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Implementation of Fuzzy-based Model for Prediction of Thalassemia Diseases

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Abstract: Thalassemia is known as one of the blood disorder diseases that is inherited by parents. There are several types of Thalassemia, namely as Thalassemia major, minor, and intermedia. Among them, Thalassemia major is the most dangerous and needs more attention. Generally, it can be detected since the child is one year old. Late detection of this disease can have adverse consequences and various complications. This study aims to develop a new model for the prediction of thalassemia for children. The model adopts a fuzzy-based rule. The novelty in this article is that our model has 4 outputs, namely thalassemia major, intermedia, minor and not thalassemia. In the previous article it only had 3 outputs. In this study, we intend to implement a model that we developed using a fuzzy-based approach to classify thalassemia diseases based on CBC data. This article describes how to build a model and implement it in software. We compare the test results with the opinion of pediatricians regarding thalassemia. The final results of testing 4 CBC data show that our proposed model has successfully identified the type of thalassemia.

Keywords: Blood disorder diseases, Thalassemia, Fuzzy-based model, Thalassemia classification, Artificial Intelligence

1. Introduction

In many countries, there have been many ¹ports of blood disorders. One of them is thalassemia. The increase in thalassemia cases is due to the lack of genetic counseling and prenatal diagnosis [1][2][3]. Actually, thalassemia can be known after the baby is born through several visible symptoms. Currently, most of thalassemia parents only know after the child is two years old. In fact, thalassemia is very dangerous if it is not treated immediately [4]. Based on clinical manifestations, thalassemia is classified into several types, namely thalassemia major, thalassemia minor or trait or carier and thalassemia intermedia [5][6][7]. Patients with thalassemia major require routine blood transfusions throughout their life. Patients with thalassemia intermedia also require blood transfusions, but not as often as thalassemia major. Meanwhile, thalassemia minor patients do not need blood transfusions but

only carry thalassemia traits[8]. Thalassemia treatment in Indonesia is currently still supportive, it has not yet reached the level of cure. Supportive treatment given to thalassemia patients aims to treat the symptoms that arise due to thalassemia. Symptoms include pale, yellowish or gray skin. In addition, there are also disturbances in growth. Lifelong routine transfusions, iron chelation administration, and psychosocial support are the main treatments for thalassemia major patients. [9][10].

A child becomes thalassemia due to genetic factors, namely a blood disorder that is inherited from parents[11][12][7]. Based on Mendel's law, it is known that if both parents or one of the two are characteristic carriers, the trait will be passed on to the child. Actually, thalassemia can be known based on visible physical symptoms. An easily seen disorder in thalassemia sufferers is that the child looks pale so that he experiences symptoms such as anemia[13][14]. In addition, thalassemia can also cause fatigue, dizziness, and shortness of breath because the blood vessels that supply the lungs are narrowed, forcing the heart to work harder to push blood in [15][16][17]. As a result, the activity of thalassemia sufferers will be disrupted. Thalassemia needs to be watched out for, especially thalassemia major because it can cause complications in the form of heart failure, stunted growth, liver problems, and even death[18]. Recent surveys show that thalassemia major occurs in infants between 300,000 and 400,000 babies born and it is known that up to 90% of births occur in poor and developing countries[5]. However, thalassemia is also very potential to occur in developed countries though.

The life of a thalassemia major patient is very dependent on blood transfusions of 1-3 bags per month. The existence of thalassemia major has a significant role in the blood supply needs of the Indonesian Red Cross. Doctors diagnose thalassemia through blood tests, including a complete blood test (CBC) and a special hemoglobin test. The CBC measures the amount of hemoglobin and various types of blood cells, such as red blood cells, in a blood sample. People who have thalassemia have fewer healthy red blood cells and less hemoglobin in their blood. People who have alpha or beta thalassemia traits may have red blood cells that are smaller than normal. The hemoglobin test measures the type of hemoglobin in a blood sample. People who have thalassemia have problems with the alpha or beta globin hemoglobin protein chains.

Usually moderate and severe thalassemia are diagnosed in early childhood. This is because the signs and symptoms such as severe anemia can be detected in the first 2 years of life. People who have mild thalassemia are usually diagnosed after routine blood tests show that they have anemia. Doctors will suspect thalassemia if someone is anemic and comes from an ethnic group at high risk of developing thalassemia.

The doctor will analyze the amount of iron in the blood to determine whether the anemia is due to iron deficiency or thalassemia. Iron deficiency anemia occurs because the body doesn't have enough iron to make hemoglobin. Anemia in thalassemia occurs due to problems with the alpha globin chain or hemoglobin beta globin, not due to iron deficiency. Through genes, thalassemia is passed from parents to children. Family genetic studies can also help diagnose the disorder.

Thalassemia in children can be detected after going through a series of blood tests in a medical laboratory or commonly called blood screening. The results of thalassemia screening contained several components of the blood test, including the complete blood count (CBC), High Performance Liquid Chromatography (HPLC) values and peripheral blood analysis. The CBC components consist of hemoglobin, hematocrit, erythrocyte, leukocyte, platelet and MC values, namely MCV, MCH, MCHC and RDW. While the HPLC components consist of HbA2 and HbF[19].

Many people think that the current cost of screening for thalassemia is quite expensive. In general, in Indonesia, not all health laboratories can provide blood screening services. Only health laboratories in big cities like Jakarta have blood screening testing facilities. In addition, it takes about 1 week to find out the results of a person's blood screening. The aim of this article is to propose a new model to determine thalassemia in children based on CBC values. Where almost every health laboratory in the area has this testing facility. Thus, thalassemia is known faster than the usual method so far. In this study, we used the fuzzy approach, which is one of the methods of artificial intelligence to solve the classification problem of thalassemia. Fuzzy approach is also known to be often used to solve various problems in health. In this study, we intend to implement a model that we developed using a fuzzy-based approach to classify thalassemia diseases based on CBC data. This article

describes how to build a model and implement it in software. We tested the model using some patient data at Abdoel Moeloek Hospital, Lampung Province. Thereafter we compare the test results with the opinion of pediatricians regarding thalassemia.

2. Literature Study

There have been many researched on applying artificial intelligence for medical field. The first article discusses the Efficiency of Data Types for Classification Performance of Machine Learning Techniques for Screening Thalassemia [20]. This article compares the MultiLayer Perceptron, K-Nearest Neighbors, Bayesian networks, NaiveBayes, and Multinomial Logistic Regression methods for the classification of thalassemia. The results obtained in this first article show that the MultiLayer Perceptron, K-Nearest Neighbors, and Bayesian networks have the best accuracy. The second article discusses the design of a fuzzy model for thalassemia disease diagnosis. The type of fuzzy used in diagnosing thalassemia is mamdani [21]. In this second article, there are 3 input variables used, namely hemoglobin, MCV and MCH. Meanwhile, the results show that the Fuzzy Inference System can be used for the classification of thalassemia. The third article discusses the Thalassemia risk prediction model using fuzzy inference systems[22]. In this third article, 3 input variables are used, namely symptoms, HbA and Hba. The results obtained are in the form of a prediction model for the severity of thalassemia patients.

Wantoro in his article entitled Implementation of fuzzy-profile matching in determining drug suitability for hypertensive patients implementing artificial intelligence techniques in the health sector. He made an application to make it easier for doctors to analyze hypertension data then provide possible drug recommendations for patients. Based on the results of comparisons with expert 1 and expert 2, the results of the average system testing are obtained with a value of 100% precision, recall 96.6% and accuracy 96.5%. Tests show that the application with a fuzzy logic approach can increase the efficiency of 9.5% compared to the interpolation weighting method[23].

3. Method

Knowledge based Development

Based on consultations conducted with experts and several literature reviews on blood screening, the criteria for developing a knowledge base for the classification of thalassemia are presented in Figure 1.

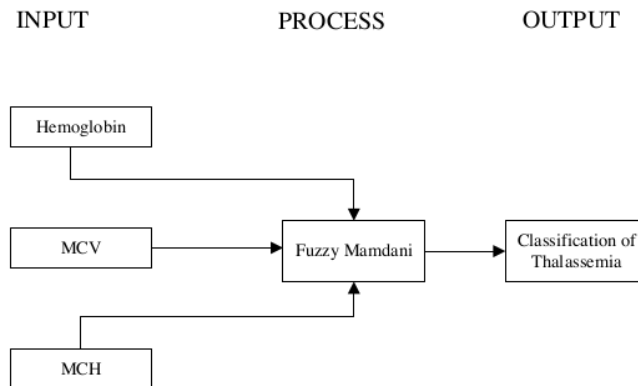


Figure 1 Knowledge for thalassemia classification

Variable

Experts think that the results of the CBC test may be used to determine thalassemia in children [18][24]. There are several values that are strongly thought to affect the results of thalassemia screening including hemoglobin, MCV, and MCH. Furthermore, these values are analyzed and used to determine the type of thalassemia in children. Meanwhile the variable names are presented in Table 1.

Table 1 Variables for determining the type of thalassemia

Hemoglobin	MCV	MCH
g/dL	fL	pg

The Evaluation of Thalassemias Classification

Based on the results of the CBC, data is needed as a basic reference for measuring the severity of thalassemia. For example, to find out a thalassemia major child has a hemoglobin value less than 7 g/dL, has an MCV value of 50 to 70 fL, and has an MCH value of 12 to 20 pg. In this case, we divide the data for each variable into four types including high, medium, low and very low. So that the ideal data used to determine the classification of thalassemia are obtained in Table 2.

Table 2 Classification data for thalassemia

Type	Hemoglobin	MCV	MCH
Mayor	<7[18]	50-70[18]	12-20[18]
Intermedia	7-10[18]	50-80[18]	16-24[18]
Minor	10-11	53-71[24]	17-25[24]
Not Thalassemia	10.8-12.8	73-101	23-31

Based on the data in Table 2, it is then used as a basis for determining the type of thalassemia using the Profile Matching evaluation method.

Input

Based on the thalassemia classification data in Table 2 and this data has been confirmed by a thalassemia specialist. Furthermore, the model for thalassemia are made using fuzzy Mamdani which is presented in Table 3.

Table 3 Model for input

Variable	Model
Hemoglobin	

Figure 2 Model for hemoglobin variable

Variable	Model
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MCV

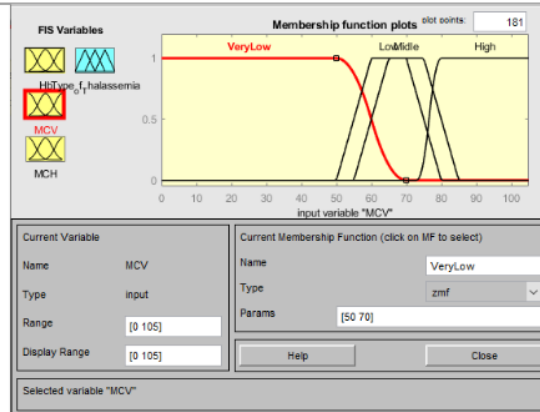


Figure 3 Model for MCV variable

MCH

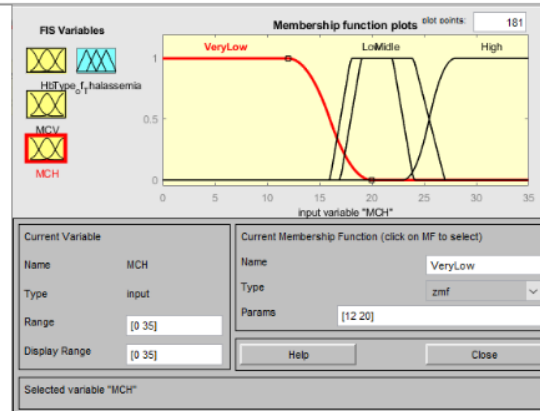


Figure 4 Model for MCH variable

Output

After the input model is created, then we also create a model to display the results of the analysis using Mamdani fuzzy. The output of the model is adjusted according to the rules as in Table 4.

Table 4 Classification of output

Linguistic Variable	Ranges	Fuzzy Set
Type_of_Thalassemia	≤ 3	Not Thalassemia
	$\geq 3; \leq 4$	Minor
	$> 3.5; \leq 8$	Intermedia
	≥ 7	Major

The output model is presented in Figure 5.

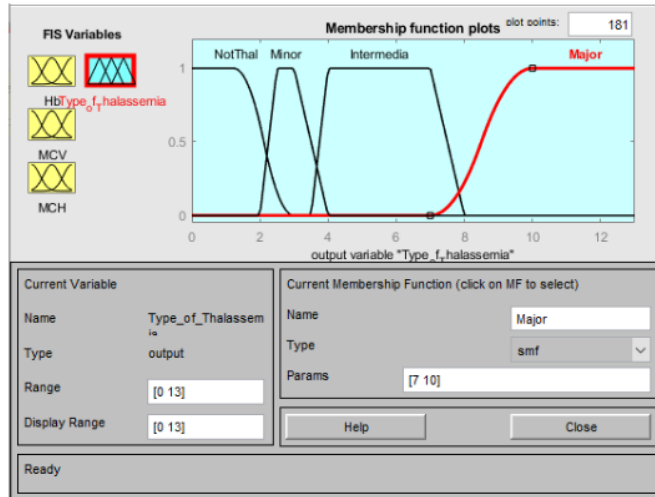


Figure 5 Model for output

Based on the membership function, the membership value for each variable is then calculated. As an example the membership values for Thalassemia Major. In this case we used data from pediatrician regarding thalassemia screening which are presented in Table 5.

Table 5 Fuzzy membership values for thalassemia major

P-001	Data	Membership values
Hemoglobin	5.0	Value (x) = (x<7) then x=1
MCV	66.7	Value (x) = (x>=50; x<=70) then x=1
MCH	20.3	Value (x) = (x>=12; x<=20) then x=0

Based on these data, it is then tested on a model that has been made. The results of the model trials are presented in Figure 6.

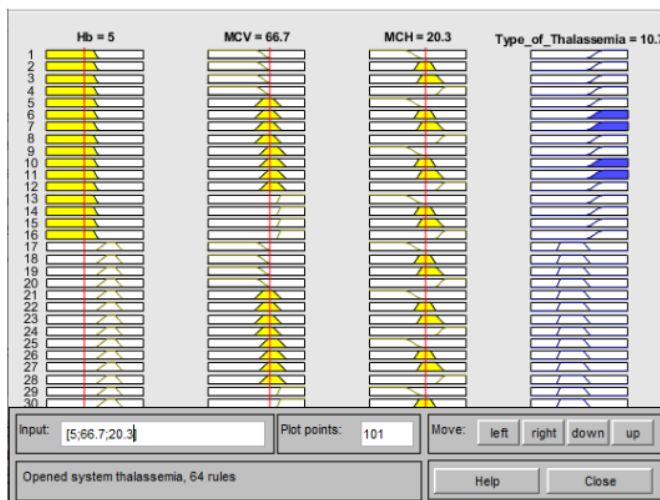


Figure 6 Tested the model using thalassemia major data

Based on the results of testing on the model, it can be seen that the results of the fuzzy calculation show a value of 10.7, so it can be concluded that the data includes thalassemia major.

4. Result and Discussions

After the model has been built, the next step is create a graphical user interface. In this case, we create a user interface as an implementation of the model we have developed. Next, we tested our proposed model. In this study we use test problem like as presented in Table 6.

Table 6 Test problem

CBC data			
	Hemoglobin (g/dL)	MCV (fL)	MCH (pg)
1	5.0	66.7	20.3
2	7.9	54.1	21.9
3	10.0	55.5	16.9
4	10.8	73.0	23.0

In testing the model we used 4 variables from CBC data. The first data contained several variable values including hemoglobin = 5.0 g/dL, MCV = 66.7 fL and MCH = 20.3 pg. The data input into the software as shown in Figure 7.

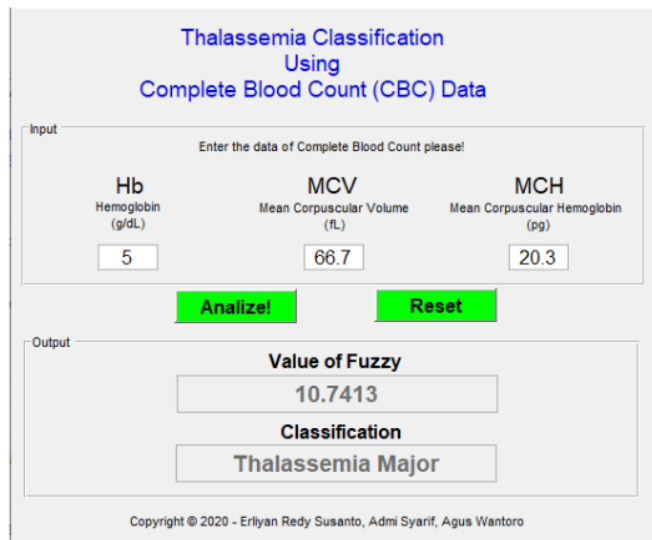
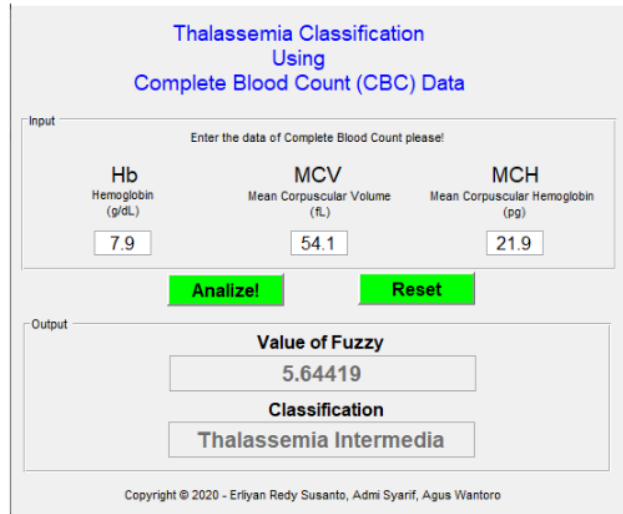


Figure 7 Display software on data testing for thalassemia major

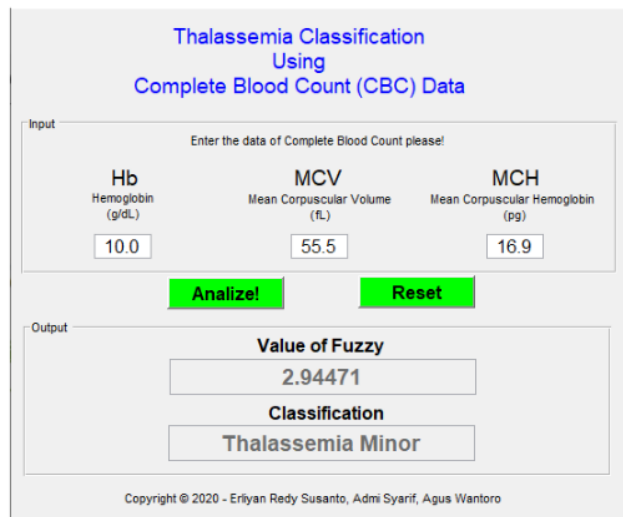
Based on the analysis results as Figure 7, the fuzzy value = 10.7413 is obtained so that the classification of thalassemia is classified as major. Second data contained several variable values including hemoglobin = 7.9 g/dL, MCV = 54.1 fL and MCH = 21.9 pg. The test results are presented in Figure 8.



The screenshot shows a web-based application titled "Thalassemia Classification Using Complete Blood Count (CBC) Data". It features an input section with three text boxes for Hb (7.9 g/dL), MCV (54.1 fL), and MCH (21.9 pg). Below the input boxes are two green buttons labeled "Analyze!" and "Reset". The output section displays "Value of Fuzzy" as 5.64419 and "Classification" as "Thalassemia Intermedia". A copyright notice at the bottom reads "Copyright © 2020 - Eriyan Redy Susanto, Admi Syarif, Agus Wantoro".

Figure 8 Display software on data testing for thalassemia intermedia

Based on the analysis results as Figure 8, the fuzzy value = 5.64419 is obtained so that the classification of thalassemia is classified as intermedia. Third data contained several variable values including hemoglobin = 10.0 g/dL, MCV = 55.5 fL and MCH = 16.9 pg. The test results are presented in Figure 9.



The screenshot shows the same web-based application as Figure 8. The input section now has Hb (10.0 g/dL), MCV (55.5 fL), and MCH (16.9 pg). The "Analyze!" and "Reset" buttons are still present. The output section displays "Value of Fuzzy" as 2.94471 and "Classification" as "Thalassemia Minor". The copyright notice at the bottom remains the same: "Copyright © 2020 - Eriyan Redy Susanto, Admi Syarif, Agus Wantoro".

Figure 9 Display software on data testing for thalassemia minor

Based on the analysis results as Figure 9, the fuzzy value = 2.94471 is obtained so that the classification of thalassemia is classified as minor. Fourth data contained several variable values including hemoglobin = 10.0 g/dL, MCV = 55.5 fL and MCH = 16.9 pg. The test results are presented in Figure 10.

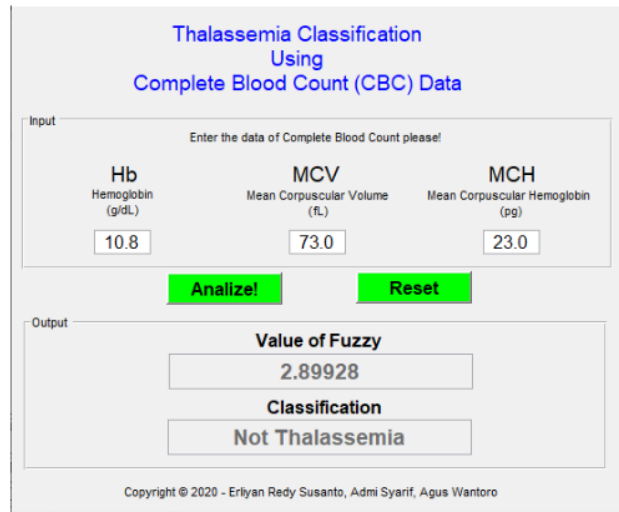


Figure 10 Display software on data testing for not thalassemia

Based on the analysis results as figure 10, the fuzzy value = 2.89928 is obtained so that the classification of thalassemia is classified as not thalassemia. The test result data is then matched with the opinion of the pediatrician. The test results on the four data are presented in Table 7.

Table 7 The Result of Testing

No	Variables			Value of Fuzzy	Classification	Expert Judgement	Result
	Hemoglobin	MCV	MCH				
1	5.0	66.7	20.3	10.7413	Major	Major	Correct
2	7.9	54.1	21.9	5.64419	Intermedia	Intermedia	Correct
3	10.0	55.5	16.9	2.94471	Minor	Minor	Correct
4	10.8	73.0	23.0	2.89928	Not Thallasemia	Not Thallasemia	Correct

The results of testing on 4 CBC data show that the model we developed is proven to be in accordance with the opinion of pediatricians regarding thalassemia.

5. Conclusion

Finally, the results show that the proposed model works well to determine thalassemia. The type of thalassemia is determined based on the results of the CBC examination based on the hemoglobin, MCV and MCH values. We have 4 output models, namely Major, Intermedia, Minor and Not Thalassemia. The results of the model prediction against 4 data were tested in accordance with the opinion of doctors about thalassemia. In order to know better accuracy, our model needs to be tested with more real data.

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