## Article Review: Effect of Electronic Cigarette on Fertility in Male

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#### ABSTRACT

Numerous researches have shown that blood plasma, sperm parameters, or a host of other fertility-related characteristics are adversely affected by the components of cigarette smoking. However, it's unclear how smoking really affects fertility in men. Depending on the well-established scientific observation that smoking enhances the amount of radical oxide organisms, which in turn causes oxidative stress (OS), smoking has an influence on the sperm variables.

Men are affected by OS as it damages seminal characteristics including survival that appearance as well as inhibits sperm functioning. Still, not every research has reached identical findings. This research evaluates the influence of tobacco use via nonsmoking methods on male infertility on clarifies the contentious relationship among smoking on fertility in men. Additionally, it explores the therapeutic consequences of the evidence—which includes recently discovered genomic or regulatory data—that tie smoking to male infertility.

Keywords- e-cigarette, reproductive health, male infertility, oxidative stress (OS), smoking, blood plasma, sperm parameters.

#### I. INTRODUCTION

Called by another name, "e-cigarette," electronic cigarettes are a unique method of delivering the nicotine that were created around that Kingdom of China in 2003. Those who use electronic cigarettes are often referred to as "e-smokers" or "vapers"3. It is presently becoming more as well as more prominent among young, affluent people living in cities all over the world<sup>[1]</sup>. Children are drawn to e-cigarettes due to its innovative technological advances; wide range of tasty flavour choices, as well as marketing that makes many promises about how much safer e-cigarettes are than traditional cigarettes and just how easy it is to quit smoking.

Even though public health officials have major concerns, e-cigarettes have become more prevalent as accepted among consumers, particularly teenagers, and sales are growing geometrically yearly<sup>[2]</sup>. The e-cigarette market is expanding rapidly, as well as in regarding ten years could overtake traditional tobacco. An electronic cigarettes industry has over 200 brands, with revenues projected to exceed US\$ 3 billion in 2013. By 2017, the worldwide e-cigarette industry was predicted to reach \$10 billion USD5.

The majority of e-cigarettes are made to look even feel as conventional cigarettes, however others could be either or longer compared this or mimic fashion items like USB sticks, lipstick, pens, or pencils[3-4]. An aerosol generator, battery, flow sensor, as well as solution (eliquid) holding container are the key parts of the majority of e-cigarettes. An e-cigarette could be dangerous because to its ingredients, which include lithium-made batteries, plastics, steel, metals, silver, ceramics, fibres, aluminium, rubber, or spume[5]. This research aims to draw attention to the medical risks associated with e-cigarette smoking in light of the widespread usage of e-cigarettes or concerns among the public.

Smoking is still the global problem even after it has been shown to have harmful effects on both overall healths as particularly. The World Health Organisation (WHO) reports that 30% of males aged 15 or older smoke[6-7]. About 46% of smokers are men who are of age to reproduce (20–39). The real number of smokers climbed from 721 million in 1980 to 967 million for 2012, despite the fact that the proportion of adult smokers, based

on regionally relevant statistics from 187 countries, declined from 41.2% in 1980 to 31.1% in 2012.

By the World Health Organisation, infertile affects 8% for spouses globally including 10% - 15% of spouses throughout developed countries, with male factor contributing to 30% - 35% of instances. Much research has been carried out to look for potential links among smoke with masculine infertility; however, certain of the findings had been inconsistent[8]. Smoking has been linked to reduced sperm motility or increased nuclear DNA damage in sperm, corresponding to various although several researches has revealed no negative consequences as well as certain instances, possibly favourable benefits on masculine infertility or semen analytical characteristics.

There are other contradictory reports of evidence addressing the impact of cigarettes on infertility among men. Furthermore, research examining how smoking affects semen parameters has not conclusively shown that smoking affects male fertility[9-10]. It is hardly unexpected if the findings are contradictory and unclear. It is feasible to formulate three theories to account for the development of contradicting findings.

First, sperm cells functioning, histological changes, sperm parameters, or additional measures are utilised to evaluate the impact of smoking on male fertility. Research may provide conflicting results if alternative evaluations are used. Second, it is unclear how smoking may affect male fertility via well-established processes[11]. Third, since it is tough to account for variables such as exposure to consumption of alcohol, illnesses, environmental pollutants, as well as socio-economic circumstance, it is impossible to precisely contrast findings between research.

Aforementioned perspective, Researchers go over the ingredients in cigarettes including the specific ways that tobacco exposure affects infertility among men[12]. The effects of smoking on various components of the male reproduction system are then described, with an emphasis on both the physiologic or pathological consequences of smoking[13-14]. Afterwards, this work provides an overview of the potential pathways by which smoking affects masculine reproduction functioning via genetic or environmental modifications—a subject that hasn't been addressed thoroughly before. Lastly, they talk about how this field of research has to develop in order to serve the therapeutic requirements of smokers who are barren.

Instead of inhaling smoke, they may inhale nicotine vapour thanks to a vaporizer and e-cigarette. A water-based liquid that could include flavorings, nicotine, and various other components is heated by the vape device[15]. The consumer inhales the resulting liquid vapour. (This functions identically to vape pens for cannabis, which use THC rather of nicotine.)

E-cigarettes have the advantage of not burning tobacco and producing smoke. This indicates that they aren't releasing carbon gases carbon monoxide and tar, https://doi.org/10.55544/jrasb.3.1.33

two of the most harmful components of tobacco smoke[16-17]. To be exact, tobacco smoke contains 7,000 different compounds, 250 from which are deemed hazardous, including 69 of them are carcinogenic (the substances that cause cancer).

#### II. THE *E-CIGARETTE* DEVICE

Several research investigations dealt with the substance of the electronic gadget and its possible repercussions especially an interest inclusion of metallic substances like metals such as nickel, copper or silver particles fragments within e-liquids as well as aerosols beginning about their threads as well as cables along with the atomizer itself[18-19]. Although a great deal of investigations connected with effects that using e-cigarettes in the health of people concentrated on the e-liquid elements along with ensuing particles generated following heating.

Silica particles from the fibreglass wicks and silicone are additional significant ingredients in the aerosol. Although it is recognised that several of these items may lead to anomalies in the respiratory system including respiratory conditions, greater research is needed[20]. Remarkably, e-liquids with a greater batteries voltage of output seem to be more harmful for A549 tissues, suggesting that the batteries input voltages could additionally affect the cytotoxic in the aerosols vapours. During a recent research, the immediate consequences of e-cigarette vapour produced by a nickel-chromium alloy (NC) and the stainless-steel atomizer (SS) heating element (containing cannabis flavoured and PG/vegetable glycerin with no nicotine) were investigated [21-22]. A final group of rodents had been exposed to fresh air over two hours, while certain rodents had just one treatment to e-cigarette vapour over two hours from the NC heating component (60 - 70 W); others experienced an identical treatment to e-cigarette vapour over a comparable amount duration through the SS generating element (60 or 70 W). Mice exposed to air or e-cigarette vapour via SS heating components did not experience respiratory discomfort.

Conversely, if the 70-W output level was used, 80% for those mice acclimated with electronic cigarettes vapour with NC heated devices had clinically severe difficulty breathing. implying that using operational units over what is advised may have unfavourable consequences[23]. However, it is certain that the harmful consequences of batteries discharge voltage are not equivalent to those imposed by CS extraction.

#### III. E-CIGARETTES: TYPES, USAGE

E-cigarettes had evolved significantly when researchers were introduced to the market (Figure 1). These gadgets, which go by numerous names, were produced in a wide variety of forms, dimensions, or kinds. To date, four distinct generations of electronic cigarettes have been created (Figure 1)[24]. In general, a device's

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primary parts are still identical as well as contain of a fluid-filled cartridges, a piece of equipment which heats the e-liquid to create the aerosol, an electrode that activates an equipment, as well as a battery that supplies the electricity needed to heat the vaporizer.

Considering the original e-cigarettes were neither refilling nor battery-operated, these are exclusively made for "one-time use." The second series of devices evolved to include replaceable cells or refillable e-liquid cartridges[25-26]. The third version allowed users to customize the ingredients in the e-liquid that was made to last for several uses. The fourth generation of ecigarettes, called Pod-Mods, arrived in a huge range of dimensions, shapes, but colors as well as had both the characteristics of the prior version.



Figure 1. Description of the four generations of e-cigarettes.

There are currently over 500 brands however 8000 flavors available on the market for e-cigarettes, and the market is constantly changing [27]. adolescents had taken a strong interest in these gadgets. Actually, 85% of persons aged 18 to 29 reported having experienced smoking in 2018, while being exposed to advertisements was closely linked to the consumption of this product. But because of new pricing and licensing, the use of ecigarettes has decreased. Varied age groups have varied perceptions about using e-cigarettes[28]. Youth or young people might think of the use of these tools as trendy on the one hand. In actuality, forty percent of young people who use e-cigarettes had never smoked. The e-cigarette was proposed as a gateway to traditional smoking of cigarettes according to these results. Yet, among older users, using e-cigarettes helped them stop smoking traditional cigarettes[29]. This resulted from the widespread perception that using electronic cigarettes is less dangerous than smoking traditional cigarettes. However, e-cigarettes include a variety of pollutants which expose users to compounds that may have an adverse effect on their overall health.

## IV. EVIDENCE ANALYSIS OF THE IMPACT OF E-CIGARETTE ON MALE REPRODUCTION

While there has been little research on how ecigarettes affect male reproductive among humans, several organizations have looked into how they affect animal models. It has been shown that e-cigarette exposure disrupts the hypothalamo-pituitary axis, which modifies gonadal function or semen quality (Figure 3). Fact, showed that masculine mice exposed to e-cigarette vapour exhibited elevated apoptotic for spermatogonia or spermatocyte, modified seminiferous epithelium shape including function, as unica albuginea abnormalities [30]. Further investigations connected the use of e-cigarettes to disruptions in steroidogenesis, worldwide disarray in the testicles, or a notable elimination of germ cells.

Furthermore, smoking e-cigarettes has been linked to lower testicles weight or a greater proportion in dead cells within the testis. Masculine mice administered e-cigarette liquid intraperitoneally, leading to cytotoxicity or inflammatory of the testicles[31]. This, in consequently, had an impact on testicular generation or effectiveness, resulting in decreased sperm density, fewer epididymal sperm, or reduced male survival.

Research revealed that smoking e-cigarettes may potentially have an impact on the quality of the sperm chromatin. In actuality, treated rats' testicles or sperm showed greater levels of DNA damages[32]. These outcomes point to possible germline carcinogenic consequences for e-cigarettes.

There are very few human research investigations that support these conclusions. The British Fertility Society Conference in 2017 featured an initial investigation that examined the impact of e-cigarette flavoring on human sperm. The results indicated a noteworthy reduction in velocity in specimens that were cultured with e-liquid flavoring[33]. Vaping may have adverse consequences on male procreation, according to the findings of this research as well as the outcomes of animal models, so people who vape as well as are considering conception ought to apply caution.

An initial paper addressing that impact on ecigarette recharge fluids, both with or without nicotine, upon rodent testes using an experimental approach animals (Figure 2). Wistar rats weighing  $160 \pm 20$  g were given intraperitoneal injections with electronic cigarettes refilling fluid every day throughout 4 weeks[34]. The findings demonstrated that e-cigarette refill fluids, whether or not these included nicotine, caused oxidative stress in rodent testes, which significantly increased the activities of antioxidant proteins such cysteine Stransferase, catalase, as well as superoxide dismutase.

Testicular tube fluids were disorganized as germ cells prematurely shed from the seminiferous the

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epithelium among other histopathologically-observed alterations in the testis architecture. Mostly testosterone, which are compounds that broadly affect the anatomy or physiology of the testicular[35-36]. Since e-liquid exposure reduced cellular messenger RNA (mRNA) conveying of two important steroid production digestive enzymes, chromium P450scc and 17 $\beta$ -Hydroxysteroid dehydrating enzymes (17 $\beta$ -HSD), circulatory the hormone testosterone the amount decreased substantially (by 50% - 30%, accordingly) [37]. The previously number or survival of sperm extracted from the epididymis cauda were significantly lower.



Figure 2: The effect of e-fluid with and without nicotine on male reproductive system.

The impact of e-liquids with or without cannabis on rodent epididymides was examined during a follow-up investigation using an identical creature model. Both in or rather than the nicotine, access to e-liquids caused a considerable drop for the quantity of epididymal sperm cells [38]. Remarkably, the sperm counts in the rats handled by the non-nicotine liquid was  $32.3 \pm 3.0$ million/mL or 38.4 ± 0.9 million/mL, accordingly, compared to the other subgroup handled with nicotine. Assessing the vitality of sperm for treated rats yielded similar findings, with 27.0%  $\pm$  4.6% to 42.8%  $\pm$  5.1% of viable sperm[39]. A morphological analysis revealed a statistically significant rise in the proportion of aberrant sperm, particularly in the rats subjected to the nicotinefree liquid compared to the nicotine-containing liquid compared the untreated group, at  $43.0\% \pm 1.0\%$  vs. 30.2% $\pm$  1.8% vs. 24.0%  $\pm$  0.9%. All experiment receiving treatment showed a significant drop in plasma androgen of testosterone following 4 weeks on therapy.

The research rats' epididymides showed impacts caused by the e-liquid upon their antioxidant state.

In an associated research, tested a novel ecigarette gadget on seven-week-old Sprague-Dawley rats. The previously instrument was loaded with a nicotine-free liquid that contained 50% vegetables glycerin (VG) as well as 50% propylene glycol, or (PG) at an impedance corresponding among, which is the least possible levels[40]. The vapour form the e-cigarettes was subjected to the 7 rodents during their control group for 28 days, three hours a day, during 11 cycles of two puffs. The e-cigarette cartridge had been filled with PG or VGbased, 10% red fruit-flavored fluid that was nicotinelike free[41]. Toxic aldehydes formaldehyde, acetaldehyde, or acrolin may be found in the vapor according to the scientific examination. Findings from the literature indicate that male animals exposed to aldehydes had changes in the activities of the steroid production enzymes. Absorption to vapor from e-cigarettes has been demonstrated to decrease the levels of two essential enzymes involved for steroid production, 3β-HSD or 17β-HSD[42]. Consequently, there were a considerable reduction for the activities of the testicles indicator proteins, sorbitol dehydrogenase (SDH) with glucose-6phosphate dehydrogenase (G6PDH), but a modest rise in the indicator of tissue destruction, lactate dehydrogenase (LDH). Major testicles from its subjected mice showed elevated concentrations of reactive oxygen species (ROS) as well as an essential rise in the production of protein carbonyls or peroxidation of lipids components[43]. Changes are additionally observed for the antioxidant as well as detoxifying metabolic pathways. The findings indicate that alterations occurred in the testicular structure

including the cellular architecture of the seminiferous the epithelium despite the fact that the appearance of the testicles hadn't been investigated. The researchers concluded by stressing that being subjected to vapor from a low-voltage, nicotine-free e-cigarette liquid caused the disruption of digestive enzymes related to seminiferous epithelial cells action and steroid production, indicating potential reproductive health disabilities.

Regarding its impact for e-cigarette usage upon spermatozoa, no conclusions are released. However, Helen O'Neill made the suggestion that harmful compounds found their spices of electronic cigarettes might harm men's infertility at the 2017, British Fertility Society Conference[44]. She presented outcomes of an investigation whereby the tastes of gum with bubbles when spice had been added to the medium as men's sperm cells were subjected to it.

The research's taste concentrations were comparable to what regular or infrequent e-cigarette users typically consume. Thirty male applicants that are enrolled in an in vitro fertilization (IVF) program had their sperm samples collected. According to these findings, spermatozoa's velocity may be considerably decreased by vaping flavours with cinnamon, which is which makes the particles move less quickly[45-46]. In the second portion of the research, male mice were given access to the tastes of bubble gum as well as cinnamon, two of the most famous foods, to see how they affected the Somniferous epithelium (O'Neill (2017) as well as her children). It was discovered when the bubble-gum taste harmed the mice's testicular germinal cells. The research found that while bubble-gum taste killed the testicular tissues that produce testosterone, cinnamon flavor affected the movement of spermatozoa.

Male mice were utilized during a research to examine overall impact of smoking and e-vapor consumption on fertility. The seminiferous epithelial had structural or physiological defects as a result of exposure to smoking and e-vapor, including vaporization, decreased sperm formation, or increased sperm cells or spermatocyte death[47-48]. Furthermore, minor alterations in the shape of sperm were noted. Rats exposed to conventional cigarette smoking showed greater alterations, whereas the effects of vapour on male reproductive systems were marginally less apparent.

Human majority of research on the impact of ecigarettes on the male reproductive systems used animal models. Nonetheless, the information presented herein points to pathogenic changes in the human reproduction system's cells, tissues, or hormones as a result of vaping[49]. Teenagers who identify as male must know that flavor- or nicotine-free e-cigarettes may have identical negative effects on their biological systems including potential for parenthood as nicotine-based ecigarettes.

By stimulating the ejection of development hormone, cortisol, blood pressure medication, or testosterone, that in turn inhibits the production of LH or https://doi.org/10.55544/jrasb.3.1.33

PRL, smoking may modify the hypothalamic-pituitary axis. While the average concentrations of T or dehydroepiandrosterone (DHEA) are similar, smoking had greater median concentrations of 17 beta-estradiol (E) or lower mean amounts of LH, FSH, or PRL compared to non-smokers[50]. People who smoke had lower the PRL or higher levels of both liberated and whole blood T, according to Trummer et al.

It has been demonstrated indicating smoke was substantially linked to a reduction in seminal PRL. A good dose-response relationship between smoking or T, LH, especially LH/free T ratios was conversely, there were no important variations on the levels of FSH, LH, as well as whole T among the 889 viable males who were classified as light, mild, and avid smokers.

#### V. NICOTINE AND MALE FERTILITY

Just nicotine by itself may be harmful. The 2017, research upon mice looked at the impact of a long-term low dosage of nicotine on the properties of sperm. It emerged that there had been a correlation between smoking or decreased sperm count, movement, or survival as well as a rise in the quantity of men with aberrant morphology[51-52]. considerably lower sperm velocity or determine, considerably higher aberrant morphology, lower libido, as well as lower litter weight as well as number of offspring per litter were all linked to nicotine-only therapy, according to another research on rodents.

One inhalation from an e-cigarette machine can have anywhere from 0 to 35 mg of nicotine, based on the manufacturer of the e-liquid. The majority of that times, that amounts of nicotine found exceeded the amount of nicotine that the manufacturer had declared[53]. Interestingly, nicotine was found in "Nicotine-Free Products" as well. There are significant differences in the amount of nicotine in e-liquid across nations since ecigarette production is often unregulated.

#### VI. VAPING AND MALE FERTILITY

Many individuals think e-cigarettes, or "vaping," is healthier than cigarette smoking because they were attracted to its sweet tastes, attempting to get themselves off of regular nicotine goods, or perhaps because it reduces hunger. However, the CDC has just issued a warning on EVALI, a freshly created term that stands for E-cigarette Vaping-Associated Lung Injury[54]. Furthermore, while the consequences of smoking on the lungs are well established, the harm it does to the human reproductive system is still being investigated.

Vaping allows unidentified or potentially hazardous substances to enter the circulatory system as well as body. Electronic cigarettes, often known as ecigarettes or e-pipes, are a kind of device that dispense nicotine electronically[55]. Other names for these devices

include vaporizers, hookah pens, vapes, or vape pens. These products employ a liquid that might contain nicotine along with different combinations of veggie glycerin, propylene glycol, or flavors. The individual breathes an aerosol that is produced by heating the liquid[56-57]. These cancer-causing substances enter the respiratory tract deeply, irrigating the bronchi, raising blood pressure or heart rate, impairing the heart's ability to receive enough blood, or constricting the arteries. In addition, the "vapor" that is breathed causes gum disease, persistent coughing, and slowed recovery from wounds. Vapour produced by using an e-cigarette contains a variety of substances, such as formaldehyde, TSNA, acrolein, glycidol, VOCs, as well as PAH. Rodent species were used to analyze formaldehyde's reprotoxicity in great detail. It demonstrated the capacity to cause sperm parameter abnormalities, change reproductive behavior, as well as change testicles anatomy. Long-term formaldehyde treatment raises oxidative stress, which has detrimental effect upon rodent reproductive а histopathology and causes a sharp decline in the quantity and size of developed ovarian follicles[58]. Formaldehyde is known to be reprotoxic, notwithstanding the paucity of human research on the subject. Men subjected to formaldehyde vapor while at work had a deleterious effect on sperm mobility. Comparably, VOC as well as PAH consumption in the environment has a negative impact on endocrine function or semen quality (sperm counts as well as morphology), which may lead to problems with reproduction . It is well recognized that volatile organic compounds (VOCs) negatively impact the development of embryos, which lowers the likelihood of successful IVF treatments due to reduced frequencies of insertion as well as conception[59]. During 2017, an investigation headed by the University College London found a connection between flavored e-cigarettes and harm to sperm. Additionally, another Danish research from 2020 discovered that males who vaped had less sperm.

A comprehensive analysis of vaping and male fertility that was published in 2020 found that the toxic ingredients in e-cigarettes may alter the morphology, operation, or hormonal equilibrium of the reproductive systems.

Additional research has shown that vaping may still be harmful to reproductive wellness in spite of cases when it contains no nicotine.

Mice were exposed to e-cigarette refilling juice in a 2016 research. This research discovered that exposure was linked to lower level of testosterone, sperm count, or fertility of sperm irrespective of when the liquid used for vaping included nicotine or not[60-61]. A 2019, research using nicotine-free e-cigarette juice on rats revealed that it had an impact on the testicles or the reproductive systems of men. This limited body of research on smoking on human reproduction suggests that vaping could have a deleterious effect on male fertility, even though additional study is required[62]. Regarding potential long-term https://doi.org/10.55544/jrasb.3.1.33

effects of vaping upon physiological wellness, there remain many unresolved Note that a large number of research on vaping with male fertility have been done using animals. Even though there is solid evidence to support the possibility that vaporization or nicotine could be dangerous, much better research remains needed in this field.

#### VII. VAPING HAS BEEN LINKED TO LOW SPERM COUNT AND EMBRYO IMPLANTATION PROBLEMS

Additionally, e-cigarettes have been shown to lower sperm from men count, motility, and integrity. For individuals seeking to become pregnant, cell eggs is almost difficult due for inadequate male reproductive cells of sperm[63-64]. And things don't stop there. Male children exposed to toxic chemicals or vapors from ecigarettes during pregnancy have an increased risk of developing reproductive health problems as adults.

Furthermore, similar poisons may cause physiological disabilities including reduced development as well as additional developmental problems in female newborns[65]. Glycerin from vegetables or polypropylene glycol was substances that exist in ecigarettes. There is evidence connecting exposure to these substances with severe infertility or reproductive problems in women.

Research reveals that the initial or persistent existence of such toxins in a human body may significantly hinder transplantation inside a female reproductive system, notably addition to cases when an egg is fertilized or prepared for insertion. Finally, vaping during pregnancy increases the likelihood that the baby growing within the woman's uterus may suffer bodily injury[66-67].

A growing health issue was the growing popularity on e-cigarettes by men or women throughout their adolescent years. Although investigation into the potential harm that vaping may do to its biological systems are still at its early stages, considering its novelty of vapers just like a social phenomena or a substitute for conventional cigarettes items, other tobacco products, preliminary findings indicate:

• A cross-sectional research of adolescents within the population as a whole found that e-cigarette use is linked to decreased numbers of sperm. In fact, corrected studies showed that daily e-cigarette users (147 million vs 91 million) and daily cigarette smokers (139 millions versus 103 million) had considerably lower overall sperm counts than non-users[68]. Significantly, this represents the first human research to show a link between decreased sperm counts and both cigarette and e-cigarette usage.

• Given that e-cigarettes are constantly thought to be less dangerous than traditional tobacco products that information may be essential to men who were attempting

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to conceive, particularly those who already have poor sperm counts.

#### VIII. THC AND MALE FERTILITY

Whatever happens once you vape THC? The problems listed above with extra components in vape juice still hold true. Additionally, issues with male reproduction are connected to THC, a primary component of medical marijuana[69]. Research from 2019 or 2021, to name a couple among numerous comparable research, revealed an adverse association between cannabinoid usage or decreased testosterone volume, quality, or shape as well as a detrimental effect on male reproduction.

#### IX. **EFFECTS OF SMOKING ON MALE FERTILITY**

In instance, the harmful consequences of smoking cigarettes have an impact not only on the cardiac or pulmonary systems yet on male fertility. The capacity of a man to generate sperm of a great calibre along with appropriate movement, count, or appearance is known as male fertility. Smoking tobacco products, nevertheless, will lower sperm motility or output[70]. Smoking, however, also results in aberrant sperm morphology. Sperm is necessary for maintaining the progeny or fertilising the egg.





Few things were documented about how using ecigarettes affected fertility, despite knowing that cigarette smoking has been linked to lower fertility among men in the past. The Journal of Human Reproduction has released a research that demonstrates the link between regular ecigarette usage and decreased sperm counts[71]. Using a population-based, cross-sectional design, the investigators of their research included 2008 males (average ages were 19) who completed a way of life evaluation which sought information on smoking habits in addition to providing a specimen of sperm as well as blood sampling. Both daily users of e-cigarettes (147 million versus 91 million) and everyday cigarette smokers (139 million versus 103 million) have substantially fewer overall the sperm levels than non-users.

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Numerous experts are perplexed by just diminishing productivity or integrity of men's semen during the prior few years. Toxic substances from the environment, inadequate food, or overweight constitute a few of the causes for this deterioration. With further research focusing on the negative effects of health-related behaviours like vaping nicotine or marijuana, additional components of the jigsaw are starting to fit together[72]. Men's reproduction had previously been shown to be negatively impacted by marijuana use. Since men make up the bulk of those who use marijuana for medicinal or social reasons, the pattern of falling fertility will only become worse as legalisation expands throughout the country.

The risks associated with maternal consumption of marijuana during her pregnancy are well documented. Upon another hand, recent research have started focusing in particular impact of parental marijuana usage upon the male reproductive epigenome including the potential for causing unfavourable birth outcomes[73]. There is even a little research that suggests a link between marijuana usage by fathers with a higher chance of autism in their children.

Further retrospective research is required to clarify the effects of mother or parental marijuana consumption on children all through infancy, youth, or teenage years, as marijuana usage becomes more common. Right now, women who are expecting or want to become pregnant are being told to cease using cannabis altogether by the American College of Gynaecologists as well as Obstetrics. It's possible that the American College of Urologists might consider recommending males who want to fathers kids on a comparable way.

#### X. SMOKING AND CHROMOSOMAL DAMAGE

Endometriosis may be caused by significant DNA damage that could impede oocytes fertilisation and the growth of the embryo, according to an array of articles on male smoking. In Golgi-phase or cap-phase spermatids, genomic to chromosomes occurred at rates of 1.15% in impotent smoker or 0.82% in sterile nonsmokers. People who smoke were discovered to have a much larger proportion for a single-strand versus doublestranded DNA Spermatozoa, along with a diminished chromosomal fertilisation of potential.

It was calculated that masculine those who smoke had a higher proportion of spermatozoa having fragmented DNA than non-smokers (4.7% vs. 1.1% in one research or 32% versus. 25.9% for others[74]. A deleterious impact of smoking was found by analysing seminal Chromosomal disintegration following stimulation, which changed the sperm swim-up procedure for choosing in users. However, among contrast to nonsmokers, some researchers could not discover a link among smoking or fragmentation of DNA in the

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spermatozoa of competent moderate to avid smokers. However, they did note a definite unfavourable tendency, particularly with regard to the disruption of the plasma membrane's phospholipid asymmetries.

Smoke has been proven to increase disomic sperm cells, which has been linked with the increased risk of anaemia, but only for the occurrence of disomy. Furthermore, Horak et al. found that amongst infertility individuals, there was a negative association between sperm motility versus number as well as an important distinction (1.7-fold increase) among current tobacco users or non-smokers.

#### XI. SMOKING AND ACROSIN

When male circulation or concentration is normal, smokers have been seen to have reduced acrosin activity. Acrosome response was found to be nonsignificantly different in sperm cells from individuals having the varicocele, although it was considerably lower in samples of sperm from smoking compared to viable individuals. When comparing varicocele-associated instances to normal men, the proportions of spermatozoa with halo production or halo dimensions were both considerably reduced.

## XII. SMOKING AND TOXIC MATERIALS IN SEMEN

Sperm count or shape is significantly impacted by nicotine. Seminal cotinine or trans-3'-hydroxycotinine concentrations among smoking have been demonstrated to be comparable with blood phases; however semin nicotine concentrations were much higher. Upward sperm mobility was linked to seminal the amino acid cot stages, whereas overall sperm motility was inversely connected with trans-3'-hydroxycotinine or seminal cotinine concentrations.

It had also been demonstrated that seminal nicotine but cotinine concentrations could be measured by response to passively exposure to ambient tobacco smoke, as well as that these levels are associated with the stated amount of exposure.

Smokers who consume more than twenty cigarettes per day have been shown to have elevated seminal cadmium (Cd), with there is a substantial negative link between Cd in blood with testicular quality as well as cigarette years. In normozoospermics, was has been demonstrated that seminal Cd had been linked to the number of cigarettes smoked daily, with smokers having greater levels than non-smokers[75]. Additionally, it has been documented that sterile smoking has greater levels of leads throughout their primordial blood than both productive males and infertile non-smokers.

Consuming increases benzo (a) pyrene diol epoxide-DNA conjugates in sperm cells; smokers have much greater amounts of these additives than nonsmokers do, suggesting a significant ecological https://doi.org/10.55544/jrasb.3.1.33

component as well. One possible cause of transmissible pre-zygotic DNA degradation to sperm cells is the creation of conjugates.

#### XIII. CONCLUSION

The potential consequences of smoking on male fertility were covered in this review. Certain conclusion could have taken, despite the fact that no exact or unequivocal findings are possible. Initially compared to sub-fertile males, productive men are more significantly affected by smoking both terms of sperm composition or functionality. This might be explained by decreased sperm cells functioning at these fundamental levels or sperm characteristics in the infertility populace. It wasn't possible to determine with certainty how smoking influenced semen characteristics based on а comprehensive evaluation of the available information. yet conception may still be impacted by typical sperm characteristics. Furthermore, a plausible explanation for how consuming affects the infertility of men is that OS or the genomic and environmental alterations it causes may be closely associated with decreased reproductive functioning or infertility. More research must be done on which association. Third, there are dose-dependent relationships between smoking or testicular or sperm quality. Every man smoking must definitely recommend quitting, particularly if he is attempting to conception with his spouse. Healthcare professionals should encourage people to stop smoking by providing information, guidance, or ongoing care. The information about smoking particularly male fertility supports the recommended preventative strategy of banning tobacco use or limiting tobacco smoke consumption for both sexes generally, as well as specifically during the reproductive process.

#### REFERENCES

[1] Rutstein S.O., Shah I.H. *Infecundity Infertility and Childlessness in Developing Countries*. World Health Organization; Geneva, Switzerland: 2004.

[2] Gore A.C., Chappell V.A., Fenton S.E., Flaws J.A., Nadal A., Prins G.S., Toppari J., Zoeller R.T. EDC-2: The Endocrine Society's Second Scientific Statement on Endocrine-Disrupting Chemicals. *Endocr. Rev.* 2015;36:E1–E150. doi: 10.1210/er.2015-1010.

[3] Segal T.R., Giudice L.C. Before the beginning: Environmental exposures and reproductive and obstetrical outcomes. *Fertil. Steril.* 2019;112:613–621. doi: 10.1016/j.fertnstert.2019.08.001.

[4] Chen R., Pierce J.P., Leas E.C., Benmarhnia T., Strong D.R., White M.M., Stone M., Trinidad D.R., McMenamin S.B., Messer K. Effectiveness of ecigarettes as aids for smoking cessation: Evidence from the PATH Study cohort, 2017–2019. *Tob. Control.* 2022 doi: 10.1136/tobaccocontrol-2021-056901.

https://doi.org/10.55544/jrasb.3.1.33

[5] Hutzler C., Paschke M., Kruschinski S., Henkler F., Hahn J., Luch A. Chemical hazards present in liquids and vapors of electronic cigarettes. *Arch. Toxicol.* 2014;88:1295–1308. doi: 10.1007/s00204-014-1294-7.

[6] Pisinger C., Døssing M. A systematic review of health effects of electronic cigarettes. *Prev. Med.* 2014;69:248–260.

doi: 10.1016/j.ypmed.2014.10.009.

[7] Thirión-Romero I., Pérez-Padilla R., Zabert G., Barrientos-Gutiérrez I. Respiratory impacy of electronic cigarettes and "low-risk" tobacco. *Rev. Investig. Clin.* 2019;71:17–27.

[8] Bertholon J.F., Becquemin M.H., Annesi-Maesano I., Dautzenberg B. Electronic cigarettes: A short review. *Respiration*. 2013;86:433–438. doi: 10.1159/000353253.

[9] Ali F.R.M., Dave D.M., Colman G.J., Wang X., Saffer H., Marynak K.L., Dench D., Grossman M. Association of e-cigarette advertising with e-cigarette and cigarette use among US adults. *Addiction.* 2021;116:1212–1223.

doi: 10.1111/add.15281.

[10] Jun J., Kim J.K. Do state regulations on e-cigarettes have impacts on the e-cigarette prevalence? *Tob. Control.* 2021;30:221–226. doi: 10.1136/tobaccocontrol-2019-055287.

[11] Sapru S., Vardhan M., Li Q., Guo Y., Li X., Saxena D. E-cigarettes use in the United States: Reasons for use, perceptions, and effects on health. *BMC Public Health.* 2020;20:1518. doi: 10.1186/s12889-020-09572-x.

[12] Kosmider L., Sobczak A., Fik M., Knysak J., Zaciera M., Kurek J., Goniewicz M. Carbonyl Compounds in Electronic Cigarette Vapors: Effects of Nicotine Solvent and Battery Output Voltage. *Nicotine Tob. Res.* 2014;16:1319–1326.

doi: 10.1093/ntr/ntu078.

[13] Goniewicz M.L., Kuma T., Gawron M., Knysak J., Kosmider L. Nicotine Levels in Electronic Cigarettes. *Nicotine Tob. Res.* 2012;15:158–166. doi: 10.1093/ntr/nts103.

[14] Fuoco F., Buonanno G., Stabile L., Vigo P. Influential parameters on particle concentration and size distribution in the mainstream of e-cigarettes. *Environ*. *Pollut*. 2014;184:523–529.

doi: 10.1016/j.envpol.2013.10.010.

[15] Kim H.-J., Shin H.-S. Determination of tobaccospecific nitrosamines in replacement liquids of electronic cigarettes by liquid chromatography–tandem mass spectrometry. *J. Chromatogr. A.* 2013;1291:48–55. doi: 10.1016/j.chroma.2013.03.035.

[16] Cheah N.P., Chong N.W.L., Tan J., Morsed F.A., Yee S.K. Electronic nicotine delivery systems: Regulatory and safety challenges: Singapore perspective. *Tob. Control.* 2012;23:119–125. doi: 10.1136/tobaccocontrol-2012-050483. [17] Etter J.-F., Zäther E., Svensson S. Analysis of refill liquids for electronic cigarettes. *Addiction*. 2013;108:1671–1679.

doi: 10.1111/add.12235.

[18] Kubica P., Wasik A., Kot-Wasik A., Namiesnik J. An evaluation of sucrose as a possible contaminant in eliquids for electronic cigarettes by hydrophilic interaction liquid chromatography-tandem mass spectrometry. *Anal. Bioanal. Chem.* 2014;406:3013–3018. doi: 10.1007/s00216-014-7690-2.

[19] Office on Smoking and Health, National Center for Chronic Disease Prevention and Health Promotion (US) 2016.

[20] Han S., Chen H., Zhang X., Liu T., Fu Y. Levels of Selected Groups of Compounds in Refill Solutions for Electronic Cigarettes. *Nicotine Tob. Res.* 2015;18:708– 714. doi: 10.1093/ntr/ntv189.

[21] Dawkins L., Turner J., Hasna S., Soar K. The electronic-cigarette: Effects on desire to smoke, withdrawal symptoms and cognition. *Addict. Behav.* 2012;37:970–973.

doi: 10.1016/j.addbeh.2012.03.004.

[22] Dawkins L., Turner J., Crowe E. Nicotine derived from the electronic cigarette improves time-based prospective memory in abstinent smokers. *Psychopharmacology*. 2013;227:377–384.

doi: 10.1007/s00213-013-2983-2.

[23] Dicpinigaitis P.V., Lee C.A., Dicpinigaitis A.J., Negassa A. Effect of electronic cigarette use on cough reflex sensitivity. *Chest.* 2015;149:161–165. doi: 10.1378/chest.15-0817.

[24] Palamidas A., Tsikrika S., Katsaounou P.A., Vakali S., Gennimata S.-A., Kaltsakas G., Gratziou C., Koulouris N. Acute effects of short term use of e-cigarettes on airways physiology and respiratory symptoms in smokers with and without airways obstructive diseases and in healthy non smokers. *Tob. Prev. Cessat.* 2017;3:5. doi: 10.18332/tpc/67799.

[25] Pellegrino R.M., Tinghino B., Mangiaracina G., Marani A., Vitali M., Protano C., Osborn J.F., Cattaruzza M.S. Electronic cigarettes: An evaluation of exposure to chemicals and fine particulate matter (PM) *Ann. Ig.* 2012;24:279–288.

[26] Williams M., Villarreal A., Bozhilov K., Lin S., Talbot P. Metal and silicate particles including nanoparticles are present in electronic cigarette cartomizer fluid and aerosol. *PLoS ONE*. 2013;8:e57987. doi: 10.1371/journal.pone.0057987.

[27] Lerner C.A., Sundar I.K., Watson R.M., Elder A., Jones R., Done D., Kurtzman R., Ossip D.J., Robinson R., McIntosh S., et al. Environmental health hazards of ecigarettes and their components: Oxidants and copper in e-cigarette aerosols. *Environ. Pollut.* 2015;198:100–107. doi: 10.1016/j.envpol.2014.12.033.

[28] McAuley T.R., Hopke P., Zhao J., Babaian S. Comparison of the effects of e-cigarette vapor and cigarette smoke on indoor air quality. *Inhal.* 

#### Toxicol. 2012;24:850-857.

doi: 10.3109/08958378.2012.724728.

[29] Goniewicz M.L., Knysak J., Gawron M., Kosmider L., Sobczak A., Kurek J., Prokopowicz A., Jablonska-Czapla M., Rosik-Dulewska C., Havel C., et al. Levels of selected carcinogens and toxicants in vapour from electronic cigarettes. *Tob. Control.* 2014;23:133–139. doi: 10.1136/tobaccocontrol-2012-050859.

[30] Jandíková H., Dušková M., Stárka L. The influence of smoking and cessation on the human reproductive hormonal balance. *Physiol. Res.* 2017;66:S323–S331. doi: 10.33549/physiolres.933724.

[31] Harlev A., Agarwal A., Gunes S.O., Shetty A., du Plessis S.S. Smoking and Male Infertility: An Evidence-Based Review. *World J. Men Health.* 2015;33:143–160. doi: 10.5534/wjmh.2015.33.3.143.

[32] Natali A., Mondaini N., Lombardi G., Del Popolo G., Rizzo M. Heavy smoking is an important risk factor for erectile dysfunction in young men. *Int. J. Impot. Res.* 2005;17:227–230. doi: 10.1038/sj.ijir.3901275.

[33] Cao S., Yin X., Wang Y., Zhou H., Song F., Lu Z. Smoking and Risk of Erectile Dysfunction: Systematic Review of Observational Studies with Meta-Analysis. *PLoS ONE*. 2013;8:e60443. doi: 10.1371/journal.pone.0060443.

[34] Sharma R., Harlev A., Agarwal A., Esteves S.C. Cigarette Smoking and Semen Quality: A New Metaanalysis Examining the Effect of the 2010 World Health Organization Laboratory Methods for the Examination of Human Semen. *Eur. Urol.* 2016;70:635–645. doi: 10.1016/j.eururo.2016.04.010.

[35] Practice Committee of the American Society for Reproductive Medicine. Penzias A., Bendikson K., Butts S., Coutifaris C., Falcone T., Gitlin S., Gracia C., Hansen K., Jindal S., et al. Smoking and infertility: A committee opinion. *Fertil Steril.* 2018;110:611–618.

[36] La Maestra S., De Flora S., Micale T. Effect of cigarette smoke on DNA damage, oxidative stress, and morphological alterations in mouse testis and spermatozoa. *Int. J. Hyg. Environ. Health.* 2015;2018:117–122.

doi: 10.1016/j.ijheh.2014.08.006.

[37] Firns S., Cruzat V., Keane K.N., Joesbury K.A., Lee A.H., Newsholme P., Yovich J.L. The effect of cigarette smoking, alcohol consumption and fruit and vegetable consumption on IVF outcomes: A review and presentation of original data. *Reprod. Biol. Endocrinol.* 2015;13:134. doi: 10.1186/s12958-015-0133-x.

[38] Håkonsen L.B., Ernst A., Ramlau-Hansen C.H. Maternal cigarette smoking during pregnancy and reproductive health in children: A review of epidemiological studies. *Asian J. Androl.* 2014;16:39–49. doi: 10.4103/1008-682X.122351.

[39] Ozbakir B., Tulay P. Does cigarette smoking really have a clinical effect on folliculogenesis and oocyte maturation? *Zygote*. 2020;28:318–321. doi: 10.1017/S0967199420000155.

https://doi.org/10.55544/jrasb.3.1.33

[40] Lyngsø J., Kesmodel U.S., Bay B., Ingerslev H.J., Pisinger C.H., Ramlau-Hansen C.H. Female cigarette smoking and successful fertility treatment: A Danish cohort study. *Acta Obstet. Gynecol. Scand.* 2021;100:58– 66. doi: 10.1111/aogs.13979.

[41] Harte C.B., Meston C.M. Acute Effects of Nicotine on Physiological and Subjective Sexual Arousal in Nonsmoking Men: A Randomized, Double-Blind, Placebo-Controlled Trial. *J. Sex. Med.* 2008;5:110–121. doi: 10.1111/j.1743-6109.2007.00637.x.

[42] El Golli N., Rahali D., Jrad-Lamine A., Dallagi Y., Jallouli M., Bdiri Y., Ba N., Lebret M., Rosa J., El May M., et al. Impact of electronic-cigarette refill liquid on rat testis. *Toxicol. Mech. Methods.* 2016;26:417–424. doi: 10.3109/15376516.2016.1163448.

[43] Rahali D., Jrad-Lamine A., Dallagi Y., Bdiri Y., Ba N., El May M., El Fazaa S., El Golli N. Semen Parameter Alteration, Histological Changes and Role of Oxidative Stress in Adult Rat Epididymis on Exposure to Electronic Cigarette Refill Liquid. *Chin. J. Physiol.* 2018;61:75–84. doi: 10.4077/CJP.2018.BAG521.

[44] Vivarelli F., Canistro D., Cirillo S., Cardenia V., Rodriguez-Estrada M.T., Paolini M. Impairment of testicular function in electronic cigarette (e-cig, e-cigs) exposed rats under low-voltage and nicotinefreeconditions. *Life* Sci. 2019;228:53–65. doi: 10.1016/j.lfs.2019.04.059.

[45] Wawryk-Gawda E., Zarobkiewicz M.K., Chłapek K., Chylinska-Wrzos P., Jodłowska-Jedrych B. Histological changes in the reproductive system of male rats exposed to cigarette smoke or electronic cigarette vapor. *Toxicol. Environ. Chem.* 2019;101:404–419.

doi: 10.1080/02772248.2019.1703989.
[46] Heger A., Sator M., Walch K., Pietrowski D.
Smoking Decreases Endometrial Thickness in IVF/ICSI
Patients. *Geburtshilfe Und Frauenheilkd*. 2018;78:78–82. doi: 10.1055/s-0043-123762.

[47] Orzabal M.R., Lunde-Young E.R., Ramirez J.I., Howe S.Y., Naik V.D., Lee J., Heaps C.L., Threadgill D.W., Ramadoss J. Chronic exposure to e-cig aerosols during early development causes vascular dysfunction and offspring growth deficits. *Transl. Res.* 2019;207:70– 82. doi: 10.1016/j.trsl.2019.01.001.

[48] Szumilas K., Szumilas P., Grzywacz A., Wilk A. The Effects of E-Cigarette Vapor Components on the Morphology and Function of the Male and Female Reproductive Systems: A Systematic Review. *Int. J. Environ. Res. Public Health.* 2020;17:6152. doi: 10.3390/ijerph17176152.

[49] Akar Y., Ahmad N., Khalid M. The effect of cadmium on the bovine *in vitro* oocyte maturation and early embryo development. *Int. J. Veter Sci. Med.* 2018;6:S73–S77.

doi: 10.1016/j.ijvsm.2018.03.001.

[50] Holden L.L., Truong L., Simonich M.T., Tanguay R.L. Assessing the hazard of E-Cigarette flavor mixtures using zebrafish. *Food Chem. Toxicol.* 2020;136:110945. doi: 10.1016/j.fct.2019.110945.

https://doi.org/10.55544/jrasb.3.1.33

[51] Marzec-Wróblewska U., Kamiński P., Lakota P., Szymański M., Wasilow K., Ludwikowski G., Kuligowska-Prusińska M., Odrowąż-Sypniewska G., Stuczyński T., Michałkiewicz J. Zinc and iron concentration and SOD activity in human semen and seminal plasma. *Biol. Trace Elem. Res.* 2011;143:167– 177. doi: 10.1007/s12011-010-8868-x.

[52] Marzec-Wróblewska U., Kamiński P., Łakota P., Szymański M., Wasilow K., Ludwikowski G., Jerzak L., Stuczyński T., Woźniak A., Buciński A. Human Sperm Characteristics with Regard to Cobalt, Chromium, and Lead in Semen and Activity of Catalase in Seminal Plasma. *Biol. Trace Elem. Res.* 2019;188:251–260. doi: 10.1007/s12011-018-1416-9.

[53] Shi X., Chan C.P.S., Man G.K.Y., Chan D.Y.L., Wong M.H., Li T.C. Associations between blood metal/ metalloid concentration and human semen quality and sperm function: A cross-sectional study in Hong Kong. *J. Trace Elem. Med. Biol.* 2021;65:126735. doi: 10.1016/j.jtemb.2021.126735.

[54] Zhang X.F., Gurunathan S., Kim J.H. Effects of silver nanoparticles on neonatal testis development in mice. *Int. J. Nanomed.* 2015;10:6243–6256.

[55] Zhang T., Ru Y.F., Bin Wu B., Dong H., Chen L., Zheng J., Li J., Wang X., Wang Z., Wang X., et al. Effects of low lead exposure on sperm quality and sperm DNA methylation in adult men. *Cell Biosci.* 2021;30:110–150. doi: 10.1186/s13578-021-00665-7.

[56] Chen C., Li B., Huang R., Dong S., Zhou Y., Song J., Zeng X., Zhang X. Involvement of Ca<sup>2+</sup> and ROS signals in nickel-impaired human sperm function. *Ecotoxicol. Environ. Saf.* 2022;231:113181. doi: 10.1016/j.ecoenv.2022.113181.

[57] Zhao L.L., Ru Y.F., Liu M., Tang J.N., Zheng J.F., Wu B., Gu Y.H., Shi H.J. Reproductive effects of cadmium on sperm function and early embryonic development in vitro. *PLoS ONE*. 2017;12:e0186727. doi: 10.1371/journal.pone.0186727.

[58] Roychoudhury S., Nath S., Massanyi P., Stawarz R., Kacaniova M., Kolesarova A. Copper-induced changes in reproductive functions: In vivo and in vitro effects. *Physiol. Res.* 2016;65:11–22. doi: 10.33549/physiolres.933063.

[59] Kong L., Tang M., Zhang T., Wang D., Hu K., Lu W., Wei C., Liang G., Pu Y. Nickel Nanoparticles Exposure and Reproductive Toxicity in Healthy Adult Rats. *Int. J. Mol. Sci.* 2014;15:21253–21269. doi: 10.3390/ijms151121253.

[60] Yiqin C., Yan S., Peiwen W., Yiwei G., Qi W., Qian X., Panglin W., Sunjie Y., Wenxiang W. Copper exposure disrupts ovarian steroidogenesis in human ovarian granulosa cells via the FSHR/CYP19A1 pathway and alters methylation patterns on the SF-1 gene promoter. *Toxicol. Lett.* 2022;356:11–20. doi: 10.1016/j.toxlet.2021.12.002.

[61] Vosoughi S., Khavanin A., Salehnia M., Mahabadi H.A., Shahverdi A., Esmaeili V. Adverse Effects of

Formaldehyde Vapor on Mouse Sperm Parameters and Testicular Tissue. *Int. J. Fertil. Steril.* 2013;6:250–267. [62] Zang Z.-J., Fang Y.-Q., Ji S.-Y., Gao Y., Zhu Y.-Q., Xia T.-T., Jiang M.-H., Zhang Y.-N. Formaldehyde Inhibits Sexual Behavior and Expression of Steroidogenic Enzymes in the Testes of Mice. *J. Sex. Med.* 2017;14:1297–1306.

doi: 10.1016/j.jsxm.2017.09.001.

[63] Wang H.-X., Wang X.-Y., Zhou D.-X., Zheng L.-R., Zhang J., Huo Y.-W., Tian H. Effects of low-dose, long-term formaldehyde exposure on the structure and functions of the ovary in rats. *Toxicol. Ind. Heal.* 2012;29:609–615.

doi: 10.1177/0748233711430983.

[64] Wang H.-X., Li H.-C., Lv M.-Q., Zhou D.-X., Bai L.-Z., Du L.-Z., Xue X., Lin P., Qiu S.-D. Associations between occupation exposure to Formaldehyde and semen quality, a primary study. *Sci. Rep.* 2015;5:15874. doi: 10.1038/srep15874.

[65] Balabanič D., Rupnik M.S., Klemenčič A.K. Negative impact of endocrine-disrupting compounds on human reproductive health. *Reprod. Fertil. Dev.* 2011;23:403–416. doi: 10.1071/RD09300.

[66] Longo V., Forleo A., Ferramosca A., Notari T., Pappalardo S., Siciliano P., Capone S., Montano L. Blood, urine and semen Volatile Organic Compound (VOC) pattern analysis for assessing health environmental impact in highly polluted areas in Italy. *Environ. Pollut.* 2021;286:117410. doi: 10.1016/j.envpol.2021.117410.

[67] Yang P., Wang Y.-X., Chen Y.-J., Sun L., Li J., Liu C., Huang Z., Lu W.-Q., Zeng Q. Urinary Polycyclic Aromatic Hydrocarbon Metabolites and Human Semen Quality in China. *Environ. Sci. Technol.* 2017;51:958–967. doi: 10.1021/acs.est.6b04810.

[68] Khoudja R.Y., Xu Y., Li T., Zhou C.J. Better IVF outcomes following improvements in laboratory air quality. *J. Assist. Reprod. Genet.* 2013;30:69–76. doi: 10.1007/s10815-012-9900-1.

[69] Chen T., Wu M., Dong Y., Kong B., Cai Y., Hei C., Wu K., Zhao C., Chang Q. Effect of e-cigarette refill liquid on follicular development and estrogen secretion in rats. *Tob. Induc. Dis.* 2022;20:36. doi: 10.18332/tid/146958.

101: 10.18332/tid/146958.

[70] Wetendorf M., Randall L.T., Lemma M.T., Hurr S.H., Pawlak J.B., Tarran R., Doerschuk C.M., Caron K.M. E-Cigarette Exposure Delays Implantation and Causes Reduced Weight Gain in Female Offspring Exposed In Utero. *J. Endocr. Soc.* 2019;3:1907–1916. doi: 10.1210/js.2019-00216.

[71] McGrath-Morrow S.A., Hayashi M., Aherrera A., Lopez A., Malinina A., Collaco J.M., Neptune E., Klein J.D., Winickoff J.P., Breysse P., et al. The Effects of Electronic Cigarette Emissions on Systemic Cotinine Levels, Weight and Postnatal Lung Growth in Neonatal Mice. *PLoS* ONE. 2015;10:e0118344. doi: 10.1371/journal.pone.0118344.

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https://doi.org/10.55544/jrasb.3.1.33

[72] Smith D., Aherrera A., Lopez A., Neptune E., Winickoff J.P., Klein J.D., Chen G., Lazarus P., Collaco J.M., McGrath-Morrow S.A. Adult Behavior in Male Mice Exposed to E-Cigarette Nicotine Vapors during Late Prenatal and Early Postnatal Life. *PLoS ONE*. 2015;10:e0137953.

doi: 10.1371/journal.pone.0137953.

[73] Greene R.M., Pisano M.M. Developmental toxicity of e-cigarette aerosols. *Birth Defects Res.* 2019;111:1294–1301. doi: 10.1002/bdr2.1571. [74] Freour T., Masson D., Mirallie S., Jean M., Bach K., Dejoie T., Barriere P. Active smoking compromises IVF outcome and affects ovarian reserve. *Reprod. Biomed. Online.* 2008;16:96–102. doi: 10.1016/S1472-6483(10)60561-5.

[75] Klonoff-Cohen H., Natarajan L., Marrs R., Yee B. Effects of female and male smoking on success rates of IVF and gamete intra-Fallopian transfer. *Hum. Reprod.* 2001;16:1382–1390.

doi: 10.1093/humrep/16.7.1382.