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Contents
Volume 10, Number 1 January 2019
A study to Assess the Knowledge Regarding the Care of Patients on Mechanical Ventilation and Prevention of VAP among Nursing Students of a Selected Nursing College
2. A Study on Job Satisfaction among Employees in Quick Service Restaurants
3. Comparitive Evaluation of the Fluoride Release and Rechargability of Chitosan Modified Glass Ionomer Cement and a Glass Ionomer Cement—An in Vitro Study
4. A Study on Determinants of Agricultural Productivity in Tamilnadu
5. Susceptibility of Gender Entrepreneurship Gap in India–A Preview
6. Analysis of Medical Tourism and its Economic Impact
7. Occurrence of Menstrual Irregularities among Adolescent Girls in Selected Area, Dehradun, Uttarakhand 2 Jyoti Kandpal, Mugdha Devi Sharan Sharma, Upma George
8. Mother's Knowledge on Nutrition and Incidence of Malnutrition
9. Mortality Pattern amongst Patient Admitted in Tertiary Health Care Center, Rajnandgaon (C.G.)
10. Effectiveness of Self-Instructional Module on Knowledge Regarding Post-Partum Psychiatric Disorders 4 <i>Leeja Bonny Thomas, Anusha Pradhan</i>
11. A Study to Evaluate the Effectiveness of Structured Teaching Programme on Obesity & its Consequences among Adolescents in Selected Private High Schools in Pune
12. Assessment of the Awareness about Effects of Cell Phone Radiations amongst Students at Symbiosis International (Deemed University) Hill Base Campus
13. A Study to Estimate the Level of Physical Activity and Perceived Benefits and Barriers to Exercise among Women in Coastal Karnataka

14.	An Empirical Study	63
15.	Confocal LASER Scanning Microscopy (CLSM) for Evaluation of Endodontic Microflora-A Review Laxmish Mallya, Kundabala M, Vinod Jathanna	69
16.	Effective Recruitment and Selection System for the IT Software Industry in India	74
17.	A Study on Innovative Recruitment Techniques and It's Impact on Job Seekers	79
18.	Emotional Intelligence and Performance of Manager in Manufacturing Industries (With special reference to Automobile Industry)	85
19.	Mechanically Induced Stump Dermatoses: High Prevalence Concern and Measures of Prevention	88
20.	A Study on Universal Precautions and Needle Stick Injuries among Nursing Staff in a Tertiary Care Hospital, Davangere	93
21.	Pattern of Employment and Consumption Expenditure in India	98
22.	A Study on Women Domestic Workers in M.g.r.nagar in Kanchipuram District	103
23.	Influence of Emotional Intelligence on Employee Performance among Seleced Restaurants, Chennai 1 V. Krishna Priya	106
24.	Preferences and Problems of Agri-Based Enterprises of Guntur (A.P., India): An Empirical Study of Farmers Advisory Committees Under Atma	111
25.	Effects of Strength Training Exercises on Physical Parameters and Quality of Life among Older Adults in Selected Geriatric Homes in Kerala, India	116
26.	Study of MRSA and ESBL Organisms Isolated from Infected Wounds Suresh P, V. Sreenivasulu Reddy, V. Praveen Kumar, P. Vamsimuni Krishna	121
27.	Prevalence of Angles Malocclusion Traits in 7-16-Year-old School Children of Mewar Region, India 1 Pradeep Vishnoi, Tarulatha R Shyagali, Prabhuraj Kambalyal, Deepak P Bhayya, Rutvik Trivedi, Jyoti Jingar	125
28.	A Study on Marketing Prospects in Promoting Cultural Tourism in Tamil Nadu	131
29.	An Objective and Subjective Evaluation of Dental Implant Impressions using Vinylsiloxanether and Polyether Impression Materials—An <i>in Vivo</i> Study	135

30.	Comparison of Serum Calcium Levels in Euthyroid, Subclinical and Overt Hypothyroid Women in the Tribal Belt of West Midnapore, West Bengal	139
31.	Role of Whistle Blowers in Health Care Industry: An Empirical Study	142
32.	Management of Talons Cusp in a Primary Maxillary Central Incisor: A Rare Case Report	147
33.	Two Way Analysis of GST : With Reference to Healthcare and Pharma Sector	151
34.	Comparison of Serological Tests in the Diagnosis of Leptospirosis in a Tertiary Care Hospital at Chidambaram, Tamilnadu, India	
35.	Traditional Use of Medicinal Plants in Puducherry for Treatment of Urinary Tract Disorders	160
36.	Awareness of Cervical Cancer among HIV Positive Women in Southern India	165
37.	Correlates of Hope and Depression among People Living with Human Immunodeficiency Virus in Chhattisgarh State Bansh Gopal Singh, Deepak Pandey	170
38.	Vitamin C Intake Improve the Anthropometric Measurements, Lipid Profile and Atherogenic Indices in Obese and Non Obese Females	177
39.	Effect of Vitamin C Supplementation on Insulin Resistance, β-cell Function and Insulin Sensitivity in Obese and Non Obese Individuals	183
40.	Serum VEGF and TNF-α Correlate Bacterial Burden in Pulmonary Tuberculosis	189
41.	Online Sales Promotions of Herbal Products and Its Effectiveness towards Tanisha.com	195
42.	Effects of Strength Training Exercises on Physical Parameters and Quality of Life among Older Adults in Selected Geriatric Homes in Kerala, India	201
43.	Impact of Quality of Work Life Dimensions on Organizational Performance: With reference to Jute Industry in Andhra Pradesh and West Bengal, India	206
44.	A Study on Customer Preferences on Green Marketing	211

45.	Why Physician's Keep Coming Back to Telemedicine: Predicting Using Unsupervised Learning Preeti Y Shadangi, Manoranjan Dash, Sunil Kar	216
46.	Practicing the Strategies of Interpersonal Conflicts Management in Business Organisations to Accede Development and Effectiveness in Personal Health	222
47.	Comparative Performance Analysis of Selected Large Cap Mutual Funds in India	227
48.	Study of Morbidity Pattern among Women Beedi Rollers Residing in Urban Area of Mangalore	233
49.	Perception, Attitude and Practices Regarding Climate Change among College Students in Coastal South India	
50.	Evaluation of Thyroid Hormone Levels Before and After Thyroidectomy	242
51.	Impact of Ambidextrous Leadership on Firm Performance: A Study on IT Sector in Hyderabad, India . Sahyaja Ch., K. S. Sekhara Rao	247
52.	Prevalence of Halitosis among Preclinical Medical and Dental Students	253
53.	The Effectiveness of Information, Education and Communication on Knowledge, Attitude, Practice Regarding Obesity among Adolescents at Selected Government Schools in Kancheepuram District Shanthi M., C. Kanniammal, Jaideep Mahendra, G. Valli	257
54.	Various Online Marketing and Promotions Strategies to Improve the Validation Towards the Organic Products in the Pharmaceutical Sectors	263
55.	Effect of Bidirectional Dyadic Association on Anxiety and Self Esteem among Patients Undergoing Mastectomy	270
56.	An Empirical Study to Improve the Service Quality for Geriatric Patients in a Tertiary Care Hospital Swathi TM , Khyathi GV	276
57.	Effects of Nudge and Purchase Intention in Online Purchasing of Electronic Products	282
58.	Discharge Planning Model with Approach of Method in Improving Patients' Readiness for Discharge in Hospitals	288
59.	Study of Model Climate Maps Using Geographic Information System (G.I.S)	293
60.	Impact of Terrorism Act on Child Psychology and Post-Traumatic Stress Disorder	298

61.	The Sociopragmatics of Preaching in an American Christian Sermon	303
62.	The Effectiveness of Extract Klika Streculiapopulifolia Cream on the Collagen of Albino Mice against Ultraviolet B Radiation	309
63.	Polymerase Chain Reaction (PCR) Method for Identification Gene Escherichia coli and Officer Depot Behavior in Drinking Water Refill	315
64.	Assessment of Eu-152 Nuclide Contaminated from Radioactive lightning Rods in Soil Samples at Kasra and Atash in Baghdad	. 321
65.	Increased Expression of Interleukin 13 in Iraqi Patients Suffer from Ulcerative Colitis	326
66.	The Protective Role of Hydatid Cyst against Colorectal Cancers	332
67.	Pulp Response Capped by Brain Derived Neurotrophic Factor (BDNF)	337
68.	Quality of Food Bacteria in School Snacks and Canteens in East Jakarta Health Office Working Area in 2017	. 341
69.	Borax Content in Foods Sold in a Campus and Its Trader Characteristics	346
70.	The Condition of Sanitation Facilities with <i>Escherichia coli</i> Contamination on Food at University Cafeteria 2015	350
71.	Method and Frequency of Stethoscope Cleaning among Respiratory Therapists in Intensive Care Units at KAMC, Riyadh	354
72.	Hepatoprotectie Effect of Bromelain against Gentamicin-Induced Hepatic Damage in Rats	
73.	Acute Appendicitis Versus Ruptured Ovarian Cyst in Female Patients Presented as Acute Abdomen Pain Wisam Mahmood Aziz, Hayder Adnan Fawzi	364
74.	Effects of Health Promotion Behavior, Self-Esteem and Social Participation Activities on Life Satisfaction of Elderly Men A Reum Lee, Hee Kyung Kim	368
75.	Change of Brief Psychiatric Rating Scale (BPRS) Value with Spiritual Qur'anic Emotional Freedom Technique (SQEFT) Therapy on Mental Disorder Patient	

76.	5. An Empirical Analysis Research on the Characteristics of Elderly Welfare Organizations on Job Fit and Contextual Performance		
77.	A Study on the Effect of Job Performance on Emotional Labor, Career Turnover Intention, Job Stress, Growth Need	385	
78.	Application of Digital Rubbing Massage in Pain Level, Comfort, and Duration of Labor Phase	391	
79.	Using Propensity Score Bootstrapping on Determining the Model of the HIV/AIDS Patients' Assistance <i>Mahdalena, Mahpolah, Ismi Rajiani</i>	396	
80.	ARCS Module (Attention, Relevance, Confidence, Satisfaction) to Increase Classroom Motivation for Pregnant Women at Public Health Center	401	
81.	Interprofessional Education Module in Achieving Ethics/Values, Roles, Responsibilities, Professional Communication Competencies, and Team Collaboration among the College of Health Students	406	
82.	Anxiety Level of Dental Care among Adolescents in Kepulauan Selayar District	409	
83.	Baby Massage With Common Cold Massage Oil on Temperature Change, Pulse Rate, Frequency of Bro Sleep Quality and Number of Streptococcus Bacteria in Toddlers with Acute Respiratory Infection Melyana Nurul W, Fatatu Malikhah, Kusmini Suprihatin, Sutarmi		
84.	Risk Factors Affecting Attention Deficit Hyperactivity Disorder among Early Childhood in the Agricultural Area in Indonesia	417	
85.	Maternl and Neonatal Outcomes of Elective and Emergency Cesarean Sections	422	
86.	Comorbidities of Phototherapy Used in Neonatal Jaundice in Diyala Governorate, Iraq	428	
87.	Effect of Thyroid Disorder on Liver Function and Some Immunological Parameters	433	
88.	The Presence of Pathogenic Leptospira sp. in Water Bodies in Klaten District	439	
89.	Dialectic Unity between Threat and Division Sociological Study	444	
90.	Job Demands, Low Back Pain, and Job Crafting Behaviors: A Proposed Framework	449	
91.	"Educational-Staff Knowledge and Attitude towards Antibiotic Use in Technical Institute of Karbala". <i>Maytham Salim AL-Nasrawii, Ali abd Al–Latif. G. Mohammed, Mohammad Abdul Baqi Abdul Mohsin, Mohammed A.Merzah</i>		

92.	Intervention of Sexual Abuse Prevention for Mother of Children with Mental Retardation in Payakumbuh Indonesia 2016	61
93.	Prenatal Tobacco Exposure and Neonate Birth Weight	67
94.	Influence of Firm's Intangible Assets Intensity on Stock Prices Volatility: Evidence from Emerging Market of Pakistan	72
95.	Perception of Job Characteristics and Internal Motivation in Medical Records Staff	78
96.	Influencing Factors and Microbial Agents Which Contribute to Acne among Students from Pathological Analysis Department/Kufa Technical Institute\Al- Najaf Government	84
97.	Analysis of Factors on Reward System in the Hospital	90
98.	Model Development of Nursing Service Loyalty	95
99.	Occupational Health Issues Faced by Women in Spinners	00
100.	Association of HLA-DRB1 Alleles with Allergic Asthma and Total Serum IgE Levels in Iraqi Adults Patients	05
101.	The Role of Serum and Follicular IL-1Beta in Predicting the ICSI Outcome in Infertile Women 5 Rihab Abbas Ali, Sahib Yahya Hasan Al-Murshidi, Dalal Mahdi Al-jarah	511
102.	Factors Related to the Satisfaction of BPJS Participants on Outpatient Services in the Regional General Hospital Dr. H. Moch Ansari Saleh Banjarmasin	17
103.	Increasing of Nutrition Status of Pregnant Women after Supplementation of Moringa Leaf Extract (Moringa Oliefera) in the Coastal Area of Makassar, Indonesia	21
104.	Assessment of the Effect of Diyala River upon the Quality of Tigris River in Baghdad Province by National Sanitation Water-Quality Index (NFS-WQI)	26
105.	Determination of the Radiation of Alpha Particles in the Air of Primary School Buildings in the City of Karbala	31
106.	The Influence of Organizational Pride on the Performance of Lecturers in Health at the Nahdlatul Ulama University in Surabaya	38

107.	The Effectiveness of Using Direct Composite Veneer Template System in Restoring Anterior Teeth Sri Wahyuni, Saluna Deynilisa, Ismalayani	543
108.	A Study on Breast Cancer Awareness in Female Students of Begum Rokeya University, Rangpur: A Cross-Sectional Study	547
109.	Failure of Speed oligo Mycobacteria to diagnose <i>Mycobacterium tuberculosis</i> Complex Directly from Sputum Samples	553
110.	Body Fat Composition as a Determinan of Cognition Functions in Elementary School Students	557
111.	Factors Associated with the Knowledge and Attitude Towards Breastfeeding in Thai Grandmothers of Pregnant Adolescents	560
112.	Cross-Sector Collaboration Indicators Development of HIV-AIDS Prevention Program in Indonesia Balqis, Hasbullah Thabrany, Kemal N Siregar	566
113.	The Relationship between Ventilation with Excess Cancer Risk (ECR) of Benzene at the Shoe Home Industry in Romokalisari Surabaya Bachtiar Chahyadhi, Abdul Rohim Tualeka	572
114.	The Prevalence of Oral Manifestation in Transgenders with HIV/AIDS in Surabaya, East Java, Indonesia Sagus Soebadi, Adiastuti Endah Parmadiati, Hening Tuti Hendarti, Desiana Radithia, Diah Savitri Ernawati	577
115.	Evaluation of the Health Policy Implementation of Indonesian Social Insurance Administration Organization in Primary Health Care Facilities Supriyana, Edy Susanto, Irmawati, Bernadus Rudy Sunindya, Asep Tata Gunawan, Ismi Rajiani	581
116.	The Relationship between Environmental Sanitation to the Incidence of Hepatitis A in Rural Areas of Central Java, Indonesia	585
117.	The Correlation between Green Open Space with Carcinogen Toxicity Score of Benzene in Shoes Home Industry Surabaya	589
118.	Frequency of Cardiac Troponin T (TNNT2) Polymorphism, a Dilated Cardiomyopathy Gene in Tabuk Population	594
119.	SLC2A2 Gene (Glucose Transporter 2) Variation is Associated with an Increased Risk of Developing T2d in an Ethnic Population of Saudi Arabia	600
120.	A Content Analysis of Original Research Articles on Public Health Published in an International Journa The Case Study of Thailand Sunanta Wongchalee, Orapin Laosee, Ratana Somrongthong	

226.	Identification of Hazard and Risk Occupational Health in Lumu-Lumu Island Fisheries		
227.	. Analysis of Ordinal Logistic Regression Model on Breast Cancer Diagnosis by Birads Mammography 119 M. Nadjib Bustan, M. Arif Tiro, Suwardi Annas, Adiatma		
228.	28. Coping Mechanism of Students Facing the Competency Exams Reviewed from the Factors Influence i Surya Mitra Husada Health College Kediri		
229.	Correlation between Pulmonary Tuberculosis (TB) Patient's Characteristics and Role of Supervisor of Drugs Swallowing (PMO) with the Risk of Transmission in Medan City 2017		
230.	Determinants of Occupational Health and Safety Problems among Seaweed Workers in Takalar Regency 1214 Yahya Thamrin, Atjo Wahyu, Masyita Muis, Syamsiar S. Russeng, Agus Bintara Birawida, Hasnawati Amqam, Andi Hardianti		
231.	Risk Prediction Model of Lung Tuberculosis Using Spatial Approach in the Coastal Area of Makassar City1220 Stang Abdul Rahman, A. Ummu Salmah, Indra Dwinata, Anwar Mallongi		
232.	Analysis of Management System of Healthy Aisle Program in Makassar City		
233.	Prevention of Delay of Decision Making as Efforts to Improve The Mother Health in Polewali Mandar, West Sulawesi Indonesia		
234.	Criminal Liability of Illegal D Rug Traffickers in Makassar City		
235.	Climate Risks and Environmental Determinants on Dengue transmission		
236.	Efforts to Improve Knowledge of Dental and Oral of Sangging in Mepandes Ceremony in Kerambitan District Bali Province, Indonesia		
237.	Relationship of Rainfall, Population Density, and Human Behavior with DHF Incidence in Makassar City 1253 Hasanuddin Ishak, Jum Dewi Sartika, Darmawansyah		
238.	Ineffective Regulation of Narcotics Crime Prevention (Criminology Perspective)		
239.	Influence of Assertiveness Training in Preventing Sexual Harassment in Nigerian Universities as Perceived by Counsellors in Kwara State		
240.	The Development of Post Traumatic Stress Disorder among Secondary School Students in Borno State Nigeria: A Systematic Review		

Analysis of Ordinal Logistic Regression Model on Breast Cancer Diagnosis by Birads Mammography

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ABSTRACT

The right diagnosis is needed for appropriate therapy. The diagnosis of breast cancer is quite ambiguous and requires high accuracy. Mammography is a method of diagnosing breast cancer using BIRADS (Breast Imaging-Reporting and Data System) assessment. This study aimed to assess the accuracy of BIRADS classification in the diagnosis of breast cancer and predictors that influence it through a logistic regression model test. The research method was cross sectional study by collecting data from the results of mammography examinations obtained from Medical Record documents, SIRS (Hospital Information Systems), and the radiologist's expertise of mammography. The data came from 47 hospital breast cancer patients that contained information on potential predictors of breast cancer namely tumor location, metastases, age, weight, and education. Logistic regression model analysis was performed to find the best statistical test model for breast cancer diagnosis classification based on BIRADS assessment. The diagnosis classification of BIRADS was consisting of normal, benign, and malignant grades. For this reason, hypothesis testing was conducted with G test for simultaneous model testing. Then, a development of an appropriate logit model by using a partial test. Followed by conducting a suitability and feasibility test model with the Goodness of Fit using the Hosmer-Lemeshow Test. The results of the analysis revealed that the ordinal logistic regression was the best model of BIRADS classification diagnosis with an accuracy value of 52.5%. The result of ordinal logistic regression model for malignant breast cancer:

$$\hat{\pi}_1(x) = \frac{\exp(-19,436+1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}{1 + \exp(-19,436+1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}$$

The result for benign cancer:

$$\hat{\pi}_2(x) = \frac{\exp(-17,696+1,538\,\text{age}-5,725\,\text{education}-16,313\,\text{occupation}+2,549\,\text{location})}{1+\exp(-17,696+1,538\,\text{age}-5,725\,\text{education}-16,313\,\text{occupation}+2,549\,\text{location})} - \frac{\exp(-19,436+1,538\,\text{age}-5,725\,\text{education}-16,313\,\text{occupation}\,\text{Pekerjaan}+2,549\,\text{location})}{1+\exp(-19,436+1,538\,\text{age}-5,725\,\text{education}-16,313\,\text{occupation}+2,549\,\text{location})}$$

A significant predictor factors were the location of the tumor, age, education, and the work of cancer patients. The conclusion of the diagnosis classification of breast cancer using BIRADS of mammography is quite accurate and assessment of diagnosis classification BIRADS should pay attention to tumor location factors, age, education, and work of breast cancer patients.

Keywords: Ordinal logistic regression, BIRADS, mammography, breast cancer diagnosis

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INTRODUCTION

Every disease requires an accurate diagnosis so that doctors can provide appropriate treatment. The diagnosis of breast cancer is quite ambiguous but still requires a high accuracy of diagnosis.¹, ²

Diagnosis of breast cancer requires several types of testing, namely physical or clinical examination,

radiological examination, histopathological examination, genetic examination, and immunology.³

Radiological examinations for the diagnosis of breast cancer using mammography were assessed for the malignancy levels by using BIRADS (Breast Imaging-Reporting and Data System) developed by the American College of Radiology (ACR) and carried out by radiologists. BIRADS assessment is scoring from 1 to 6 with the meaning that 1 is negative, 2 is benign, 3 is probably benign, 4 is suspicious for malignancy, 5 is highly suggestive of malignancy, 6 = known biopsy malignancy.^{4,5}

For developing the model and assessing the accuracy of the mammography examination results, a statistical approach could applying the Ordinal Regression Logistics.⁶

The results of the model analysis will find the best model, the accuracy of the selected model, and determine the predictor factors that influence the presence of breast cancer.⁷

Ordinal Regression Logistics is one of the statistical methods for analyzing ordinal scale of response variables consisting of three or more categories. Predictor variables used in this model in the form of category data or quantitative data.⁸

Ordinal Regression Logistic Model for ordinal data response variables are often referred to as cumulative logistic models. The response variable in the cumulative logistic regression model is in the form of multilevel data represented by numbers 1, 2, 3, ..., k. With k is the number of categorical response variables. The cumulative logistic regression model will compare cumulative opportunities, ie opportunities less than or equal to the jth response category on p predictor variables expressed in vector of the x_i . $P(Y \le j | x_i)$, with opportunities greater than the response category j, x_i , $P(Y \ge j | x_i)$.

Cumulative opportunity forms are defined as follows:

$$\pi_{k}(x_{k}) = P(Y \le j | x_{i}) = \frac{\exp[g_{j}(x_{k})]}{1 + \exp[g_{j}(x_{k})]}$$

$$= \left(\frac{\exp(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k})}{1 + \exp(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k})}\right);$$

$$\text{dengan } k = 1, 2, ..., j, ..., r$$

$$\pi_k(x_k) = P(Y \le j|x_j) = p_1 + p_2 + \dots + \pi_k$$

The formula for general logistic distribution function

is:
$$F(x) = \frac{1}{1 + e^{-x}} = \frac{1}{1 + e^{x}}$$

If $P(Y \le j)$ is compared with the probability of a respons variabel on category (j+1) until category r, the result is:

$$\frac{P(Y \le j)}{P(Y > j)} = \frac{P(Y \le j)}{1 - P(Y \le j)} = \frac{\frac{\exp\left(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k}\right)}{1 + \exp\left(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k}\right)}}{1}$$

$$= \exp\left(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k}\right)$$

$$= \exp\left(\beta_{0j} + \sum_{k=1}^{r} \beta_{k} x_{k}\right)$$

$$\frac{P(Y \le j)}{P(Y > j)} = \frac{P(Y \le j)}{1 - P(Y \le j)} = \frac{\pi_{1} + \pi_{2} + \dots + \pi_{j}}{\pi_{j+1} + \pi_{j+2} + \dots + \pi_{r}}$$

Next, execute logistic transformation to be logit model of ordinal regression logistic:

Logit
$$[P(Y \le j)] = \log \frac{P(Y \le j)}{1 - P(Y \le j)}$$

$$= \log \frac{\pi_1 + \pi_2 + \dots + \pi_j}{\pi_{j+1} + \pi_{j+2} + \dots + \pi_r}$$
Logit $[P(Y \le j)] = \beta_{0j} + \sum_{k=1}^r \beta_k x_k$

with the value of β_k , for k = 1, 2,...,r to each of ordinal regression logistic is the same.

MATERIALS AND METHOD

To conduct a model analysis, data on breast cancer patients was needed. The source of data collection came from medical records documents, SIRS (Hospital Information System), and mammography images. Data containing information about patient identity and potential determinants in the term of age, tumor location, metastases, education, employment, and supplemented by the results of reading mammography expertise. The BIRADS assessment results are converted to ordinal data where 1 was normal, 2-3 were benign, 4-5-6 were malignant. The study design was a cross sectional study that collected data on breast cancer patients who were treated and registered at one Makassar hospital, Indonesia. The collected data was analyzed to find the best model. The analytical steps taken include: estimating parameters; - testing logit model parameters with simultaneous testing, partial test and logistic analysis; - and testing the suitability and accuracy of the model with the Goodness of Fit (GOF) test and the Hosmer-Lemeshow test.⁹

RESULTS

Parameter estimation was conducted by using simulataneus test of ordinal regression logistic by G test

with the formula G =
$$-2 \ln \frac{\left(\frac{n_1}{n}\right)^{n_1} \left(\frac{n_0}{n}\right)^{n_0}}{\prod_{i=1}^n \hat{\pi}_i \left(1 - \hat{\pi}_i\right)^{(1-y_i)}}$$

where
$$n_1 = \sum_{i=1}^n y_i$$
, $n_0 = \sum_{i=1}^n (1 - y_i)$, $n = n_1 + n_0$.

Rejection area Ho if $G > \chi^2_{(\nu,\alpha)}$ with ν degree of freedom is equal with the number of parameters in the model without β_o .

Simultaneous testing obtained a calculated value of -2log-likelihood model of 61,146. Because p-value is equal 0,001 is smaller than $\alpha = 0.01$, then Ho is rejected, which means that at least one predictive variable has a significant effect on the classification of breast cancer BIRADS. In this case the variables of age, education, occupation, and tumor location significantly influence the classification of breast cancer BIRADS.

Table 1: Statistical Output of G Test of Ordinal Regression Logistic

Model	-2 Log Likelihood	Chi- Square	df	P-Value
Intercept Only	89,529			
Final	61,146	28,382	9	0,001

To identify the role of each variable, parietal test is conducted with the result as the follows:

Table 2: Statistical Outputs of Partial Test of Ordinal Regression Logistic

	Estimation	Std. Error	Wald	P-Value
BIRADS Malignant	-19,436	4,222	21,192	0,000
BIRADS Benign	-17,696	4,230	17,505	0,000
Age	1,538	0,740	4,326	0,038
Education	-0,069	0,022	10,152	0,001
Occupation	-16,313	1,516	115,797	0,000
Location of tumor	2,549	1,086	5,510	0,019

From the results of the parameter estimation above, it is found that there are four predictor variables that have a significant effect on the variable level of malignancy BIRADS namely, age, tumor location, education and occupation.

The logit model of its statistical outputs are:

$$g_1$$
 (malignant) = $-19,436 + 1,538$ age $-5,725$ education $-16,313$ occupation $+2,549$ location

$$g_2$$
 (benign) = $-17,696 + 1,538$ age $-5,725$ education $-16,313$ occupation $+2,549$ location

Based on the three logit functions above, logit 1 is a logit function for malignant BIRADS and logit 2 is a logit function for benign BIRADS. Furthermore, from the two logit functions, the probability function of each category is obtained.

The logit model formula could be used to calculate the probability formulation for each response variable. The probability formula for malignant breast cancer BIRADS is as follows.

$$\hat{\pi}_1 \text{ (ganas)} = \frac{\exp(g_1(x))}{1 + \exp(g_1(x))}$$

Formulation of probability BIRADS benign breast cancer is:

$$\hat{\pi}_2(x) = \frac{\exp(g_2(x))}{1 + \exp(g_2(x))} - \frac{\exp(g_1(x))}{1 + \exp(g_1(x))}$$

So:

For Y=1 (BIRADS malignant)

$$\hat{\pi}_{1}(x) = \frac{\exp(-19,436+1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}{1 + \exp(-19,436+1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}$$
For Y=2 (BIRADS benign)

$$\hat{\pi}_{2}(x) = \frac{\exp(-17,696 + 1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}{1 + \exp(-17,696 + 1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})} - \frac{\exp(-19,436 + 1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}{1 + \exp(-19,436 + 1,538 \,\text{age} - 5,725 \,\text{education} - 16,313 \,\text{occupation} + 2,549 \,\text{location})}$$

To determine the model formed from the above predictor variables is appropriate or not in accordance with the data, the suitability model of Goodness of Fit is used by using Hosmer-Lemeshow test:

$$\hat{C}$$
 (Hosmer – Lemeshow) = $\sum_{k=1}^{g} \frac{\left(o_k - n_k \, | \, \overline{\pi}_k\right)^2}{n_k \, | \, \overline{\pi}_k \, \left(1 - \overline{\pi}_k\right)}$

Rejection area H_0 : $\hat{C} > \chi^2_{(\alpha,g-2)}$ or *p-value* $<\alpha = 0.01$

Table 3: Goodness of Fit Test of Ordinal Regression Logistic Model

	Chi-Square	Df	Sig.
Pearson	58,427	73	0,893
Deviance	56,987	73	0,916

The p-value results for Pearson and Deviance are more than α > 0.01, with values of 0.893 and 0.916, respectively. Ho is not rejected, which means that the model obtained is in accordance with the data or there is no significant difference between the results of the observation with the possible predictions of the model.

Thus, variables that significantly influence the increase in breast cancer BIRADS are variables of age, education, occupation, and location of the tumor.

To find out the model that is formed is feasible, it can be seen from the R2 value.

Table 4: Pseudo R-Square of Ordinal Regression Logistic Model

Cox and Snell	0,453
Nagelkerke	0,525
McFadden	0,303

Based on the table "Pseudo R-Square" the value of Nagelkerke R Square is 0.525. In other words, the resulting model with five predictor variables, the variables of age, education, occupation and location that have a significant effect while body weight variables did not significantly influence the increase risk in breast cancer BIRADS. In addition, the model was also able to explain the variation of breast cancer BIRADS classification.

In addition, the model was also able to explain the variation of breast cancer BIRADS classification amounting of 30.3%.

DISCUSSION

The accuracy test of diagnosis can be done using logistic regression models. There are three main types of logistic regression known, namely binary logistic regression, multi-nominal logistic regression and ordinal logistic regression.¹⁰

The selection of logistic regression types depends on the measurement scale of dependent variable data. Because the diagnosis of breast cancer BIRADS is categorical and ordinal (normal, benign, malignant), ordinal logistic regression is chosen. The results of the model analysis show that the Ordinal Regression Logistic model along with six predictors are only able to explain the variation of breast cancer BIRADS classification by 30.3

This happens because this model data still requires some important potential predictors such as marital status, age of menarche, menopausal status, and others.^{13,14}

CONCLUSION

The conclusion of the diagnosis of classification of breast cancer using BIRADS of Mammography is quite accurate, and assessment of classification diagnosis BIRADS should pay attention to tumor location factor, age, education, and work of breast cancer patients.

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