotypes	People	-
I_1	Population who suffered from Serotype 1 Dengue Fever (DENV-1)	People	-
I_2	Population who suffered from Serotype 2 Dengue Fever (DENV-2)	People	-
R_1	Population who have cured from Serotype 1 Dengue Fever (DENV-1)	People	-
R_2	Population who have cured from Serotype 2 Dengue Fever (DENV-2)	People	-
<i>S</i> ₁	Population who have never suffered from Serotype 2 Dengue Fever and susceptible to the serotype	People	-
S_2	Population who have never suffered from Serotype 1 Dengue Fever and susceptible to the serotype	People	-
<i>I</i> ₁₂	Population who suffered from Dengue Fever for the second time	People	-
R_{12}^{12}	Population who have cured from Dengue Fever for the second time	People	-
S_v	Population of mosquitoes who have never suffered from Dengue Fever and susceptible to the disease	Head	-
I_v	Population of mosquitos who suffered from Dengue Feve	Head	-
μ_{S}	Per capita death rate for S group	Per day	0.00015
μ_{I_1}	Per capita death rate for I_1 group	Per day	0.00030
μ_{I_2}	Per capita death rate for I_2 group	Per day	0.00025
μ_{R_1}	Per capita death rate for R_1 group	Per day	0.00022
μ_{R_2}	Per capita death rate for R_2 group	Per day	0.00024
μ_{S_1}	Per capita death rate for S_1 group	Per day	0.00020
μ_{S_2}	Per capita death rate for S_2 group	Per day	0.00021

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$\mu_{I_{12}}$	Per capita death rate for I_{12} group	Per day	0.00040
$\mu_{R_{12}}$	Per capita death rate for R_{12} group	Per day	0.00023
μ_{S_n}	Per capita death rate for S_v group	Per day	0.00050
μ_{I_v}	Per capita death rate for I_v group	Per day	0.00050

GENETIC ALGORITHM

Genetic Algorithm is designed to find out value of eleven parameters out of 21 parameters of equation (3) assuming that other parameters are constants. They are the per capita mortality rate in each group. They are μ_S , μ_{I_1} , μ_{I_2} , μ_{R_1} , μ_{R_2} , μ_{S_1} , μ_{S_2} , $\mu_{I_{12}}$, $\mu_{R_{12}}$, μ_{S_0} and μ_{I_0} . Each death rate cannot exceed 1, then the death rate must be between 0 and 1. The solutions for the equation of the SIR and SIRS model for dengue fever will be calculated using Runge-Kutta fourth-order method (RK4) with initial conditions t = 0 are

 $S=950, I_1 = I_2 = R_1 = R_2 = S_1 = 0,$ $S_2 = I_{12} = R_{12} = I_v = 0, S_v = 1000,$ time t = 150 and stepsize h = 0.05.

The form of ith chromosome is as follows:

 $C(i) = (S, I_1, I_2, R_1, R_2, S_1, S_2, I_{12}, R_{12}, S_\vartheta, I_\upsilon)$

The Genetics algorithm is as follow Initial population

While condition is true

Evaluation Selection Crossover Mutation

End

Interprete the solution

Initial population is the first process to have initial population in Genetic Algorithm. That is generate n number of chromosome in random. Since the values of gene are small number and to speed up the processing time of Genetic Algorithm then initial value of each gene in a chromosome is started with multiplying random numbers between 0 and 1 by 0.001. All chromosome is stored in matrix C.

Evaluation is done by using fitness function. Fitness function of i^{th} chromosome is defined with equation (4)

$$f(i) = 1 - \left(\frac{1}{11}\right) \left(\frac{\sum_{i=1}^{11} (C(i,j) - c'(i,j))^2}{\sqrt{\sum_{i=1}^{11} A'(i,j)^2}}\right)$$
(4)

C'(i, j) is true solution using Runge-Kutta with values from Table 1.

Selection is done with elitism algorithm by maintaining a number of the best chromosome of every generation [3]. That is sval% \times n number of chromosomes in population. Selection is carried out using the following algorithm:

- a. Sort the *C* matrix based on its fitness value with the Quicksort algorithm.
- b. Create new ten chromosomes
- c. The remainder chromosomes in new population are chosen randomly from previous population.

Crossover is done by two-point crossover, exchanging segments in between two randomly selected crossover point positions [3]. Crossover done is carried out using the following algorithm

- a. Determine number of chromosome are going to be crossovered. That is sval% \times n number of chromosomes in population.
- b. Determine pairs of chromosome are going to be crosovered randomly.
- c. Determine two random crossover point positions for each pair parent.
- d. Swap segments between two crossover point positions for each parent pair

Mutation is done by changing genes in chromosomes randomly. Mutation is carried out using the following algorithm

- a. Determine the number of gene that will undergo mutation. That is mval \times 11 \times n number of chromosomes in population.
- b. Determine the mutated gene randomly.
- c. Value of each mutated gene is filled by multiplying 0.001 to random number 0 to 1.

Repetition will stop when at least one of the chromosomes has the fitness value greater than 0.99999.

RESULT AND DISCUSSION

Experiment is started by determining population size n, selection value sval, crossover value cval and mutation value mval. The goal of experiment is not only to find the best chromosome but also the best combination of genetic parameter that is n, sval, cval and mval. The experimen is to determine what is the best population size with sval = 0.4; cval = 0.7 and mval = 0.1. The experimen result is shown in Table 3. The best population size is n = 500 because it is not only having the lowest error but also the smallest average generation.

Table 3. Experiment result

Avorago	Number of pupulation (n)				
Average	100	200	300	400	500
Average Error	0.64	0.41	0.41	0.50	0.22
Lowest Error	0.27	0.27	0.26	0.18	0.15
Average Time (sec)	857	912	3394	4711	3127
Average Generation	304	312	394	552	246
Average tipe perstep (Sec)	3.23	3.01	7.94	7.61	14.18

The best chromosome is C=(0.00022, 0.00022, 0.00021, 0.00021, 0.00021, 0.00021, 0.00021, 0.00021, 0.00018, 0,00050, 0.00050. 0.00050) with h = 0.05, error = 0.26, t = 5049 sec.

From the experiment can be seen that the average error and the best error is inversely proportional to n. This means the results are getting better if the larger population n increase. Incontrary, average time is

Diterima <23082023> | Revisi <10012023> | Diterima untuk publikasi <28012023> Lisensi: Creative Commons Attribution 4.0 International (CC BY 4.0) increase while *n* increase except for n = 500.

Figure 3 is the actual solution using Runge-kutta fourth order method and Fig. 4 using Genetic Algorithm.





Fig. 4. GA solution

CONCLUSION

In this paper a mathematical model has been made for the spread of Dengue Fever with two serotypes based on the SIR Model and SIRS Model. The solutions are listed in Table 3 with an error of 0.26.

DAFTAR PUSTAKA

- "Infodatinsituasi-demam berdarah-dengue," Kementerian Kesehatan Republik, 2018, Indonesia.Available at http://www.depkes.go.id/resources/download/pusda tin/info datin/InfoDatin-Situasi-Demam-Berdarah-Dengue. [Diakses pada 22 September 2019].
- [2] "Fact sheet dengue and severe dengue," World Health Organization, 2019. Available at https://www.who.int/newsroom/factsheets/detail/dengue-and-severe-dengue. [Diakses pada 22 September 2019].
- [3] Melanie Mitchell. An introduction to genetic algorithms. MIT press, 1998.
- [4] Matt J Keeling and Pejman Rohani. Modeling infectious diseases in humans and animals. Princeton University Press, 2011.
- [5] Erwin Kreyszig, Herbert Kreyszig, and E. J. Norminton. Advanced Engineering Mathematics. Wiley, Hoboken, NJ, tenth edition, 2011