ORIGINAL ARTICLE 1

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Primary infertility of male and female factors, polycystic ovary syndrome and 3 4 oligoasthenoteratozoospermia dominate the infertile population in agricultural and 5 industrial areas in Karawang Regency, West Java Province, Indonesia 6 Assangga Guyansyah¹, Raditya Wratsangka¹, Denny Dhanardono¹, Muhammad Farid 7 Ghazali², Hosea Jaya Edy³, Haryo Ganeca Widyatama⁴, Dietha Kusumaningrum⁵, 8 9 David Tjahyadi⁶, Edy Parwanto^{7*} 10 11 ¹Department of Obstetrics and Gynecology, Faculty of Medicine, Universitas Trisakti, Indonesia. 12 ²Policlinic of Infertil, Mitra Bunda Amanda Karawang Clinic, Karawang Regency, West 13 Java, Indonesia 14 15 ³Study Program of Pharmacy, Faculty of Math and Natural Sciences, Universitas Sam Ratulangi, Indonesia 16 17 ⁴Medical doctor of Bhakti Mandala Clinic, Tangerang Regency, Banten Province, Indonesia. 18 19 ⁵Medical doctor of Batari Husada Clinic, Duren Sawit, East Jakarta, Special Capital 20 Region of Jakarta, Indonesia 21 ⁶Department of Histology, Faculty of Medicine, Universitas Trisakti, Indonesia. 22 ⁷Department of Biology, Faculty of Medicine, Universitas Trisakti, Indonesia. 23 24 25 * Corresponding to: Edy Parwanto, Department of Biology, Faculty of Medicine, Universitas Trisakti 26 Kyai Tapa Street, Kampus B, No.260 Grogol 11440, Jakarta-Indonesia. 27 28 Email: edyparwanto@trisakti.ac.id; edy.parwanto@gmail.com

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30	
31	ABSTRACT
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33	Introduction: Indonesia is a country with a large agricultural and industry, known to
34	utilize various types of pesticides, as well as several other industries with uncontrolled
35	pollution levels distributed across the nation. Besides, numerous studies have stated the
36	adverse effects of chemicals substances used in daily life and industrial waste on the health
37	of living things, including humans. This study aimed to determine the infertility
38	characteristic in the agricultural and industrial areas in Karawang Regency, West Java
39	Province, Indonesia.
40	Methods: The study was conducted retrospectively on medical records. Therefore, this
41	study determined the infertility characteristics based on sperm analysis, the etiology of the
42	causes of infertility in female, and the diagnosis of infertility. Data collection was obtained
43	from patients' medical records in the Infertile Poly of Mitra Bunda Amanda Hospital
44	Karawang, Karawang Regency, West Java Province, Indonesia.
45	Result: The results showed infertility was most prevalent in males aged 30-40 years
46	(55.79%) and females below 30 years (61.05%). Furthermore, the male and female's most
47	prevalent educational qualification (33.68% and 36.84%, respectively) was discovered to
48	be high school diploma. In terms of occupation, most male (56.84%) were laborers, while
49	the female was mostly housewives (36.84%). Meanwhile, oligoasthenoteratozoospermia
50	was the most analyzed sperm type (33.68%), and polycystic ovary syndrome was the most
51	common etiology of infertility in females (26.32%). The most prevalent diagnosis was
52	primary infertility factors, male and female (45.26%).
53	Conclusion: Primary infertility of male and female factors, polycystic ovary syndrome and
54	oligoasthenoteratozoospermia dominate the infertile population in agricultural and
55	industrial areas in Karawang Regency, West Java Province, Indonesia.
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57	Keywords: pollutants, sperm analysis, infertility, oligoasthenoteratozoospermia,
58	polycystic ovary syndrome.

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61 **INTRODUCTION**

Pollutants are harmful to human health and damage the environment. Pollutants in the human environment have various effects that are detrimental to health to cause disease.¹ The character of disease risk due to exposure to multi-pollutants can be determined using an "environmental risk score".² Environmental pollutants with estrogenic effects generally have biological effects in women, as pollutants with anti-androgenic effects that affect male fertility.³

Pollutants from agricultural and industrial activities can pollute the air, water and soil. 69 70 Of course, pollutants can move places. Pollutants in the air can move to water and land or vice versa. Directly or indirectly, pollutants can cause human health problems. Air 71 72 pollution can occur because it contains particles with aerodynamic diameters below 10 and 73 2.5 µm (PM10 and PM2.5). Apart from that, it can also contain NO, NO2, NOx and SO2. 74 It has been proven that air pollutants cause endocrine disorders and hormonal disturbances. 75 Women exposed to high concentrations of air pollutants, namely PM 2.5, NO, NO2, NOx and SO2, have a high risk of developing polycystic ovary syndrome.⁴ Besides that, it has 76 been shown that particles less than 0.3 µm in diameter can dominate the acute effects of 77 particulate air pollution resulting in cardiac autonomic dysfunction.⁵ 78

Furthermore, it was reported that PM interferes with energy metabolism, thereby 79 disrupting the endocrine glands and becoming a risk for cardiovascular disease.⁶ In this 80 regard, we have reported a case of aortic enlargement on the cadaveric heart and great 81 vessel dimensions.⁷ Because that, studies are still needed on the effect of pollutants on 82 83 aortic enlargement. That is important because cardiovascular disease as a risk factor has been shown to have a strong correlation with a history of infertility in women of 84 childbearing age and menopause.⁸ Moreover, chemicals used in the textile industry are 85 believed to produce persistent organic pollutants (POPs). These chemicals include 86 87 dichlorodiphenyltrichloroethane (DDT), dichlorodiphenyldichloroethylene (p,p'-DDE) and polychlorinated biphenyls (PCBs). POP is a stable lipophilic compound found in the 88 89 environment. These pollutants are difficult to break down, insoluble in water, and accumulate in the human body. Furthermore, pollutants in the body can cause human 90 health problems.⁹ How are people living in agricultural and industrial areas exposed to 91 pollutants?. 92

93 Of course, people living in the area are exposed to various pollutants from agricultural 94 and industrial activities. One proof that agricultural and industrial activities in Karawang Regency, West Java Province, Indonesia, impact the environment can be seen in the water 95 quality of the Citarum river. The Citarum River is a large and long river in West Java that 96 crosses Karawang Regency. It has been reported that the water in the Citarum river is of 97 poor quality, making it unsuitable for drinking. This fact illustrates that the water in the 98 Citarum river contains high pollutants.¹⁰ It should be noted that 18 sub-districts in 99 100 Karawang Regency are crossed by the downstream segment of the Citarum River. 101 Furthermore, it is shown that the high pollutant load of COD, BOD, phosphate and nitrate 102 in the downstream section of the Citarum river. The high pollutant load found in the 103 Citarum river downstream is caused by excess waste from domestic, agricultural and industrial activities.¹¹ 104

105 Epidemiological studies show that pollutants affect animal and human life. It has been shown that air pollution plays a role in infertility. Factory waste is a disruptive endocrine 106 hormone able to damage the body's endocrine system through various mechanisms.¹² 107 Moreover that pollutants disrupt spermatogenesis, leading to decreased reproductive 108 capacity in exposed populations.¹³ The results of previous studies show the effect of 109 environmental lead pollution on blood lead and sex hormone levels in the electronic waste 110 disposal area.¹⁴ Previous studies have also shown that pollution affects chromosomes, 111 thereby affecting infertility and sex hormone levels.¹⁵ Infertility is still a problem for many 112 113 married couples. It was also stated that the average age of childbearing in women was increasing.¹⁶ Moreover that infertility in Indonesia occurs in about 10-15% of couples of 114 childbearing age.¹⁷ 115

Based on the researchers' data, it is necessary to conduct a study on the characteristics of infertile communities living in agricultural and industrial areas. The purpose of this study was to determine the characteristics of infertility in agricultural and industrial areas. The infertility characteristics include the results of sperm analysis, the etiology of the causes of infertility in females, and the diagnosis of infertility in the agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia.

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125 METHODS

This research is a retrospective study with descriptive analysis. This research is part of a research project on infertility characteristics in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia, in 2015-2020.

The research material is secondary data obtained from the medical records of patients. The study was conducted from June to November 2019 in the Infertile Poly of RSIA Mitra Bunda Amanda Karawang, Karawang Regency, West Java Province, Indonesia. Collection of medical record data used from January 1st to December 31st, 2015. The individuals whose data were used in this study all live in Karawang Regency, West Java, Indonesia.

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136 **RESULTS**

Age	N (190 Subjects)	%	
Male	95		
<30 years	35	36.84%	
30–40 years	53	55.79%	
>40 years	7	7.37%	
Female	95		
<30 years	58	61.05%	
30-40 years	35	36.84%	
>40 years	2	2.11%	

137 **Table 1.** Age distribution of Research Subjects

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Based on the Table 1, showed infertility was most prevalent in males between 30 and 40 years (55.79%), followed by the age group below 30 years (36.84%) and above 40 years (7.37%). Meanwhile, in the female, infertility was most prevalent in the age group below 30 years (61.05%), followed by female aged 30-40 years (36.84%), and above 40 years (2.11%).

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145	Table 2.	Educational	qualification	of Research	Subjects
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Age	N (190 Subjects)	%	
Male	95		
Junior High School	20	21.05%	
High School	32	33.68%	
Academy	25	26.32%	
Bachelor	18	18.95%	
Female	95		
Junior High School	30	31.58%	
High School	35	36.84%	
Academy	22	23.16%	
Bachelor	8	8.42%	

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According to Table 2, the most common educational qualification possessed by male education is high school diploma (33.68%), followed by the academy (26.32%), junior high school (21.05%), and bachelor (18.95%) degrees. Similarly, the most prevalent educational qualification possessed by the female was high school diploma (36.84%), followed by junior high school (31.58%), academy (23.16%), and bachelor (8.42%) degrees.

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154 **Table 3.** Characteristics of Research Subjects Based on Occupation

Age	N (190 Subjects)	%	
Male	95		
Laborer	21	22.11%	
Factory Employees	54	56.84%	
Entrepreneur	28	29.47%	
Civil servants	12	12.63%	
Female	95		
Laborer	12	12.63%	

Factory Employees	22	23.16%
Entrepreneur	18	18.95%
Civil servants	12	12.63%
Housewife	35	36.84%

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Based on Table 3, infertility was discovered to be most prevalent in factory employees (56.84%), followed by entrepreneurs (29.47%), laborers (22.11%), and civil servant males (12.63%) in terms of occupation. Meanwhile, in the female, infertility cases were most prevalent in housewives (36.84%), followed by factory employees (23.16%), laborers (12.63%), and civil servants (12.63%).

161 **Table 4.** Infertility diagnosis based on the research Subjects

Diagnosis	(N=95 couples)	%
Primary Infertility ex Male Factor	30	31.58%
Primary Infertility of Male and Female Factors	43	45.26%
Primary Infertility ex Female factor	18	18.95%
Secondary Infertility	4	4.21%

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Based on Table 6 shows the most common infertility diagnosis was primary infertility male and female factor (45.26%), followed by primary infertility ex male factor (31.58%), primary infertility ex female factor (18.95%), and secondary infertility (4.21%).

167 **Table 5.** The Etiology of Infertility in the Female Research Subjects

Etiology	N (95 Subjects)	%	
Tubal Factor	19	20.0%	
PCOS	25	26.32%	
Myoma	16	16.84%	
Endometriosis	20	21.05%	
Ovulation	15	15.79%	

168 Abbreviations: PCOS=polycystic ovary syndrome

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According to Table 5, the etiology of infertility in females was discovered to be majorly due to polycystic ovary syndrome (PCOS), 26.32%, followed by endometriosis 21.05%, tubal factors 20.0%, myoma 16.84%, and ovulation 15.79%.

Based on Table 6 shows that oligoasthenoteratozoospermia being the most prevalent sperm type (33.68%), followed by oligoteratozoospermia (23.16%), oligoasthenospermia (21.05%), teratospermia (7.37%), oligospermia (6. 32%), normospermia (4.23%), and asthenospermia (4.23%).

Age	N (95 Subjects)	%	
Normospermia	4	4.21%	
Oligospermia	6	6.32%	
Asthenospremia	4	4.21%	
Teratospermia	7	7.37%	
Oligoasthenospremia	20	21.05%	
Oligoteratozoospremia	22	23.16%	
Oligoasthenoteratozoospermia	32	33.68%	

177 Table 6. Sperm Analysis of Research Subjects

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180 **DISCUSSION**

This study showed that infertility is most prevalent in males between 30 and 40 years 181 (55.79) and females below 30 years (61.05%). Ordinarily, these age groups ought to be 182 183 rather reproductive. However, on the contrary, these are the groups with the most infertility problems. Meanwhile, the highest educational qualification possessed by the male and 184 185 female is High School diploma (33.68% and 36.84%, respectively). That is a possible early indication of the factories' harmful impact in work environments and around residences. It 186 187 has been reported that demographic factors such as gender, education, income and geographic location influence the prevalence of infertility in infertile Chinese men and 188 189 women.¹⁸ Besides, the general levels of education, knowledge, and socioeconomic development within the region are currently low. Consequently, many people are ignorant 190 or forced to live near factories and to utilize polluted water sources. In terms of occupation, 191 the males were mostly laborers (56.84%), while the female was mostly housewives 192

(36.84%). The occupation of laborers is a possible cause of infertility, especially in exposure to heat and direct contact with heat sources, often encountered in the manufacture of the metal rim, tires, steel plates, zinc, machine operators, motorcycle body frames, forklifts, and other products. This exposure of male reproductive organs to heat is possibly associated with reduction in sperm quality. That can occur because high temperatures cause an increase in testicular metabolism so that sperm is damaged.¹⁹

199 Pollution has detrimental effects on health, not only by direct inhalation of pollutants 200 but also through other means of exposure, including ingesting contaminated water or skin 201 contact. One easy example is carbon monoxide as a pollutant from industrial activities. In 202 humans, carbon monoxide poisoning affects the cardiovascular, neurological, and affective systems.²⁰ The most common health effects are respiratory infections. However, pollutants 203 204 affect all body systems, including reproduction. The exact pathophysiology of the pollutant 205 effect on ovaries is not currently known. However, pollutants bind to hemoglobin during blood circulation and cause toxicity upon entering body organs.²¹ We already know that 206 agricultural and industrial activities produce pollutants as a by-product. Therefore the 207 208 negative effects of pollutants on the population must be avoided. Also, the government has long-established technical guidelines for industrial estates (Pedoman Teknis Kawasan 209 Industri).²² 210

Based on the diagnosis of infertility, this study showed that the main factor of male 211 212 and female infertility has the biggest role compared with the other factors (Table 4). We 213 already know that various hormones play a role in the reproductive process, including gonadotrophin-releasing hormone (GnRH), follicle-stimulating hormone (FSH), luteinizing 214 215 hormone (LH), estrogen, progesterone, testosterone, and inhibin. It has been proven that 216 estrogen plays a role in the reproductive system of women and men. Apart from that, 217 estrogen also plays a role in the neuroendocrine, skeletal, vascular and immune systems. Therefore, estrogen has implications for infertility and other diseases.²³ Therefore, 218 219 exogenous estrogenic compounds have the potential to interfere with the reproductive system. In this regard, the effects of diethylstilbestrol (DES) and methoxychlor (MXC) 220 221 have been investigated on female rhesus monkeys' peripubertal period. These studies' results indicate that DES had a striking effect on adolescent maturation, and MXC also 222 altered development during this period. The pattern of effects across agents and doses may 223 be based on specifics of estrogenic action.²⁴ On the other hand, it has also been proven that 224

225 xenoestrogen is involved in the decrease in the number and quality of human sperm,
226 consequently contributing to a decrease in fertility and decline in the proportion of male
227 births. Xenoestrogens have also been shown to increase the occurrence of abnormalities in
228 the male reproductive tract. Moreover, it has also been shown that xenoestrogens play a
229 role in increasing spontaneous abortion.²⁵

230 It has been stated that primary infertility is associated with protein that binds with sex 231 hormones. In humans, some proteins bind with sex hormones in the circulatory system and 232 the testes. The protein that binds with sex hormones in the circulating system is called sex 233 hormone-binding globulin (SHBG). Proteins that bind to sex hormones in the testes are 234 called androgen binding proteins (ABP). SHBG in the circulatory system has a function to bind sex steroid hormones and mediate the work of these hormones to target cells outside 235 236 the testes, while ABP functions to mediate the action of sex steroid hormones in the testes.²⁶ It is shown that the distribution of SHBG concentrations is broad-based on age and 237 body mass index (BMI) values in primary infertile men. From these two variables, it turns 238 out that the relationship between BMI and a decrease in SHBG levels is stronger than the 239 relationship between age and increased levels of SHBG.²⁷ The other study showed that the 240 levels of SHBG, total testosterone, free testosterone and percent of free testosterone have a 241 242 negative correlation with age, but the insulin and free testosterone index do not correlate 243 with age. The decrease in SHBG levels per decade in healthy Indonesian men was 8.19%, while the decrease of total testosterone levels per decade in healthy Indonesian men was 244 9.8%.²⁸ The results of previous studies show that low total testosterone levels can increase 245 fasting blood glucose levels in adult men, but SHBG levels do not predict fasting blood 246 glucose levels.²⁹ Although it has been stated that SHBG levels are influenced by many 247 factors, including genetic factors such as the genetic polymorphism of SHBG.²⁹ 248

249 Research has been carried out concerning primary infertility to reduce SHBG levels in postmenopausal women, namely by isoflavone supplementation.³¹ We recommend that 250 this method be implemented in women of childbearing age to increase fertility. Also, 251 women of childbearing age in industrial areas also need special attention to BMI, 252 253 especially those less than 18.5 kg/m². We recommend that women of childbearing age in these areas have a normal BMI. We need to present this matter because our results show 254 that women of reproductive age with a BMI <18.5 kg/m² and having a heterozygous 255 variant SHBG genotype (W/v) is undernutrition. Moreover, it has also been shown that 256

women of childbearing age with a BMI <18.5 kg/m² and having the heterozygous variant SHBG genotype (W/v) have lower protein, fat and carbohydrate intake.³² It has been stated that gene mutations cause abnormalities in protein metabolism in cells. Disorders of protein metabolism in cells cause various forms of organ abnormalities, resulting in congenital abnormalities³⁰ and morphological variations.³¹ Therefore, it is necessary to improve nutrition for reproductive women in agricultural and industrial areas such as in Karawang Regency, West Java Province, Indonesia.

Various natural ingredients can be used as a source of protein. Proteins that are sourced from natural materials can be developed to meet protein intake. Moreover, it has also been shown that proteins from natural ingredients contain several enzymes with the potential for therapy.³⁵ All the above studies' results that reveal the role of SHBG in both men's and women's reproductive systems clarify the relationship between SHBG and primary infertility. Apart from hormones and SHBG, which can affect primary infertility, it is necessary to discuss pollutants that affect populations in agricultural and industrial areas.

271 Based on the etiology of infertility in female subjects, this study indicates that PCOS ranks top, which is 26.32% of the total subjects. PCOS is potentially valuable indicators of 272 cultural, environmental, and genetic factors that may contribute to excess risk in certain 273 world regions. It has been proven that the prevalence of PCOS is determined by region and 274 275 race/ethnicity.³² The results of a study in the US showed that the prevalence of PCOS in the southern region was 47.5%, in the central region at 23.0%, while in the western region 276 it was 18.7% and in the northeast region 10.3%.³³ Also, it has also been stated that genetic 277 and environmental (lifestyle) factors are associated with the pathophysiology of PCOS 278 after prenatal exposure to androgens.³⁴ Moreover, environmental toxins, dietary diet, 279 obesity, and geographical variations are associated with PCOS.³⁵ Besides these pollutants, 280 281 bisphenol A {2, 2,-bis (4-hydroxyphenyl) propane=BPA)} is made by combining acetone 282 and phenol. BPA is used in food packaging and in general as an industrial ingredient. BPA 283 exposure to humans can be through inhalation, skin and digestive tract. BPA has weak estrogenic, anti-androgenic, and antithyroid activity, although it can accumulate in various 284 285 human body tissues. It has been reported that BPA affects metabolism and the reproductive system in humans. It is more detailed than BPA decreases male and female fertility.³⁶ In 286 more detail, it shows the impact of 2,2-bis 4-hydroxyphenyl propane (BPA) as a water and 287 soil pollutant with PCOS incidence.³⁷ The results of previous studies showed that the 288

women with PCOS had higher blood levels of BPA than the control group.³⁸ With the high percentage of primary infertility in this study, research on various pollutants in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia, should be conducted.

Oligoasthenoteratozoospermia in this study reached 33.68% of the population (N=95 293 294 subjects). The results of this study are different from study results in India. A study in India of 105 295 showed that 3.8% men with fertility problems experienced oligoasthenoteratozoospermia.39 296 We suspect that high prevalence the of oligoasthenoteratozoospermia in the group of infertile men in this study is related to 297 298 environmental pollutants. It has been explained previously that high pollutant loads are found in the downstream part of the Citarum river, which crosses the Karawang Regency. 299 300 Our statement follows the research results, which state a significant positive correlation between seminal total PCB level and the percentage of single-stranded DNA in sperm.⁹ 301

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303 CONCLUSION

304 Primary infertility of male and female factors, polycystic ovary syndrome and 305 oligoasthenoteratozoospermia dominate the population in agricultural and industrial areas 306 in Karawang Regency, West Java Province, Indonesia. Therefore, it requires supervision 307 and protection from the government, society, factory owners, and related health workers. 308 This study is intended to overcome the impact of pollutants that threaten residents who live 309 and work in agricultural and industrial areas in Karawang district, West Java Province, 310 Indonesia. Of course, this is also applied in the other agricultural and industrial areas in 311 Indonesia.

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313 ACKNOWLEDGEMENTS

We express our deep gratitude to the Leaders and Staff of Mitra Bunda Amanda Hospital,
Karawang, Karawang Regency, West Java Province, Indonesia, for facilitating data
collection and data validation.

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318 CONFLICTS OF INTEREST

319 The authors declare that they have no competing interests.

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321 FUNDING

322 All authors have no support or funding to report.

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324 AUTHORS CONTRIBUTIONS

- 325 Conceptualization: AG, RW, DD. Data acquisition: AG, DD, HGW and DK. Data analysis
- 326 or interpretation: AG, RW, HJE, HGW, DK and DT. Drafting of the manuscript: AG, RW
- and EP. Critical revision of the manuscript: DD, HJE, HGW and DT. Approval of the final
- 328 version of the manuscript: all authors.
- 329

330 ETHICS APPROVAL AND CONSENT TO PARTICIPATE

- 331 Not applicable.
- 332

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473		



Accepted with Revision (BaliMedJ) Primary infertility of husband and wife factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia

34 pesan

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: ML EDY Parwanto <edy.parwanto@gmail.com> 30 Januari 2021 pukul 14.20

Dear Authors,

Thank you for submitting your article entitled: "Primary infertility of husband and wife factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia"

Based on our author guidelines, Your article fulfilled the minimal required structure,

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In order to have a better-structured article, we suggest you edit based on a checklist, the simplest way, you may use the Publons review checklist.

According to the new International regulation, please input your detail in the article:

1. Ethical clearance number. (Send a Copy) If your work is a review, please fill out the ICJME for that can be found on author guidelines.

2. Please state your conflict of interest in the paper. (Confirmed)

3. Please state the funding (if any) in your paper. (Confirmed)

4. Please state each author's contribution. (Confirmed)

I also need to comment on your English academic writing, make sure to take your time, and avoid little mistyping, since we found more than **227 critical grammar errors**.

If you want us to fix this for you, we will charge an extra **200 USD** for proofreading and editing (by native English Medical Speaker).

Thank you for trusting us with your hard work

Best regards Bali Medical Journal (BaliMedJ) P-ISSN: 2089-1180 E-ISSN 2302-2914 Indexed at: Web of Science (WOS) Clarivate Analytics SCOPUS Elsevier All Indexing Organisation







MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com>

Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Thank you for your information. [Kutipan teks disembunyikan] MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 31 Januari 2021 pukul 08.29 Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Dear Editor, Here we will send the ICMJE form. Thank you for the good cooperation. Best regards, Parwanto MLE [Kutipan teks disembunyikan] ICMJE-AG.pdf 1225K MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 31 Januari 2021 pukul 08.43 Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Dear Editor, Here we will send the ICMJE form. Thank you for the good cooperation. Best regards, Parwanto MLE [Kutipan teks disembunyikan] ICMJE-AG-1.pdf 443K Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 1 Februari 2021 pukul 16.06 Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com>

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Thank you for the ICJME form, after having a discussion based on our initial review, and adding some comment from our associate editor,

we decided to accept your article with some revisions required. To continue to the peer review process, please confirm that you have paid the article processing charge.

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[Kutipan teks disembunyikan]

30 Januari 2021 pukul 16.05

Yes, I agree [Kutipan teks disembunyikan]

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com>

4 Februari 2021 pukul 02.17

8 Februari 2021 pukul 15.29

9 Februari 2021 pukul 10.17

11 Februari 2021 pukul 13.18

11 Februari 2021 pukul 13.45

11 Februari 2021 pukul 13.46

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Reminder [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Dear editor, Thank you for reminding us. We're taking care administration at our institution. Thank you, Best regards, [Kutipan teks disembunyikan]

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com>

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Thank you [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Thank you! [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Assangga Guyansyah <assanggaguyansyah@trisakti.ac.id> Cc: Assangga Guyansyah <assangga ag@yahoo.com>

[Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

The following updated invoices are not for us: https://www.paypal.com/invoice/p/#GCPBWDMUD87K3PPS. Please verify. Thank you, Greetings, Parwanto MLE [Kutipan teks disembunyikan] Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 19 Februari 2021 pukul 19.00 Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> **Dear Author** We are terribly sorry for the technical issue, The correct billing is still the old link: https://www.paypal.com/invoice/p/#L4XN3K287L8MSQDQ Thank you for trusting us with your hard work Best regards [Kutipan teks disembunyikan] MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 19 Februari 2021 pukul 19.28 Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> No problem, thank you! [Kutipan teks disembunyikan] Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 23 Februari 2021 pukul 10.02 Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> **Dear Author** We have received your payment, please keep this pdf as legal payment documentation. Your manuscript is now currently being processed by our reviewer and editor. Please patiently wait until we send you the revised version of your manuscript. Do inform us if you need the letter of acceptance prior to the publication. Thank you for trusting us with your hard works. Best regards [Kutipan teks disembunyikan] Invoice - 1096.pdf 59K MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 23 Februari 2021 pukul 10.23 Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Dear Editor, Thank you for the information. I need a letter of acceptance. Thank you, Best regards, Parwanto, MLE [Kutipan teks disembunyikan]

Dear editor,

Automatic Reply: Letter of Acceptance

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[Kutipan teks disembunyikan]



704 Letter of Acceptance 2281 Assangga Guyansyah.pdf 94K

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 24 Februari 2021 pukul 19.34

Dear Editor,

I have received the LOA for our manuscript, but the title of the manuscript is wrong (confused with the script we haven't revised).

Please change the title in the LOA according to the text we have revised. The titles that we have revised are: Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia (our revised manuscript attached).

Thank you, Best regards, Parwanto, MLE

[Kutipan teks disembunyikan]

3 lampiran

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3-BALI MED J-PAID-Invoice - 1096.pdf

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 24 Februari 2021 pukul 21.11

Dear authors,

We had received your revision and had been under the editing process. We will inform you soon about the future progress of your manuscript.

Best regards,

Editorial Board Member [Kutipan teks disembunyikan]

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 25 Februari 2021 pukul 08.00

Dear Author

We are terribly sorry for the technical issue, your manuscript title has been updated on our system, but the Letter of Acceptance is still written with the old title. Hereby we resend to you the revised Letter of Acceptance.

Thank you for trusting us with your hard works Best regards [Kutipan teks disembunyikan] MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Thank you for your information.

[Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Dear Editor,

Previously I apologized for the inconvenience.

Please allow me to change the author's name from:

Mauritius Lmabertus Edy Parwanto Becomes Edy Parwanto (manuscript attached).

There is no change other than the change in the author's name above.

This is necessary to synchronize data at our institution. Thank you. Greetings, Edy Parwanto,

[Kutipan teks disembunyikan]

PRIMAY INFERTILITY-4-submit ke Bali Med J-REV-15-Maret-2021.doc 423K

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 16 Maret 2021 pukul 08.50

Dear author,

Thank you for your confirmation. We will change the author's name becomes Edy Parwanto. If there is anything to change more, please do not hesitate to contact us. Thank you.

Warm regards,

Editorial Board Member [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Thank you for your understanding. Edy Parwanto [Kutipan teks disembunyikan]

25 Februari 2021 pukul 08.32

15 Maret 2021 pukul 20.27

16 Maret 2021 pukul 08.53

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MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com>

Well received with thanks. [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 9 April 2021 pukul 20.10

17 Maret 2021 pukul 08.28

Dear Editor,

Previously I apologized for the inconvenience.

Please allow me to change the author's name from:

Mauritius Lmabertus Edy Parwanto Becomes Edy Parwanto (manuscript attached).

This is necessary to synchronize data at our institution.

In addition, we are also waiting for our manuscript to be published in Bali Med J in the issue of Vol. 10 Number 1, 2021. After that, we will immediately order the hard copy of the journal.

Thank you. Greetings, Edy Parwanto, [Kutipan teks disembunyikan]

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Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com< td=""></edy.parwanto@gmail.com<>

13 April 2021 pukul 12.36

Dear author,

Thank you for your confirmation. We will change the author's name becomes Edy Parwanto. If there is anything to change more, please do not hesitate to contact us. Your manuscript has been reviewing and editing process. We will notify you of further progress. Thank you.

Warm regards,

Editorial Board Member [Kutipan teks disembunyikan]

Dear Author(s),

Attached are the plagiarism report (13%) and the commentary file of your submitted article: Primary infertility of male and female factors, polycystic ovary syndrome, and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency. The commentary file is a summary of the reviews of your manuscript. There were several revisions suggested by our reviewer. Therefore, several necessary changes will be made. Our editor has fixed all sections in your manuscript according to the reviewer's suggestion. We gladly said your manuscript was appropriate for publication. We attached the final manuscript file. We hope you check again your final manuscript before we continue the galley approval process. Please let us know If you agree to publish your final manuscript. If you have any fixation, please send us the final version immediately.

Warm Regards, Editorial Board Member [Kutipan teks disembunyikan]

3 lampiran				
 Turnitin - Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency,.pdf 41K 				
Commentary_BMJ_Primary infertility of male and female factors, polyce oligoasthenoteratozoospermia dominate the infertile population in agr Karawang Regency,.docx 130K	cystic ovary syndrome and icultural and industrial areas in			
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MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com></editorbalimedicaljournal@gmail.com></edy.parwanto@gmail.com>	16 April 2021 pukul 14.10			
Dear Editor Bali Med J Please allow us to send the final manuscripts, please check again. Furthermore, thank you for publishing our manuscript. Best regards, Edy Parwanto [Kutipan teks disembunyikan]				
2 lampiran				
PRIMAY INFERTILITY-4-submit ke Bali Med J-REV-15-Maret-2021 - edit 392K	by BMJ editor.doc			
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Dear authors,

Thank you for your confirmation. We will continue your manuscript to the galley layout process. Thank you for trusting us with your hard work.

Warm regards,

Editorial Board Member

[Kutipan teks disembunyikan]

Thank you for your support! [Kutipan teks disembunyikan]

MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> Kepada: Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> 22 April 2021 pukul 07.26

Dear Editor,

Greetings healthy,

Please inform us if our manuscript entitled "Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia" has been published. Thank you.

Best regards, Edy Parwanto [Kutipan teks disembunyikan]



Accepted with Revision (BaliMedJ) Primary infertility of husband and wife factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia

Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 16 April 2021 pukul 09.48

Dear Author(s),

Attached are the plagiarism report (13%) and the commentary file of your submitted article: Primary infertility of male and female factors, polycystic ovary syndrome, and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency. The commentary file is a summary of the reviews of your manuscript. There were several revisions suggested by our reviewer. Therefore, several necessary changes will be made. Our editor has fixed all sections in your manuscript according to the reviewer's suggestion. We gladly said your manuscript was appropriate for publication. We attached the final manuscript file. We hope you check again your final manuscript before we continue the galley approval process. Please let us know If you agree to publish your final manuscript. If you have any fixation, please send us the final version immediately.

Warm Regards, Editorial Board Member [Kutipan teks disembunyikan]

3 lampiran

Turnitin - Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency,.pdf 41K

Commentary_BMJ_Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency,.docx

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PRIMAY INFERTILITY-4-submit ke Bali Med J-REV-15-Maret-2021 - edit by BMJ editor.doc 566K



Commentary

Dear Sir/Madam,

Here are some commentaries to the manuscript entitled "**Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia**".

No.	Section	Commentary	
A	Title and Affiliation	 The title writing was well written and appropriate for publication. It did not require any revision. Please write the author affiliation number in the same number if the authors come same department. → has edited by BMJ editor Some affiliations were written in <i>Bahasa Indonesia</i>, please rewrite them in English. → has edited by BMJ editor The corresponding section was well written and 	
В	Abstract	 appropriate for publication. It did not require any revision. 1. The abstract was well written and appropriate for publication. It did not require any revision. 	
C	Introduction	1. The introduction was well written and appropriate for publication. It did not require any revision.	
D	Method	1. The method was well written and appropriate for publication. It did not require any revision.	
E	Result	 It would be better if the manuscript describe generally the data about Karawang Regency than just present the figure of the Karawang Regency or it would be better if did not use the figure of Karawang Regency. Please fix this problem. → has edited by BMJ editor 	
F	Discussion	1. Please rewrite sentence which use <i>Bahasa Indonesia</i> in <i>Italic</i> format.	
G	Conclusions	1. The conclusion was well written and appropriate for publication. It did not require any revision.	



Η	Table, figure	1. Our journal adopts the "Vancouver Superscript" as the		
	and	choice of citation format. Please format your <i>inline citation</i>		
	Reference	and <i>bibliographic</i> as an example given below in: \rightarrow has		
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		Inline citation		
		Ponten et al., showed that fasciocutaneus flap could be		
		utilized to cover lower leg soft tissue defects. ¹		
		Bibliographic		
		1. Pontén B. The fasciocutaneous flap: its use in soft tissue		
		defects of the lower leg. Br J Plast Surg.		
		1981;34(2):215–20. Available from:		
		http://www.ncbi.nlm.nih.gov/pubmed/7236984		
Ι	Others	1. There were numerous grammatical errors with a total 186		
		errors. Please improve this condition. \rightarrow has edited by BMJ		
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------Study design specific checklist goes here-----



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Proofread, edit format, and BaliMedJ Guidelines Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia	1	200,00 \$	200,00 \$
		Subtotal	1.200,00 \$
		Total	1.200,00 \$
		Amount paid	-1.200,00 \$
		Amount due	0,00 \$ USD

Notes

After an initial cross-checks, it's a pleasure for me to inform you that your manuscript entitled: Primary

infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency, West Java Province, Indonesia.

is Worthy of Publication.

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Editor Bali Medical Journal <editorbalimedicaljournal@gmail.com> Kepada: MAURITIUS LAMBERTUS EDY PARWANTO <edy.parwanto@gmail.com> 23 Februari 2021 pukul 10.02

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Do inform us if you need the letter of acceptance prior to the publication. Thank you for trusting us with your hard works. Best regards [Kutipan teks disembunyikan]





Letter of Acceptance 24 February 2021

Dear: Assangga Guyansyah¹, Raditya Wratsangka¹, Denny Dhanardono¹, Muhammad Farid Ghazali², Hosea Java Edv³, Harvo Ganeca Widyatama⁴, Dietha Kusumaningrum⁵, David Tjahyadi⁶, Mauritius Lambertus Edy Parwanto^{7*} ¹Department of Obstetrics and Gynecology, Faculty of Medicine, Universitas Trisakti, Indonesia ²Policlinic of Infertile, Mitra Bunda Amanda Karawang, Karawang Regency, West Java, Indonesia ³Study Program of Pharmacy, Faculty of Math and Natural Sciences, Universitas Sam Ratulangi, Indonesia ⁴Medical doctor of Bhakti Mandala Clinic, Tangerang Regency, Banten, Indonesia ⁵Medical doctor of Batari Husada Clinic, Duren Sawit, East Jakarta, Special Capital Region of Jakarta, Indonesia ⁶Department of Histology, Faculty of Medicine, Universitas Trisakti, Indonesia ⁷Department of Biology, Faculty of Medicine, Universitas Trisakti, Indonesia *Corresponding author: edyparwanto@trisakti.ac.id

I am very excited to accept your paper entitled: "Primary infertility of husband and wife factors, polycystic ovary syndrome, and oligoasthenoteratozoospermia." Your paper will be published in the issue of Vol. 10 Number 1, 2021. http://dx.doi.org/10.15562/bmj.v10i1.2281 (Online Link: http://balimedicaljournal.org/index.php/bmj/article/view/2281).

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Please do not hesitate to contact us if you need anything. It has been a pleasure for us to proofread and edit your work, and we are looking forward to your colleagues and your other papers in the near future.



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Submission date: 13-Apr-2021 06:58PM (UTC+0700) Submission ID: 1558060258 File name: PRIMAY_INFERTILITY-4-submit_ke_Bali_Med_J-REV-15-Maret-2021.doc (423K) Word count: 5100 Character count: 32105 Primary infertility of male and female factors, polycystic ovary syndrome and oligoasthenoteratozoospermia dominate the infertile population in agricultural and industrial areas in Karawang Regency,

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