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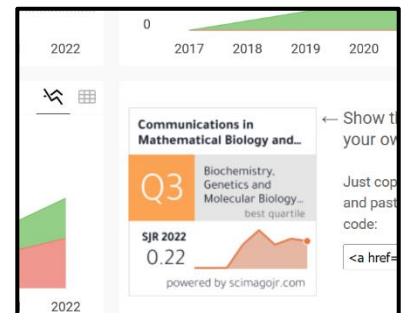
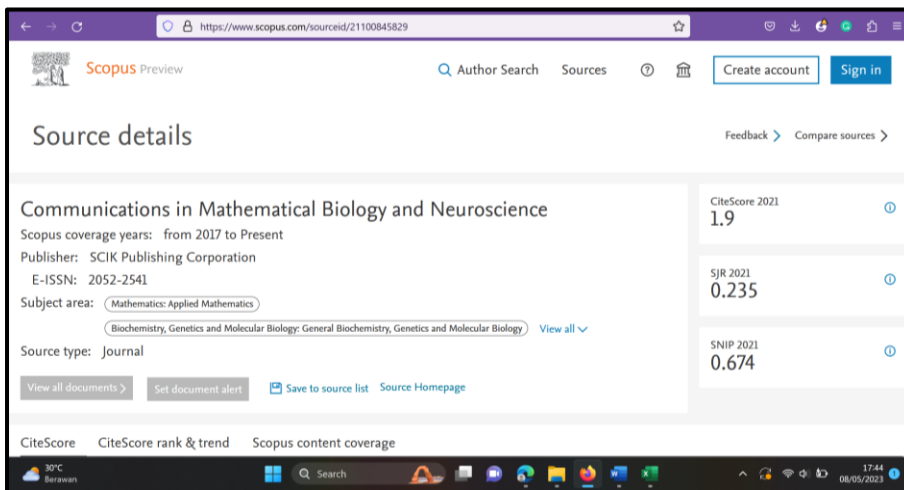
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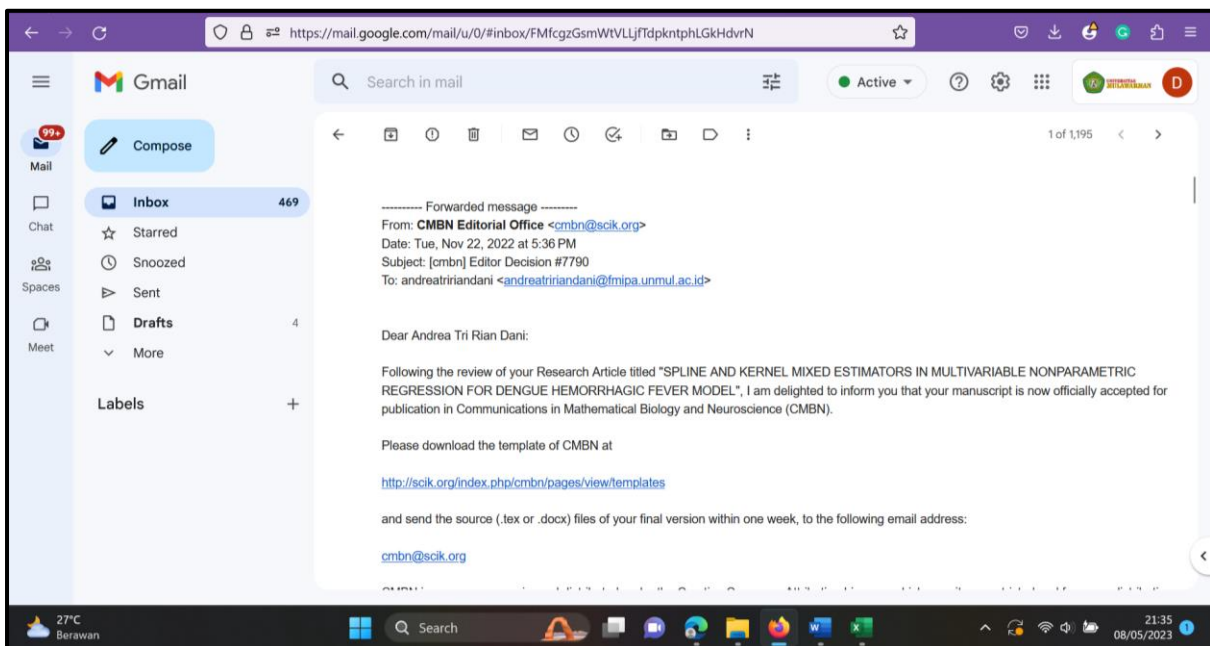
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SPLINE AND KERNEL MIXED ESTIMATORS IN MULTIVARIABLE NONPARAMETRIC REGRESSION FOR DENGUE HEMORRHAGIC FEVER MODEL

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Abstract: This article discusses statistical innovations implemented in the health sector. The research is being
conducted on the treatment and prevention of Dengue Hemorrhagic Fever (DHF), focusing on the factors contributing
to the increase in DHF. Create a nonparametric regression model with a mixed estimator, truncated spline, and
Gaussian Kernel to estimate the regression curve. In multiple nonparametric regression, this method can handle
differences in data patterns between predictors. Truncated splines are polynomial segments with segmented and
continuous properties. Truncated splines contain knot points that can locate their estimated data no matter where the
data pattern moves. In addition, the Gaussian Kernel estimator is dependent on bandwidth, which regulates the
regression curve's smoothness. The mixed estimators of truncated spline and Gaussian Kernel could model DHF cases
according to an empirical analysis of DHF data. The most effective model has a Coefficient of Determination (R^2) of
88.46%. Simultaneous hypothesis testing indicates that the model contains at least one significant predictor variable.

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27 **Keywords:** dengue hemorrhagic fever (DHF) modeling; Gaussian kernel; maximum likelihood (ML); mixed
28 estimators; nonparametric regression; truncated spline.

29 **2020 AMS Subject Classification:** 92C60.

30

31 1. INTRODUCTION

32 The relationship between the response and predictor variable, whose purpose is unknown, can
33 be identified statistically using nonparametric regression [1], [2]. Nonparametric regression is not
34 rigid in defining the regression function [3], does not require certain assumptions like linear
35 regression, where the error must be normally distributed, and does not force the regression curve
36 to be linear. This is in contrast to parametric regression, which causes the regression curve to follow
37 a specific model, such as a linear model. The observational data determine the advantage of
38 nonparametric regression, the regression curve without being forced to adjust a specific function
39 [4], [5]. Nonparametric regression assumes that the data derive their form of estimation from the
40 regression curve without regard for the researcher's subjectivity [6]. As a result, the nonparametric
41 regression model approach is both adaptable and objective [2], [3].

42 The estimator approach used in nonparametric regression includes truncated spline and Kernel.
43 Truncated spline is polynomial pieces that have segmented and continuous properties [7], [8]. One
44 of the advantages of the truncated spline is that this model tends to find its own estimate of the
45 data wherever the data pattern moves [9], [10]. This advantage occurs because, in the truncated
46 spline, knot points indicate changes in data behavior patterns [4], [5]. While the Kernel estimator
47 has the advantage that it is flexible [11], the mathematical form is easy and can achieve a relatively
48 fast level of convergence [12]. The Kernel approach depends on bandwidth, which controls the
49 smoothness of the estimation curve [5], [12]–[14]. Estimating the regression curve with the Kernel
50 estimator approach adjusts from the value of the smoothing parameter λ .

51 According to Budiantara et al. [15], the nonparametric and semiparametric regression models
52 developed by the researchers so far, if explored more deeply, basically there are very heavy and
53 basic assumptions in the model. Each predictor in the multi-predictor nonparametric regression is

54 considered to have the same pattern, so the researchers force the use of only one form of the model
55 estimator for all predictor variables. Therefore, using only one form of the estimator in various
56 forms of different data relationship patterns will certainly result in the resulting estimator not being
57 compatible with the data pattern. As a result, the estimation of the regression model is not good
58 and produces a significant error. Therefore, to overcome this problem, several researchers have
59 developed a nonparametric mixed regression curve estimator in which an appropriate curve
60 estimator approximates each data pattern in the nonparametric regression model. There are several
61 studies that have developed and reviewed mixed estimator models, including [1], [16]–[19].

62 Dengue Hemorrhagic Fever is one of the problems in Indonesia's health sector. DHF is caused
63 by the bite of the *Aedes Aegypti* mosquito [20], which usually attacks tropical and subtropical
64 areas of the world [21], [22], one of which is Indonesia. Based on data from the World Health
65 Organization (WHO), Indonesia has the 2nd rank with the most significant DHF cases among 30
66 endemic areas (Ministry of Health, 2018). In 2020, there were 108303 patients with DHF cases,
67 and 747 died. Meanwhile, the number of DHF cases in 2021 was 73518, and 705 died. The number
68 decreased by 32.12% compared to the previous year, but this case still needs special attention.

69 Based on the description that has been explained, so the purpose of this study is to conduct a
70 study of the nonparametric regression Mixed Estimator of Truncated Spline and Gaussian Kernel
71 (MTs-GK) model in the additive multi-predictor nonparametric model and the implementation of
72 the model in the case study of Dengue Hemorrhagic Fever (DHF) with a special issue of the factors
73 that influence the increase in DHF.

74

75 **2. PRELIMINARIES**

76 *A. Mixed Estimators of Truncated Spline and Gaussian Kernel*

77 A mixed estimator is a multi-predictor nonparametric regression model that uses two or more
78 types of estimators to approximate the regression curve [15], [16]. Budiantara et al. [15] were the
79 first to develop a mixed truncated spline and Kernel nonparametric regression model. A Mixed
80 estimator is a model approach in nonparametric regression where more than one estimator is used
81 [18], [23], [24]. The form of the regression curve for each relationship between the predictor and

82 the response variable will be approximated by two or more estimators according to the
83 characteristics of the relationship [25].

84 For example, given paired data (x_i, v_i, y_i) and the relationship between predictor variables
85 (x_i, v_i) with response variable (y_i) following a nonparametric regression model:

$$y_i = \mu(x_i, v_i) + \varepsilon_i \quad (1)$$

86 $\mu(x_i, v_i)$ is the regression curve, with assumed to be unknown, smooth, and follows an additive
87 model so that $\mu(x_i, v_i)$ we can write in the form in Equation (2).

$$\mu(x_i, v_i) = m(x_i) + h(v_i) \quad (2)$$

88 Based on Equation (2), the regression curve $m(x_i)$ will be estimated with a truncated spline
89 estimator, while the regression curve $h(v_i)$ with a Kernel estimator.

90 The truncated spline estimator is a segmented polynomial model [26], [27]. For example, given
91 paired data (x_i, y_i) where the relationship from the predictor (x_i) and response variable (y_i)
92 follow a truncated spline nonparametric regression model [7], [28].

$$y_i = \beta_0 + \sum_{j=1}^m \sum_{p=1}^q \beta_{jp} x_{pi}^j + \sum_{k=1}^r \sum_{p=1}^q \beta_{(m+k)p} (x_{pi} - K_{kp})_+^m + \varepsilon_i \quad (3)$$

93 The truncated function is:

$$(x_{pi} - K_{kp})_+^m = \begin{cases} (x_{pi} - K_{kp})^m & x_p \geq K_{kp} \\ 0 & x_p < K_{kp} \end{cases}$$

94
95 In matrix form, Equation (3) is as follows:

$$\mathbf{y} = \mathbf{X}(K)\boldsymbol{\beta} + \boldsymbol{\varepsilon} \quad (4)$$

96 As a result, the regression curve estimation using the truncated spline estimator can be written in
97 Equation (5).

$$\hat{\mathbf{m}}(x) = \mathbf{X}(K)\hat{\boldsymbol{\beta}} \quad (5)$$

98 Kernel estimator has a good ability to model data that does not have a certain pattern [11], [29],

99 [30]. For example, given paired data, (v_i, y_i) where is the relationship between the predictor
 100 (v_i) and with response variable (y_i) following the Kernel nonparametric regression model.

$$y_i = h(v_i) + \varepsilon_i \quad (6)$$

101 The regression curve $h(v_i)$ will be estimated with the Kernel estimator in Equation 7.

$$\begin{aligned} \hat{h}_\lambda(v_i) &= \frac{1}{n} \sum_{i=1}^n \frac{K_\lambda(v-v_i)}{\frac{1}{n} \sum_{i=1}^n K_\lambda(v-v_i)} y_i \\ &= \frac{1}{n} \sum_{i=1}^n R_{\lambda i}(v) y_i \end{aligned} \quad (7)$$

102 With

$$103 \quad R_{\lambda i}(v) = \frac{K_\lambda(v-v_i)}{\frac{1}{n} \sum_{i=1}^n K_\lambda(v-v_i)}$$

$$104 \quad K_\lambda(v-v_i) = \frac{1}{\lambda} K\left(\frac{v-v_i}{\lambda}\right)$$

105 K is an abbreviation for Kernel Function. The Gaussian Kernel function is used in this research:

$$K(v) = \frac{1}{\sqrt{2\pi}} \exp\left(\frac{1}{2}(-v^2)\right) \quad (8)$$

106 Based on equations (7) and (8), we can write them in matrix form as:

$$\hat{\mathbf{h}}(v) = \mathbf{D}(\lambda)\mathbf{y} \quad (9)$$

107 Furthermore, based on Equation (2) and the form of each estimator in Equations (5) and (9), we
 108 can write:

$$\mathbf{y} = \mathbf{X}(K)\boldsymbol{\beta} + \mathbf{D}(\lambda)\mathbf{y} + \boldsymbol{\varepsilon} \quad (10)$$

109 Vector $\boldsymbol{\varepsilon}$ has a size $(n \times 1)$, so that based on Equation (10), then:

$$\begin{aligned} \boldsymbol{\varepsilon} &= \mathbf{y} - (\mathbf{X}(K)\boldsymbol{\beta} - \mathbf{D}(\lambda)\mathbf{y}) \\ &= (\mathbf{I} - \mathbf{D}(\lambda))\mathbf{y} - \mathbf{X}(K)\boldsymbol{\beta} \end{aligned} \quad (11)$$

110 The parameter estimation of $\boldsymbol{\beta}$ used the Least Squares (LS). The estimated results for $\hat{\boldsymbol{\beta}}$ is:

$$\hat{\boldsymbol{\beta}} = \left(\mathbf{X}(K)^T \mathbf{X}(K) \right)^{-1} \mathbf{X}(K)^T (\mathbf{I} - \mathbf{D}(\lambda)) \mathbf{y} \quad (12)$$

111 Equation (12) can be summarized as:

$$\hat{\boldsymbol{\beta}} = \mathbf{A}(K, \lambda) \mathbf{y} \quad (13)$$

112 Where $\mathbf{A}(K, \lambda) = \left(\mathbf{X}(K)^T \mathbf{X}(K) \right)^{-1} \mathbf{X}(K)^T (\mathbf{I} - \mathbf{D}(\lambda))$.

113 In Equation (5), the regression curve estimation is written using the truncated spline estimator

114 is $\hat{\mathbf{m}}(x) = \mathbf{X}(K) \hat{\boldsymbol{\beta}}$, then based on Equation (12), we can write:

$$\begin{aligned} \hat{\mathbf{m}}(x) &= \mathbf{X}(K) \hat{\boldsymbol{\beta}} \\ &= \mathbf{X}(K) \left[\left(\mathbf{X}(K)^T \mathbf{X}(K) \right)^{-1} \mathbf{X}(K)^T (\mathbf{I} - \mathbf{D}(\lambda)) \mathbf{y} \right] \end{aligned} \quad (14)$$

115 A brief summary of Equation (14) is:

$$\hat{\mathbf{m}}(x) = \mathbf{S}(K, \lambda) \mathbf{y} \quad (15)$$

116 According to Equation (15) and the shape of the estimator for each component in Equation (9) and

117 (15), the mixed estimator of truncated spline and Gaussian Kernel will be obtained as follows:

$$\begin{aligned} \hat{\boldsymbol{\mu}}(x, v) &= \hat{\mathbf{m}}(x) + \hat{\mathbf{h}}(v) \\ &= (\mathbf{S}(K, \lambda) + \mathbf{D}(\lambda)) \mathbf{y} \\ &= \mathbf{B}(K, \lambda) \mathbf{y} \end{aligned} \quad (16)$$

118 Matrix $\mathbf{B}(K, \lambda)$ very dependent on $\mathbf{S}(K, \lambda)$ which is a component of the truncated spline

119 estimator, where the optimal location and number of knot points must be determined, and matrix

120 $\mathbf{D}(\lambda)$, which is a component of the Kernel estimator, needs to find the correct bandwidth value.

121 In this study, the method used to select the optimal knot point and bandwidth is Unbiased Risk

122 (UBR) [4], [17], [18] with the formula in Equation (17).

$$UBR(K_{opt}, \lambda_{opt}) = \frac{1}{n} \left(\left\| (\mathbf{I} - \mathbf{B}(K, \lambda)) \mathbf{y} \right\|^2 + \frac{\hat{\sigma}^2}{n} \text{trace}[\mathbf{I} - \mathbf{B}(K, \lambda)]^2 + \frac{\hat{\sigma}^2}{n} \text{trace}[\mathbf{B}(K, \lambda)^2] \right) \quad (17)$$

123 Where:

$$124 \hat{\sigma}^2 = \frac{\|(\mathbf{I} - \mathbf{B}(K, \lambda))\mathbf{y}\|^2}{tr((\mathbf{I} - \mathbf{B}(K, \lambda))\mathbf{y})}$$

125 *B. Simultaneous Testing Hypothesis*

126 Hypothesis testing can only be done on the truncated spline estimator component based on the
127 mixed estimator model in Equation (11) using the Likelihood Ratio Test (LRT).

128 Hypothesis Formulation:

$$129 H_0 : \beta_1 = \beta_2 = \dots = \beta_{(m+r)p} = 0$$

$$130 H_1 : \text{there is at least one } \beta_j \neq 0, \quad j = 1, 2, \dots, (m+r)p$$

131 In summary, the following ANOVA table gives the simultaneous hypothesis testing process for the
132 parameters in the mixed estimator model.

133

Source	Degree of Freedom (df)	Sum of Squares (SS)	Mean Squares (MS)	F-Test
Regression	$(m+r)p$	$SSR = \sum_{i=1}^n (\hat{y}_i - \bar{y})^2$	$MSR = \frac{SSR}{(m+r)p}$	$F_{\text{test}} = \frac{MSR}{MSE}$
Error	$n - ((m+r)p) - 1$	$SSE = \sum_{i=1}^n (y_i - \hat{y}_i)^2$	$MSE = \frac{SSE}{n - ((m+r)p) - 1}$	
Total	$n - 1$	$SST = \sum_{i=1}^n (y_i - \bar{y})^2$		

134

TABLE 1. ANOVA

135

136 Under the null hypothesis H_0 , test statistics from F_{test} following the F Distribution, with the
137 degree of freedom (df) is $((m+r)p, n - ((m+r)p) - 1)$.

138

139 3. RESEARCH METHODOLOGY

140 A. Data Sources

141 The research used secondary data from Wahab Syahrani General Hospital (AWS Hospital) in
142 Samarinda. The variables of this study are described in Table 2.

143

Variable	Notation	Description	Unit	Data Scale
DHF patient's platelet count when first blood check	y	The platelet count of DHF patients when they first do a blood check	μl	
Number of Hematocrits	x_1	The number of hematocrits found in patients with DHF	%	Continuous
Number of Hemoglobin	x_2	The number of hemoglobin cells found in patients with DHF	g/dL	
Number of Leukocyte	x_3	The number of leukocytes found in patients with DHF	g/dL	

144 TABLE 2. Description of Study Variables and Unit Data

145

146 B. Data Analysis Technique

147 To answer the research purposes, necessary to develop research steps. The research steps used
148 in this study are:

- 149 1. Create a scatter plot to show the relationship between each predictor variable and the
150 response.
- 151 2. Determine the predictor variables for the truncated spline and Kernel components.
- 152 3. Modeling case data of patients with dengue fever with the response variable (y) being the
153 patient platelet count using a mixed truncated spline and Gaussian Kernel estimator model
154 based on Equation (16).

DENGUE HEMORRHAGIC FEVER MODEL

- 155 4. Select the optimal knot point and bandwidth based on the minimum UBR value with the
 156 Formula in Equation (17). Each predictor variable in this study has the same number of knot
 157 points (1 to 3). The bandwidth values tested are in the interval of 0.05 to 5.
 158 5. Determine the best model of the mixed estimator truncated spline and Kernel based on the
 159 minimum UBR value and then calculate the Coefficient of Determination (R^2) value.
 160 6. Simultaneous hypothesis testing for the best model based on ANOVA in Table 1.

161

162 **4. MAIN RESULTS**

163 In this section, we will explain the results of the study mixed estimator truncated spline and
 164 Gaussian Kernel applied to data on the platelet count of DHF patients.

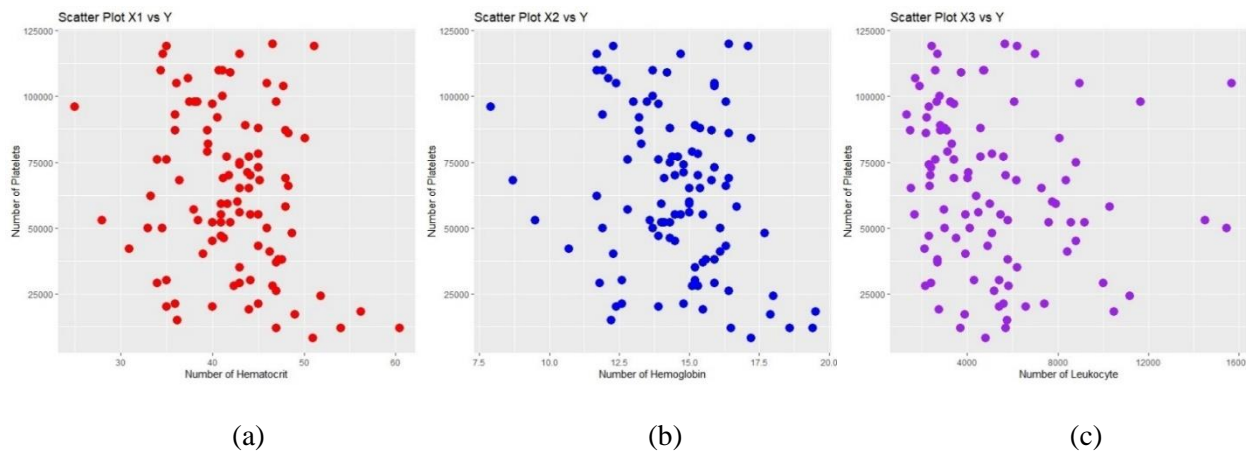
165 *A. Scatter Plot*

166 The first step in the modeling process using a mixed estimator is creating a scatter diagram for
 167 each variable. The scatter diagram for each predictor variable to the response variable is shown in
 168 Figure 1.

169

170

FIGURE 1. Scatter Plot



171

172

173

174 Based on Figure 1, it can be determined which type of estimator will be used for each predictor
 175 variable. A more detailed summary of the results of determining the estimator is presented in Table

176 3.

Variable	Notation	Description	Estimator
Predictor	x_1	Number of Hematocrits	Truncated Spline
	x_2	Number of Hemoglobin	
	v_1	Number of Leukocyte	Kernel

177 TABLE 3. Components of Truncated Spline and Gaussian Kernel Estimator

178 *B. Modeling using Mixed Estimator Truncated Spline and Gaussian Kernel*

179 The best model from the mixed estimator truncated spline and Gaussian Kernel was selected
 180 by comparing the smallest Unbiased Risk (UBR) value among various models based on the number
 181 of knot points and bandwidth. In this study, the number of knots used is the same for each predictor
 182 variable, namely, 1 to 3. The bandwidth values tested are in the interval of 0.05 to 5. The modeling
 183 results using a mixed estimator truncated spline and Gaussian Kernel are in Table 4.

184

Number of Knot Point	Knot Point Location		Bandwidth	UBR Value
	x_1	x_2	v_1	
1 knot	37.21	11.90	0.55	97.46
2 knots	35.98	11.50	2.80	96.00
	37.21	11.90		
3 knots	44.53	14.30	2.55	99.26
	45.75	14.70		
	46.97	15.10		

185 TABLE 4. Summary of Modeling Results

186 Based on Table 4, the minimum UBR value is 96.00 with an optimal bandwidth of 2.80, and the
 187 optimal knot point location for each predictor variable modeled with a truncated spline estimator
 188 is:

189 Variable x_1 (Number of Hematocrit)

190 $K_1 = 35.98 \quad K_2 = 37.21$

191 Variable x_2 (Number of Hemoglobin)

192 $K_1 = 11.50 \quad K_2 = 11.90$

193 Using the knot point and bandwidth optimal, a nonparametric regression model with mixed
194 estimator truncated spline and Gaussian Kernel is written in Equation (18).

$$\begin{aligned}
\hat{y}_i &= \hat{\beta}_0 + \hat{\beta}_{11}x_{1i} + \hat{\beta}_{12}x_{2i} + \hat{\beta}_{21}(x_{1i} - K_{11})_+ + \hat{\beta}_{31}(x_{1i} - K_{21})_+ + \hat{\beta}_{22}(x_{2i} - K_{12})_+ + \\
&= \hat{\beta}_{32}(x_{2i} - K_{22})_+ + \frac{1}{100} \sum_{i=1}^{100} \frac{K_{\lambda}(v - v_i)}{\frac{1}{100} \sum_{i=1}^{100} K_{\lambda}(v - v_i)} y_i \\
\hat{y}_i &= 22487.37 - 541.58x_{1i} + 284.55x_{2i} + 2718.32(x_{1i} - K_{11})_+ - \\
&= 1603.31(x_{1i} - K_{21})_+ - 16156.97(x_{2i} - K_{12})_+ + \\
&= 13711.35(x_{2i} - K_{22})_+ + \frac{1}{100} \sum_{i=1}^{100} \frac{K_{2.80}(v - v_i)}{\frac{1}{100} \sum_{i=1}^{100} K_{2.80}(v - v_i)} y_i \tag{18}
\end{aligned}$$

195 The coefficient of determination (R^2) of this model is 88.46%. This means that 88.46% of the
196 platelet count of patients with Dengue Hemorrhagic Fever (DHF) can be explained by the variables
197 Number of Hematocrit, Number of Hemoglobin, and Number of Leukocyte in the mixed estimator
198 model of truncated spline and Gaussian Kernel with two-knot points and optimal bandwidth.

199 *C. Simultaneous Testing Hypothesis*

200 The next step will be to simultaneously test the hypothesis for the parameters in the model based
201 on the best model from the mixed estimator truncated spline and Gaussian Kernel.

$$202 \quad H_0 : \beta_1 = \beta_2 = \dots = \beta_{(m+r)p} = 0$$

$$203 \quad H_1 : \text{there is at least one } \beta_j \neq 0, \quad j = 1, 2, \dots, (m+r)p$$

204 The ANOVA table for the results of hypothesis testing is presented in Table 5.

205

Source	Degree of Freedom (df)	Sum of Squares (SS)	Mean Squares (MS)	F-Test	P-Value
Regression	6	82542921847	13757153641	118.78	2.26e-41
Error	93	10771268700	115820094		
Total	99	91430590000			

206

TABLE 5. Summary of ANOVA

207 Based on Table 5, it can be seen that the F_{test} (118.78) is greater than the $F_{(0.05;6;93)}$ (2.19) or P-
 208 Value (2.26e-41) is smaller than the value $\alpha = 0.05$, so the decision is rejected H_0 . This means
 209 simultaneously; there is at least one $\beta_j \neq 0$ or at least one significant predictor variable in the
 210 model.

211

212 5. CONCLUSION

213 A mixed estimator truncated spline and Gaussian Kernel model was used to successfully model
 214 the cases of Dengue Hemorrhagic Fever (DHF) patients. A nonparametric regression model of
 215 mixed estimator truncated spline and Gaussian Kernel with 2-knot points and optimal bandwidth
 216 is the best model based on the lowest UBR value.

$$\begin{aligned}
 \hat{y}_i &= 22487.37 - 541.58x_{1i} + 284.55x_{2i} + 2718.32(x_{1i} - K_{11})_+ - 1603.31(x_{1i} - K_{21})_+ - \\
 &= 16156.97(x_{2i} - K_{12})_+ + 13711.35(x_{2i} - K_{22})_+ + \frac{1}{100} \sum_{i=1}^{100} \frac{K_{2.80}(v - v_i)}{\frac{1}{100} \sum_{i=1}^{100} K_{2.80}(v - v_i)} y_i
 \end{aligned}$$

218 The best model's coefficient of determination (R^2) is 88.46%. Based on the results of simultaneous
 219 hypothesis testing, it can be concluded that simultaneously there is at least one significant predictor
 220 variable in the model.

221

222 CONFLICT OF INTERESTS

223 The authors declare that there is no conflict of interest.

224

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DENGUE HEMORRHAGIC FEVER MODEL

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