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ABSTRACT

End-stage renal disease (ESRD) is one of the public health problems in Indonesia. This study aimed to determine the clinical condition of premorbid and the incidence of ESRD. The study design is a case-control. The number of volunteers involved in this study was 100 patients. The cases were ESRD patients who were determined by the criteria of Pernefri and the controls were patients from the same hospital who were not ESRD by the requirements of ESRD Pernefri. Inclusion criteria of the study sample were ESRD sufferers need routine hemodialysis and doctor's diagnosis supported by laboratory data, 15-75 years old and willing to fill out inform consent. Exclusion criteria for the study sample were patients with congenital kidney disease, Have a history of kidney transplants, and mental disorder. We collected demographic data, and patient's clinical and drug history from medical records. Chi-square analysis with 2x2 tables and the Fisher's exact test were used to determine the relationship of clinical conditions and history of disease with the incidence of ESRD. The results showed that anemia, hyperglycemia, and hypertriglyceridemia were related to the prevalence of ESRD. History of hypertension, diabetes mellitus, dyslipidemia, and family history of ESRD are associated with an increased incidence of ESRD.

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1. INTRODUCTION

End-stage renal disease (ESRD) now and in the future is world health problems including in Indonesia [1-2]. Terminal chronic renal failure occurs when the glomerular filtration rate (LFG) is less than 15 ml/minute/1.73 m2 for three months or more or creatinine clearance<5 ml/minute and serum creatinine levels are more than or equal to 10 mg/mL [3-5]. In Indonesia, the incidence and prevalence of ESRD are at 100-150 and 200-250 per 1 million population per year. It was determined that there were more than 7,000 ESRD patients in Indonesia [6-8]. The mortality rate of patients with ESRD 5-10x is not ESRD. The mortality rate of ESRD patients in the first year of illness reached 20% and increased to 60% in the fifth year [9-12]. In 2030 ESRD is estimated to contribute 30% to the cause of death in the world [1, 11, 13]. Besides the increasing incidence, prevalence, and mortality rate of ESRD, what needs to be considered is the significant cost of care for ESRD patients [14-15]. ESRD does not only cause physical, economic and psychological damage to sufferers but also a burden on families, as well as the state so that efforts are preventive and promotive [7, 16]. In line with the results of other studies, previous reseracher [17] reported

that nearly 50% of ESRD patients who do hemodialysis at PKU Muhammadiyah Yogyakarta hospital have a low quality of life [14, 18].

Until now, treatment or treatment of patients with ESRD rests on two actions, namely kidney transplantation or undergoing dialysis to replace permanently damaged kidney function [19]. Both kidney transplantation and hemodialysis are actions that require high costs, and the results are not satisfactory. Due to the increasing number of ESRD sufferers in Indonesia, it is necessary to develop a preventive and promotive program in the field of nephrology. Identifying premorbid clinical risk factors and other biological factors as a basis for clinical decision making should be identified [6, 20-22]. However, not all ESRD risk factors in Indonesia have yet to be identified. Therefore this research needs to be done in Indonesia. The purpose of the study, in general, was to determine the relationship of premorbid clinical conditions with the incidence of ESRD.

2. RESEARCH METHOD

2.1. Type and design of the study

This study was an observational analytic study with a case-controlled design. ESRD patients as a case group, while non ESRD volunteers who are by or without smoking as a control. Data gathered at PKU Muhammadiyah Yogyakarta Hospital, Bantul public Hospital, Sleman public Hospital, and Panti Rapih Hospital in Special Region of Yogyakarta (SRY).

2.2. Research subject

The population was patients diagnosed with terminal chronic renal failure for no more than three months, with diagnostic criteria namely creatinine clearance<5ml/minute or blood serum creatinine level greater or equal to 10 mg/dl which can be known from medical records and requires continuous hemodialysis. Inclusion criteria for research subjects: Indonesians (Javanese, Sundanese, Malay); ages 15-65 years old; willing to participate in research by filling out and signing the agreement and cooperative statement sheets; and new ESRD patients namely establishing a diagnosis of ESRD in patients with or less than three months. Research subjects who have been selected through inclusion criteria will be excluded from the study subjects if: have congenital kidney disease, kidney transplant history, mental illness. The minimum number of samples needed for the case is 44 people. With a comparison of between cases and control is 1:2, the number of controls group is 88 patients. The minimum number of research subjects is 132 people. Sampling is done by consecutive random sampling.

2.3. Tools and materials.

The tools and materials in this study include equipment for primary data collection (questionnaires), secondary data collection forms, chemicals, and glassware.

2.4. Operational variables and definitions of research

Independent variables were gender, body weight (BW), age and smoking activity as well as blood cotinine levels. The dependent variable is ESRD. Terminal chronic kidney disease is a persistent (more than three months) kidney function disorder and requires a kidney transplant or routine dialysis to replace kidney function; kidney abnormalities are measured by a decrease in GFR<5 ml/minute or serum creatinine levels greater than or equal to 10 mg/dL. YES if the respondent meets laboratory criteria or requires a kidney transplant or undergoes hemodialysis. NO, if the respondent does not meet laboratory criteria or needs a kidney transplant or undergoing hemodialysis.

2.5. Collection data procedure

Measuring instruments used in this study include (i) Questionnaire, used to obtain primary data demographic characteristics and smoking habits of respondents. (ii) The data collection form is used to collect secondary data taken from medical record books or patient status which include name, age, gender, previous pain and medication diagnosis data, laboratory data and patient clinical manifestations.

2.6. Operational limitations

In general, data collection in this study was carried out by interviews, observation, and direct measurements. Primary data about ESRD pain history and smoking history were collected by structured interviews with guided questionnaires. Original data about the characteristics and clinical manifestations of the subject is obtained by direct measurement. Secondary data on demographic characteristics and pain history and treatment of research subjects were taken from medical records and supplemented by interviews with assisted forms of data collection and questionnaires.

The implementation phase includes activities in the field as well as in the laboratory as follows: (i) Primary data collection in the sample by interview using a questionnaire provides for the identity of the respondent, socio-economic, smoking behavior, disease data obtained from the patient's medical record and other related data with research variables. Subjects/volunteers who have fasted 8-12 hours measured blood pressure, waist circumference, body weight, and height for the determination of body mass index (BMI). Prospective subjects who met the inclusion criteria have explained the purpose of the study and were asked to fill out an informed consent sheet as proof of the participation of the subjects in the research and fill out the questionnaire. The case/control subjects were taken for 5 ml of blood from the anterior cubital median vein. 3 ml of blood was centrifuged at 3000 g for 15 minutes. The serum is used to examine creatinine, BUN, random blood glucose, triglyceride, cholesterol levels, and other examinations. Examination of creatinine and laboratory levels was carried out by the spectrophotometer method, as was done by previous researchers [23].

2.7. Data analysis

Chi-square analysis with 2x2 or 3x2 tables with Fisher's exact test was used to assess the relationship of clinical or biological conditions and illness history to the incidence of ESRD.

2.8. Research ethics

Because it involves human respondents, it explained to all prospective respondents about the purpose and objectives of the research, the benefits and expected uses and the consequences of the respondent (informed consent), in addition to the management of ethical clearance (No KEPK/847/EC).

3. RESULTS AND DISCUSSIONS

3.1. Overview of respondents

The research subjects consisted of 53 ESRD patients as the cases and 106 non ESRD patients as the controls, originating from 4 hospitals in Special Region of Yogyakarta (SRY), namely PKU Muhammadiyah Yogyakarta private Hospital, Sleman regional public Hospital, and Bantul regional public Hospital. The demographic conditions of research subjects according to age group, sex, type of work, and level of education as well as those responsible for medical expenses are presented in Table 1.

No	Respondents characteristics		Respondent status		Total	Р	
		Respondents characteristics	Cases (CKD)	Control	Total	Ľ,	
1		PKU Yogyakarta hospital	19 (33.3%)	38 (66.7%)	57 (100%)		
	Hospital	PKU Gamping hospital	6 (33.3%)	12 (66.7%)	18 (100%)	1	
1		Bantul regional hospital	25 (33.3%)	50 (66.7%)	75 (100%)	1	
		Sleman regional hospital	3 (33.3%)	6 (66.7%)	9 (100%)		
2	Sex	Male	35 (33.3%)	70 (66.7%)	105 (100%)	0.568	
2		Female	18 (33.3%)	36 (66.7%)	54 (100%)	0.308	
		16-30 year	5 (31.2%)	11 (68.8%)	16 (100%)		
3	Age group	31-45 year	15 (33.3%)	30 (66.7%)	45 (100%)	0.998	
3		46-60 year	25 (33.8%)	49 (66.2%)	74 (100%)	0.998	
		>60 year	8 (33.3%)	16 (66.7%)	24 (100%)		
		No	3 (21.4%)	11 (78.6%)	14 (100%)		
		elementary	14 (28.0%)	36 (72.0%)	50 (100%)		
4	Education	Yunior high school	5 (21.7%)	18 (78.3%)	23 (100%)	0.267	
		Senior high school	21 (44.7%)	26 (55.3%)	47 (100%)		
		University	10 (33.3%)	15 (66.7%)	9 (100%)		
	Job	Military, police or Government employees	13 (52.0%)	12 (48%)	25 (100%)		
5		Private and self-employed	9 (36.0%)	16 (64.0%)	25 (100%)	0.085	
5		Farmers, laborers, and artisans	15 (23.8%)	48 (76.2%)	63 (100%)	0.085	
		No job	16 (34.8%)	30 (65.2%)	46 (100%)		
	Assurance	No	1 (5.3%)	18 (94.7%)	19 (100%)		
6		Private health Assurance	19 (54.3%)	16 (45.7%)	35 (100%)	0.001	
		Public health assurance	33 (31.4%)	72 (68.6%)	105 (100%)		
7	Marital	Yes	52 (33.5%)	103 (66.5%)	155 (100%)	0.502	
/	status	No	1 (25.0%)	3 (75.0%)	4 (100%)	0.593	

Table 1. The demographic characteristics of ESRD patients in the Special Region of Yogyakarta.

Most of the research subjects were between 46-60 years old (50.3%), followed by the general group 31-45 years (27%). >60 years (18.2%) and 16-30 years (7.4%). The difference in the distribution of research subjects based on age groups between ESRD patients and not ESRD was not different (p>0.05).

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3.2. Overview of respondents' clinical conditions

An overview of clinical and laboratory conditions is presented in Table 2. The table shows that the average Hb, erythrocytes, leukocytes, and hematocrit levels of the ESRD group were lower than the control group (p<0.05). Blood glucose, urea, and creatinine levels in the ESRD group were higher than the control group (p<0.05). Anemia is one of the distinguishing factors in handling ESRD patients [24].

No	Respondents characteristic	Cases	Control	р		
1	Age	48.34±11.15	48.18±11.32	0.962		
2	Body weight	57.23 ± 8.06	57.62±9.01	0.787		
3	BMI	22.21±2.79	22.33±3.27	0.809		
4	Hb level	8.24±1.46	12.93±1.81	0.000*		
5	Eritrocytes	2.89 ± 0.56	4.58±0.58	0.000*		
6	Leucocytes	7.08 ± 2.06	9.24±3.21	0.000*		
7	Thrombocytes	242.26±65.23	249.33±68.17	0.533		
8	Haematocrit	23.54±2.90	38.73±5.62	0.000*		
9	Blood glucose ad random (RBG)	120.87±23.22	107.72±13.87	0.000*		
10	hypertriglyseridemia	118.46 ± 56.18	115.63±38.18	0.742		
11	Ureum level	155.76 ± 42.81	21.65 ± 7.82	0.000*		
12	Creatinine level	8.83 ± 2.97	0.75±0.24	0.000*		
13	Glomerular filtration rate (GFR)	5.81±2.97	104.89 ± 20.05	0.000*		
Noto:*-p<0.05						

Note:*=p<0.05

3.3. Bivariate analysis of clinical and biological conditions with ESRD events

The relationship of the biological and clinical characteristics of respondents with the incidence of ESRD is presented in Table 3. This table reveals that the incidence of ESRD is related to clinical conditions. Anemia is related to the incidence of ESRD Anemic conditions (Hb<12 mg%) have a 37x ESRD risk compared to not anemic. RBG status>120 mg/ml increases the risk of ESRD 2.79x (p<0.05). The condition of hypertriglyceridemia (TG level>200 mg/dl) increases the risk of ESRD 5.42x. The body mass index is not related to the incidence of ESRD. The results of the study are in line with other studies [23, 25].

Table 3. Relationship between the clinical characteristic (BMI, haemoglobin, RBG, and triglycerides) with ESPD events

with ESKD events							
Clinical biology characteristic		Cases	controlled	Total	OR; CI;95%	Р	
	Lean	6(31.6%)	13(68.4%)	19(100%)	1	0.856	
BMI classification	Normal	37(35.6%)	67(64.4%)	104(100%)	0.84 (.29-2.38)	0.737	
Divit classification	Fat	6(28.6%)	15(71.4%)	21(100%)	1.15 (.29-4.47)	0.836	
	Obesity	4(26.7%)	11(73.3%)	15(100%)	1.27 (.28-5.68)	0.755	
Hb level	<12	51(54.3%)	43(45.7%)	94(100%)	37 (8.63-161.68)	0.000	
no level	>=12	2(3.1%)	63(96.9%)	65(100%)			
RBG	>120	21(51.2%)	20(48.8%)	41(100%)	2.79 (1.337-5.817)	0.005	
KDU	<=120	32(27.4%)	85 (72.6%)	117(100%)	2.79 (1.557-5.617)		
Urmontniloonidonnio	>200	5(71.4%)	2(28.6%)	7(100%)	5.42 (1.014-8.922)	0.042	
Hypertrilgeridemia	<=200	48(31.6%)	104(68.4%)	152(100%)	3.42 (1.014-8.922)		

3.4. Bivariate analysis relationship between respondent's history and ESRD events

Pre-morbid conditions that increase the risk of ESRD events are presented in Table 4. The results of the study showed that some history of the disease was related to the incidence of ESRD. History of DM, hypertension, hypercholesterolemia has been shown to increase the incidence of ESRD. The family history of the ESRD is also related to the prevalence of ESRD. Opportunities for ESRD events in respondents with a history of DM 34x than without a history of DM. Opportunities for ESRD events in respondents with a history of hypertension 8x than respondents did not have a history of hypertension. Family history of ESRD pain is 8.6 times more than families without a history of ESRD. The results of the study are in accordance with the results of previous studies [1, 26], except obesity. Diabetes mellitus (DM), hypertension (HTN), obesity and cardiovascular disease have attributed to the rapid rise in the incidence of CKD in most parts of the world, especially in developing countries [1].

Table 4. Relationship of previous illness history with ESRD events in respondents

Illness history		Cases	controlled	Total	OR, CI 95%	р
DM	Yes	21 (91.3%)	2(8.7%)	23 (100%)	34 (7.58–153.48)	0.000
DW	no	32 (23.5%	104 (76.5%)	136(100%)		
Hypertension	Yes	37(62.7%)	22(37.3%)	59 (100%)	8 (4.17-18.71)	0.000
Hypertension	no	16(16.0%)	84(84.0%)	100 (100%)		
Hipercholesterolemia	Ya	18(58%)	13(42%)	31(100)	3 (1.6-8.29)	0.001
nipercholesterolenna	Tidak	35(27%)	93(73%)	128(100%)		
ESRD in family	Yes	4(80%)	1(20%)	5 (100%)	8 (0.93-78.71)	0.043
ESKD III faililly	No	49(31.8%)	105(68.2%)	154 (100%)		

4. CONCLUSION

The clinical conditions of hyperglycemia, hypertriglyceridemia, and anemia are associated with the incidence of ESRD. History of diabetes, hypercholesterolemia, hypertension, and family history with ESRD are also related to the prevalence of ESRD.

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