

Tue, May 17, 2022 at 9:53 PM

Fwd: PLEASE REVISE_5723

5 messages

adinda rahma <adinda.rahma.triyaniarta-2019@fkm.unair.ac.id> To: Santi Martini <santi-m@fkm.unair.ac.id>

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Terima kasih...Dinda.

Apakah bisa dikirimkan juga email konfirmasi ketika submit?

Wass.,

Santi Martini Faculty of Public Health Universitas Airlangga (www.fkm.unair.ac.id) Surabaya INDONESIA

Preventing disease, Prolonging Life, and Promoting Health through the Organized Efforts of Society *****

[Quoted text hidden]

Tue, May 17, 2022 at 10:57 PM

To: adinda rahma <adinda.rahma.triyaniarta-2019@fkm.unair.ac.id>

Adinda,

Berikut manuskrip yang disubmitkan ya. Silakan cek revisi saya. Tks

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OK....tqvm.....Dinda.

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1	Determinants of Type-2 Diabetes Mellitus in Passive Smokers	
2		
3	Adinda Rahma Triyaniarta ¹ , Santi Martini ^{2*} , Kurnia Dwi Artanti ² , Sri Widati ² , Rizma Dwi	
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14		
15	Abstract	
16	Type-2 Diabetes Mellitus (T2DM) leads to complications to other diseases. The modifiable risk	
17	factors for T2DM are overweight, physical activity, hypertension, unhealthy diet and smoking.	
18	This study aims to analyse the determinants of T2DM incidence in passive smoker among some_	Commented [j1]: aimed
19	factors. This research was conducted at X Hospital in Surabaya and was conducted in September	
20	- April 2020. These variables were univariate, bivariate, and multivariate. Case samples were	
21	T2DM sufferers and passive smokers, while control samples were non-T2DM sufferers and	
22	passive smokers, 52 respondents per group, total respondents were 104 Variables that were	
23	statistically significant related to the incidence of T2DM in passive smokers were age >45 years,	

24 level of education not attending primary school, less physical activity, and hypertension. The 25 variable that had no statistically significant related with the incidence of T2DM in passive 26 smokers were gender, occupation, sedentary lifestyles, income, and genetic. The multivariate 27 analysis showed that age is a major factor contributing to T2DM incidence in passive smokers at 28 X Hospital in Surabaya in 2019. It can be concluded that age is the most dominant risk factor of 29 T2DM incidence in passive smokers

30

31

Keywords: Passive smoker, determinants, type-2 diabetes mellitus

Commented [j2]: Urutkan sesuai abjad.

32 Introduction

33 Type-2 diabetes mellitus (T2DM) has been counted for about 90% of all cases of diabetes. In 34 Type-2 DM, there is decreased ability of insulin to stimulate glucose uptake by peripheral tissues 35 and to inhibit glucose production by the liver, and this is defined as insulin resistance.¹ The cause 36 of insulin resistance is often due to obesity and lack of physical activity and aging. Another event 37 that lead to T2DM is pancreatic cell dysfunction, cells will show a disturbance in the first phase 38 of insulin secretion, where insulin secretion fails to compensate for insulin resistance. If not 39 treated properly, there will be damage to pancreatic cells. This will occur progressively and will 40 cause insulin deficiency, so that eventually patients require exogenous insulin. If pancreatic cells 41 cannot produce insulin secretion immediately and quickly to compensate for insulin resistance, 42 fasting hyper glycemia and diabetes will appear.² Diabetes Mellitus can cause complications to 43 other diseases, such as blindness, heart attack, stroke, kidney failure and leg amputation.³ 44 According to the World Health Organization, arterial hypertension and T2DM are the two most common cardiovascular risk factors in the global population. Despite the single roles as 45 independent cardiovascular risk factors, hypertension and T2DM often coexist in the same 46 47 patient. This coexistence multiplies the patient's risk of experiencing major acute cardiovascular

48 events and accelerates the development of chronic heart and kidney failure.⁴ Diabetes Mellitus 49 and the complications can be managed and prevented, especially when detected early. It's even 50 better to take precautions by making lifestyle changes, such as improving diet and physical 51 exercise.

52 In recent years, gender, which is male, has been regarded as a risk factor for the 53 development of type 2 diabetes. The reason that men are more prone than women to the development of this disease is not known yet. The development of T2DM is the result of the 54 55 interaction between environmental factors and a strong genetic component. Environmental risk 56 factors known as influence of the development of T2DM include obesity, sedentary lifestyle, 57 birth weight, and stress.⁵ In addition, diabetes is also a disease with a high socioeconomic 58 pattern, favoring the beneficiary group, especially in developed countries. According to a 59 systematic review on socioeconomic positions and incidence of diabetes, low education is most consistently associated with an increased risk of diabetes when compared to other socioeconomic 60 61 indicators, the knowledge and skills acquired through education determine the responsiveness to health information.6 62

63 Physical activity is one of the risk factors of Diabetes Mellitus. Physical activity is body 64 movement produced by skeletal muscles that releases energy. Physical activity consists of 65 strenuous activity, moderate activity and light activity. Lack of physical activity is estimated to be the main cause of around 21-25% of breast and colon cancer, 27% of diabetes, and about 30% 66 of the global burden of ischemic heart disease.⁷ In addition, smoking is also a risk factor for 67 diabetes. This is in line with a cohort study conducted by Kowall et.al, which found that being an 68 active smoker and a passive smoker increases the likelihood of developing type-2 diabetes 69 70 mellitus.8

71 Meanwhile, 28.11% of the population aged >10 years old in East Java are classified as smokers with the intensity of smoking every day.9 With the increasing number of smokers, the 72 73 problem of passive smoking is also increasing. 85% of households in Indonesia are exposed to 74 cigarette smoke, with an estimated eight smokers dying from active smoking, one passive 75 smoker dying from exposure to other people's cigarette smoke.¹⁰ In passive smokers, exposure to 76 cigarette smoke can increase the risk of type-2 diabetes mellitus by several mechanisms that are 77 almost the same as those that occur in active smokers. This mechanism involves 5000 chemicals, 78 including 50 types of carcinogenic and toxic materials. Endothelial function disorders due to 79 smoking cause changes in blood circulation which result in decreased blood flow to skeletal 80 muscles which can lead to insulin resistance.8

81 Cigarette smoke can increase blood sugar levels. The effect of nicotine stimulates the adrenal 82 glands and can increase glucose levels.¹¹ Nicotine can also inhibit insulin secretion. The 83 mechanism of nicotine in inhibiting insulin secretion is when nicotine attaches to the nicotinic 84 acetylcholine receptor (nAChR) on pancreatic cells. Nicotine attached to these receptors causes 85 an increase in oxidative stress (ROS) in cells and which leads to changes in the function and structure of cell mitochondria.¹² These changes will interfere with the ATP formation process in 86 secreting insulin, resulting in pancreatic cell apoptosis.¹³ Pancreatic cell apoptosis is the death of 87 88 cells in the pancreas so that the function of insulin secretion in the pancreas decreases and insulin decreases in the body.14 89

Diabetes Mellitus is a serious problem in Indonesia and the world. Based on Riskesdas 2018, East Java is one of the provinces with the fifth highest incidence of diabetes mellitus in Indonesia, which is 2.6%. This number has increased when compared to the results of Riskesdas 2013 which was 2.1%. Surabaya as the capital of East Java province is the city with the highest **Commented [j3]:** Tulis dalam Bahasa Inggris. Mohon ubah seluruh penggunaan kata seperti ini.

number of diabetes cases, which is 3.4% in the 2018 Riskesdas results. From year to year the 94 95 number of diabetics in Surabaya continues to increase. Likewise with the prevalence of physical activity. Based on the results of Riskesdas 2018, in East Java there were 61.5% of the population 96 97 with sufficient physical activity and 33.5% with less physical activity. This number illustrates an increase when compared to the percentage of physical activity in Riskesdas 2013, which is 98 99 26.1% of the population doing less physical activity. The increasing number of smokers in 100 Indonesia results in tendency to increase the risk of developing T2DM. This study aims to 101 analyse the determinants of T2DM incidence in passive smoker among some factors, which are 102 age, sex, education, occupation, income, heredity, sedentary lifestyle, physical activity, and 103 hypertension.

104 Method

105 This research uses analytical observational research design with a case control approach. This 106 research was conducted in outpatient polyclinic and inpatient polyclinic at X Hospital in 107 Surabaya. The population of this study consisted of a case population of type-2 diabetes mellitus 108 and all patients who are passive smokers at X Hospital in Surabaya. While the control population 109 in this study were all non-patients with Type-2 DM and other smoking-related diseases at X 110 Hospital in Surabaya who are passive smokers. The sample of this study were all patients who 111 are classified as passive smokers at X Hospital in Surabaya. In this study, the sample was divided 112 into 2, namely: 1.) Case samples are patients with Type-2 Diabetes Mellitus and passive 113 smokers. 2.) The control sample is patients with no Type-2 DM and other smoking-related 114 diseases who are passive smokers.

115 The sampling technique used is simple random sampling. The inclusion criteria for this study 116 were patients diagnosed with type-2 diabetes mellitus or non-smokers with smoking-related 117 diseases at X Hospital in Surabaya who participated in the study. In addition, the second 118 inclusion criteria are that the respondent is a passive smoker who has a family member who 119 smokes or co-workers smoke at work every day. Based on research data, the total population is 120 obtained 400 people consisting of 300 population cases of smoking-related diseases and 100 121 people population control. The amount of the sample according to the research criteria, namely 122 patients with Type 2 DM in passive smokers as many as 159 people as the case population, while non-patients with Type 2 DM in passive smokers as a control population as many as 52 people. 123 124 This amount is then adjusted by doing simple random sampling with lottery method to 159 case 125 population in order to obtain 52 respondents in accordance with the number of control samples. 126 The final sample for each group consists of 52 respondents and the total sample in this study is as 127 many as 104 respondents.

128 Data retrieval through secondary data. The data collected was then inputted and cleaned, 129 so that they could be analyzed. These variables were univariate, bivariate, and multivariate. 130 Univariate analysis was done for each variable with the proportional results of each. Bivariate 131 analysis with chi-square with the results whether there was a relationship between each 132 independent variable and the dependent variable if the p value was less than 0.05 was considered 133 to statistical significance. Multiple logistic regressions with backward determinant modeling 134 shows multivariate result to find significant determinant factors. The research was conducted at 135 X Hospital in Surabaya from September to April 2020. This study will use secondary data based 136 on interviews with people with Type-2 Diabetes Mellitus which have been carried out in an 137 research group by lecturers of the Faculty of Public Health, namely Dr. Santi Martini, dr., 138 M.Kes. in 2019 entitled "The Burden of Diseases Due to Cigarettes in East Java". The lack of physical activity measurement questionnaire consists of questions taken from the Riskesdas 2018 139

which is a modification of the WHO Global Physical Activity Questionnaire (GPAC) which is
part of the WHO STEPS instrument for measuring and monitoring risk factors for noncommunicable diseases.

143 Results

From January to June 2019, the number of Diabetes Mellitus patients at X Hospital in Surabaya was 8060 people, while Type-2 Diabetes Mellitus (T2DM) patients were 7875 people. The average visit for one month is 1313 people. This number represents the number of DM patients in general. The study was conducted on Type-2 Diabetes Mellitus patients who made outpatient visits at the Internal Medicine polyclinic and inpatients at X Hospital in Surabaya.

149 Based on table 1, most of the respondents are in the age range above 45 years as many as 150 58 people (44.2%). The gender distribution of respondents shows that the majority (80.8%) are 151 female. From the table 1 is known that the most respondents are with high school education 152 level, which is 33 people (31.7%) and the least is with junior high school education level, which 153 is 17 people (16.3%). These data indicate that most of the respondents are classified as having a 154 low level of education. It was also found that most of the respondents are passive smokers who 155 are going to work. While the income of the respondents is balanced between below the Surabaya 156 minimum wage and above the Surabaya minimum wage. Most of the respondents are passive 157 smokers who do sufficient physical activity. Most of the respondents are passive smokers who 158 do not excessive sedentary behavior (69,20%). Most of them have no history of diabetes in the 159 family (60,58%) and also a history of hypertension (71,15%).

160Table 1. Characteristics of the Sample (Passive Smokers Patients) at the X Hospital in161Surabaya in 2019

Characteristics	n = (Total Sample)	%
Age (Year)		
18 -45	46	44.23

	50	
\geq 46	58	55.77
Gender		
Male	20	19.23
Female	84	80.77
Education		
Not in School/ Elementary	31	29.81
School		
Junior High School	17	16.35
High School	33	31.73
College	23	22.11
Occupation		
Unemployed	48	46.15
Employed	56	53.85
Physical Activity		
Not Enough	19	18.30
Enough	85	81.7
Sedentary Lifestyle		
≥ 6 hours	32	30.80
< 6 hours	72	69.20
Income		
> Minimum Wage	34	32.70
< Minimum Wage	70	67.30
Genetic		
Yes	41	39.42
No	63	60.58
Hypertension		
Yes	30	28,85
No	74	71,15

Table 2. Results of Bivariate Analysis of Type-2 Diabetes Mellitus Incidence in Passive Smokers

162 163 164

	Γ	Disease	Incide	ence X	p- value	OR value (95%CI)
Variable		Yes	No			
	n	%	n	%	_	
Age (Year)						
18 - 45	4	7.7	43	82.7	<0.001	57.3
≥ 46	48	92.3	9	17.3	< 0.001	(16.4-199.6)
Gender						
Male	14	26.9	6	11.5	0.002	2,825
Female	38	73.1	46	88.5	0.082	(0.99 - 8.059)
Education						
Not in School/ Elementary School	26	50	5	9.6	< 0.001	8,09 (2.268 -28.853)
Junior High School	5	9.6	12	23.1	>0.05	0.65 (0.170-2.470)

High School	12	23.1	21	40.4		0.89 (0.297-2.663)
College	9	17.3	14	26.9		1
Occupation						
Unemployed	26	50	22	42.3	0.555	1.364
Employed	26	50	30	57.7	0.555	(0.629-2.955)
Physical Activity						
Not Enough	14	26.9	5	9.6	0.042	3.463
Enough	38	73.1	47	90.4	0.042	(1.1145-10.477)
Sedentary Lifestyles						
\geq 6 hours	15	28.8	17	32.7	0.922	0.835
< 6 hours	37	71.2	35	67.3	0.832	(0.362-1.922)
Income						
> Minimum Wage	35	67.30	35	67.30	1 000	1.000
< Minimum Wage	17	32.70	17	32.70	1.000	(0.441-2.269)
Genetic						
Yes	23	44.23	18	34.52	0.422	0.668
No	29	55.77	34	65.38	0.422	(0.303 - 1.472)
Hypertension						
Yes	30	57.69	0	00.00	-0.001	0,423
No	22	42,31	52	100,00	<0,001	(0,308-0,581)

165 Notes: OR: Odd Ratio; CI: Confidence Interval; *Variables that were significantly related, p-value < 0.05

Based on table 2, it shows the distribution of study participants according to the presence of 166 167 incidence of T2DM in passive smokers. Bivariate analysis showed that variable age (p value = 168 <0,001, OR= 57,3), education not in school/elementary school graduated (p value = <0,001, OR= 8,09), physical activity (p value = <0,042, OR= 3,463), and hypertension (p value = 169 170 <0,001, OR= 0,423) were significantly correlated with the incidence of T2DM in passive 171 smokers. It shows too that variable gender, junior high school graduated, senior high school 172 graduated, college graduated, occupation, sedentary lifestyles, income, and genetic were not significantly correlated with the incidence of T2DM in passive smokers (p value >0,05). 173

Table 3. Multivariate Analysis of Variable Associated with Type-2 Diabetes Mellitus 174 175 **Incidence in Passive Smokers**

	Variables	β	р	OR	95% Confidence
					Interval
Step 1 ^a	Age	3.992	.001	54.143	4.866 - 602.418
	Gender	-3.035	.013	.048	.004521

	Physical Activity	.841	.507	2.318	.193 - 27.793
	Education	.098	.954	1.103	.039 - 30.904
	Hypertension	22.747	.997	7566889697.486	.000
	Constant	-27.384	.996	.000	
Step 2 ^a	Age	4.010	.000	55.122	12.218 - 248.673
	Gender	-1.519	.077	.219	.041 - 1.178
	Physical Activity	1.482	.123	4.404	.668 - 29.029
	Education	.680	.506	1.974	.266 - 14.631
	Constant	-7.800	.001	.000	
Step 3 ^a	Age	4.301	.000	73.768	17.635 - 308.569
	Gender	-1.317	.137	.268	.047 - 1.518
	Physical Activity	1.465	.108	4.327	.724 - 25.882
Step 4 ^a	Age	4.189	.000	65.987	16.975 - 256.509
	Physical activity	1.694	.058	5.442	.943 - 31.389
	Constant	-9.014	.000	.000	
Step 5 ^b	Age	3.937	.000	51.250	9.360 - 280.613
	Physical activity	1.506	.145	4.509	.595 - 34.199
	Hypertension	21.503	.997	2180215011.610	.000
	Constant	-29.018	.997	.000	-

Notes :

a. Variable(s) entered on step 1: Age, Physical activity, Education, hypertension, gender.

b. Variable(s) entered on step 5: hypertension.

176

177 Table 4. Multivariate Final Model of Variable Associated with Type-2 Diabetes Mellitus

178 Incidence in Passive Smokers

Variables	β	р	OR	95% Confidence		
				Interval		
Age	3.937	.000	51.250	9.360 - 280.613	 	Commented [j4]: Umur yang mana? Umur 18<45 atau
Gender	-1.317	.137	.268	.047 - 1.518		lebih dari 45?
Physical Activity	1.506	.145	4.509	.595 - 34.199	 ×.,	
						Commented [15]. Laki laki atau perempuan

¹⁷⁹

Based on Table 4, variable age, gender, and physical activity were significant with the incidence of type 2 diabetes mellitus with p value < 0.25. Multivariate analysis was applied to determine the dominant factors of T2DM incidence in passive smokers. Backward Wald method is carried out on all independent variables that meet the requirements included in the model. Variables that are not significant are excluded gradually, starting from the variable with the Commented [j5]: Laki-laki atau perempuan

Commented [j6]: Ini not enough atau enough?

highest p value. In Table 4, the age variable (p-value= 0.000) is a factor that is significantly related to T2DM incidence in passive smokers after being controlled with age, gender, education, hypertension, and physical activity. The final stage of logistic regression results obtained variable age as the most important risk factor for T2DM incidence in passive smoker (OR= 51,520, 95% confidence interval [CI]= 9,360-280,613).

191 192

193 Discussion

194 The majority of respondents with Type-2 Diabetes Mellitus and passive smokers are >45 195 years old (54.8%) with p value = <0.001 which means that there is a significant relationship 196 between age in passive smoking and the incidence of Type-2 Diabetes Mellitus. classified as 197 elderly (elderly). Along with increasing age, the risk of diabetes mellitus and heart disease 198 increases.¹⁵ This is in accordance with the results of a report from the IDF in 2017 which stated 199 that as long as the increasing age, the prevalence of Diabetes Mellitus also get higher.¹⁶ This is 200 because Diabetes Mellitus often appears after a person enters a vulnerable age, especially after 201 the age of 45 years and those who are overweight, so that the body is no longer sensitive to 202 insulin. The aging process results in changes in the body's anatomical, physiological and 203 biochemical systems that can cause insulin resistance. This condition will get worse if it is accompanied by complications of other diseases, especially in the elderly group.¹⁷ As we age, the 204 205 body's metabolism slows down naturally, which causes decreased physical activity. Low 206 mobility will speed up the replacement of muscle mass with body fat. This condition can lead to 207 obesity, which is one of the risk factors for Type-2 Diabetes Mellitus.

The majority of respondents with Type-2 Diabetes Mellitus had no education/graduated from elementary school (50%). The results of data processing also showed that the level of education that did not go to school/graduated from elementary school was related to the incidence of Type211 2 Diabetes Mellitus ($p = \langle 0.001 \rangle$). Meanwhile, respondents with junior high and senior high 212 education levels had a p value of > 0.05 so that there was no significant relationship between 213 junior high school, senior high school, and Universities education levels than those with no 214 education level/graduated from elementary school. This is in accordance with previous research 215 by Irawan (2010) which stated that people with a low level of education were 1.27 times at risk 216 of suffering from DM than people with higher education.¹⁸ Although elementary and junior high 217 school education levels are included in the type of basic education, both have differences in 218 terms of level of competence and knowledge taught. The level of education is believed to be an 219 important factor for someone to be able to understand the management of blood sugar control, 220 overcome symptoms that arise with appropriate treatment and prevent complications generally 221 related to knowledge. Patients with high education are believed to have better knowledge about 222 diabetes and its effects on health compared to low education, so sufferers will respond in a 223 positive way and will try to recover.19

224 This study also revealed that physical activity is associated with the incidence of Type-2 DM 225 in passive smokers (p value = 0.042). This shows that there is a significant relationship between 226 physical activity of passive smoking and the incidence of Type-2 DM. In the physical activity 227 variable, the OR value is 3.4. This shows that people with less physical activity have a 3.4 times 228 greater risk of Type-2 DM than people with sufficient physical activity. From these data, it was 229 found that most of the respondents were passive smokers who did sufficient physical activity. 230 These results are the same as the research by Sipayung, Siregar and Nurmaini which showed 231 82.8% of the 120 respondents did sufficient physical activity.

Absorption of glucose by body tissues at rest requires insulin, whereas in active muscles, although the need for glucose increases, it is not accompanied by an increase in insulin levels. 234 This is because when a person is physically active, there is an increase in insulin receptor sensitivity in active.²⁰ The condition of insulin resistance results in glucose not being able to 235 236 enter the cells. When a person does physical activity, there will be muscle contractions. This can 237 make it easier for glucose to enter cells, so that when a person is physically active, there will be a decrease in insulin resistance and will ultimately reduce blood sugar levels.²⁰ Based on the 238 239 research of Siti Cholishotul Himmah et al (2020) at the Aulia Jombang Clinic, it was found that physical activity was an influential variable with Type-2 Diabetes Mellitus.²¹ Patients who had 240 241 high physical activity experienced the greatest decrease in sugar levels, namely a decrease of 242 53.6 mg/dl. Decrease in blood sugar levels in patients who have moderate physical activity by 243 6.73mg/dl. While the decrease in blood sugar levels in patients who have low physical activity is 4.3 mg/dl.21 244

245 From this study we also know that no hypertension has the significant (p value = <0,001, 246 OR= 0,423) relationship with the T2DM in passive smokers. It can maybe occur, because most 247 of the case group has the hypertension history (57,69%). These results are the same as the study 248 conducted by Nainggolan, Olwin, et al (2011) which stated that hypertension was significantly 249 associated with the incidence of T2DM (p = 0.00). Respondents with no hypertension have 250 protective value 0,423 times compared to those with hypertension for diabetes. Hypertension is a 251 factor that causes DM disease.²² Hypertension and DM are health problems that are closely related and both need to be handled carefully. high blood pressure causes the distribution of 252 253 sugar blood in the cells do not run optimally, so that occur accumulation of sugar and cholesterol 254 in the blood. The point is if blood pressure is good, sugar blood will also be awake. Insulin acts as a substance controlling blood pressure and water levels in the body, so that insulin levels are 255 256 enough to maintain blood pressure (Alfiyah, 2011).

257 Meanwhile, the gender variable did not have a significant relationship with the incidence of 258 Type-2 DM in passive smokers. However, the table shows that the majority of respondents are 259 female passive smokers, as many as 84 people (80.8%) of respondents. The actual results in the 260 field show that there are more women affected by Type-2 Diabetes Mellitus than men. In 261 addition, the p value is almost close to significant (p value = 0.082) indicating that there is a 262 possible relationship between the gender of passive smokers and the incidence of Type-2 263 Diabetes Mellitus. This result is also in accordance with several previous studies, namely 264 Nordstrom (2016) which stated that the prevalence of Type-2 Diabetes Mellitus in males was higher larger than women.²³ The IDF report in 2017 stated that the prevalence of Type-2 265 266 Diabetes Mellitus in the world in the male sex was greater than the female gender.¹⁵

The results of data processing also shows that some variables are not associated with the incidence of Diabetes Mellitus Type 2. Like the occupation of passive smokers (p = 0.431), The sedentary behavior excessive (≥ 6 hours) in passive smokers with the incidence of Type 2 Diabetes Mellitus (p = 0.832). The income of passive smokers (p = 1,000), and the genetic of passive smoker (p = 0,422).

272 The results showed that age is a major factor contributing to T2DM incidence in passive 273 smokers at X Hospital in Surabaya in 2019. It was found that respondents who were over 45 274 years of age and a passive smoker had an 51,52 times higher risk of developing type 2 diabetes 275 mellitus compared to those who were under 45 years of age and were passive smokers (OR= 276 51,520, 95% CI= 9,360-280,613). American Diabetes Association (ADA) (2011) stated that risk 277 Type 2 diabetes mellitus increases with age. Mechanism underlying higher risk type 2 diabetes 278 mellitus in individuals who being older is increase in body fat composition which accumulates in 279 the abdomen, so that lead to central obesity. Central obesity further triggers occurrence of insulin resistance is the initial process of diabetes mellitus type 2 (Suastika et al, 2012). WHO also stated that after a person reaches the age of 40 years, blood glucose levels rise 1-2 mg% per year during fasting and will rise approximately 5.6 - 13 mg% at 2 hours after eating. From that, it's not surprising if the age factor is a major factor in the occurrence of increasing prevalence of diabetes mellitus especially type II and impaired tolerance glucose.²⁴

Based on previous research conducted by Wei, et al (2014), it can be conluded that passive smoking is also one of the risks of T2DM.²⁵ The research by Pan, et al (2015) stated that both active and passive smoking are associated with significantly increased risks of type 2 diabetes. The risk of diabetes is increased in new quitters, but decreases substantially as the time since quitting increases.²⁶ Therefore, interventions to prevent exposure to secondhand smoke remain an urgent priority.²⁷

However, in conducting this research, there are still some limitations. The limitations of the study include the absence of data regarding the length of exposure to passive smokers who have co-workers who smoke, so that it cannot be seen that there is an influence of the variable length of exposure with the age of the respondent when it is associated with the incidence of Type-2 DM.

296	Conclusion	 Commented [j7]: Make it concise/shorter.
297	From the results of the above discussion, it can be concluded that age is the most dominant	
298	risk factor of T2DM incidence in passive smokers. People with age >45 years old are having 57	
299	times risk of developing Type-2 DM than the age of 18-45 years. The aging process results in	
300	changes in the body's anatomical, physiological and biochemical systems that can cause insulin	

resistance. This condition will get worse if it is accompanied by complications of other diseases,especially in the elderly group.

303	The level of education in passive smokers relate to the incidence of Type-2 Diabetes
304	Mellitus. The level of education of passive smoker respondents that does not go to
305	school/graduated from elementary school has a risk of developing Type-2 DM 8.09 times than
306	the level of college education. Physical activity in passive smokers has relationship with the
307	incidence of Type-2 Diabetes Mellitus. Less physical activity in passive smokers rises the risk of
308	developing Type-2 Diabetes Mellitus 3.463 times than sufficient physical activity. Respondents
309	with no hypertension have protective value 0,423 times compared to those with hypertension for
310	diabetes. Meanwhile, there is no relationship between the gender, occupation, and sedentary
311	lifestyles of passive smoking and the incidence of Type-2 DM at X Hospital in Surabaya in
312	2019.
313	According to the results, the following suggestions are given by researchers for the Surabaya
314	City government to reduce the incidence of Type-2 DM. First, the government is expected to
315	improve the health promotion and education programs about the effects of smoking, especially
316	related to passive smokers who are affected by exposure of cigarette smoke and age above >45
317	or among the elderly. Second, increasing and activating the efforts for the establishments of Non-
318	Smoking Areas in various public places and facilities. Third, always aggressively and actively
319	socializing CERDIK behavior (regular health checks, getting rid of cigarette smoke, diligent
320	activity, balanced diet, getting enough rest, managing stress) to the wider community to increase
321	awareness of Diabetes Mellitus. And finally, providing sports facilities or jogging tracks in every
322	area such as parks or green open areas so that people, especially passive smokers, can be
323	motivated to do physical activity.

324 Abbreviations

- 325 ATP : Adenosine triphosphate
- 326 DM : Diabetes Mellitus

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- 327 GPAC : Global Physical Activity Questionnaire
- 328 nAChR : Nicotinic Acetylcholine Receptor
- 329 ROS : Oxidative Stress
- 330 T2DM : Type-2 Diabetes Mellitus
- 331 WHO : World Health Organization
- 332
- 333 Ethics Approval and Consent to Participate
- 334 This research was approved by the ethics committee of RSUD dr. Soetomo Surabaya with letter
- number 0727/KEPK/X/2018 and has received research permission from X Hospital in Surabaya,
- all subjects in the study also agreed to be respondents.
- 337 Competing Interest
- 338 We declare no competing interests.
- 339 Availability of Data and Materials
- 340 All data set generated and analysis are available in the article.
- 341 Authors' Contribution
- 342 SM designed the study and wrote the protocol. All authors did the study. SM supervised all the
- 343 steps in the review process. All authors interpreted the findings. ART and RDN drafted the
- 344 manuscript. SM supervised the writing. KDA, SW, RDN, provided feedback.

345 Acknowledgment

- 346 We thank to all the organization that involve in this research
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