

# Systematic review of the association between oil and natural gas extraction processes and human reproduction

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This systematic review identified 45 original published research articles related to oil and gas extraction activities and human reproductive endpoints. Reproductive outcomes were categorized as [1] birth outcomes associated with maternal exposure, [2] semen quality, fertility, and birth outcomes associated with adult paternal exposure, [3] reproductive cancers, and [4] disruption of human sex steroid hormone receptors. The results indicate there is moderate evidence for an increased risk of preterm birth, miscarriage, birth defects, decreased semen quality, and prostate cancer. The quality of the evidence is low and/or inadequate for stillbirth, sex ratio, and birth outcomes associated with paternal exposure, and testicular cancer, female reproductive tract cancers, and breast cancer, and the evidence is inconsistent for an increased risk of low birth weight; therefore, no conclusions can be drawn for these health effects. There is ample evidence for disruption of the estrogen, androgen, and progesterone receptors by oil and gas chemicals, which provides a mechanistic rationale for how exposure to oil and gas activities may increase the health risks we have outlined. The results from this systematic review suggest there is a negative impact on human reproduction from exposure to oil and gas activities. Many of the 45 studies reviewed identified potential human health effects. Most of these studies focused on conventional oil and gas activities. Few studies have been conducted to evaluate the impact of unconventional oil and gas operations on human health. The impact of unconventional oil and gas activities may be greater than that of conventional activity, given that unconventional activities employ many of the same approaches and use dozens of known endocrine-disrupting chemicals in hydraulic fracturing. (Fertil Steril® 2016;106: 795–819. ©2016 by American Society for Reproductive Medicine.)

**Key Words:** Birth defects, cancer, oil and natural gas, human reproduction, hydraulic fracturing, endocrine disrupting chemicals, environmental pollution, fracking, hormonal activity, prenatal exposure, preterm birth, semen quality

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he world's need for energy is met substantially by natural gas and oil. In February 2016 alone, the United States produced 9 million barrels of oil and 92 billion cubic feet of natural gas per day (1, 2). As of 2014 there were 514,786 producing gas wells, and natural gas has been forecast to be the leading source of energy by 2040 (3). Increased production of oil and gas has been facilitated by the use of unconventional oil and gas extraction, which involves directional drilling and hydraulic fracturing to access previously unreached sources of oil and gas (tight gas, coal bed methane, and shale gas). Unconventional oil extraction is projected to increase from

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Fertility and Sterility® Vol. 106, No. 4, September 15, 2016 0015-0282/\$36.00 Copyright ©2016 American Society for Reproductive Medicine, Published by Elsevier Inc. http://dx.doi.org/10.1016/j.fertnstert.2016.07.1099 35% of the U.S. oil production in 2008 to 50% by the year 2019 (2). With this increased production, there is the potential for increased exposure to chemicals and products from oil and gas extraction, processing, and wastewater, which may increase the risk for adverse human health effects.

Conventional and unconventional oil and natural gas extraction activities, including drilling, hydraulic fracturing, extraction, processing, transportation and disposal of wastewater, lead to many opportunities for environmental contamination with the chemicals used in and produced during these processes. Potential routes of human exposure to these chemicals and products include

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inhalation, dermal, and oral exposure. Chemicals can be volatilized or aerosolized during extraction and by active evaporating pits, flares, surface spills, processing, and transportation (4–6). Oil and gas industry activities can contaminate surface, ground, and drinking water through the drilling process, hydraulic fracturing, failure of well casings, wastewater spills, and structural failure in abandoned wells (5–10). This has led to concerns about health risks to those exposed to the chemicals used and produced in the oil and gas industry either via residential proximity or occupational exposure (11–13).

One health effect of particular concern is the impact on human reproduction. Human reproduction is controlled by hormones in the hypothalamic, pituitary, and gonadal axis. We have shown that many chemicals used in oil and gas extraction processes can disrupt hormone receptors, the endocrine system, and development (5, 14–16). We previously reviewed the literature on a limited number of chemicals associated with unconventional oil and gas extraction and found that many are associated with adverse reproductive outcomes (17). More recently, Elliott et al. (18) performed a systematic evaluation of chemicals used in hydraulic fracturing fluids for reproductive or developmental toxicity in the ReproTox database. Out of 1,021 chemicals identified in hydraulic fracturing fluids, 240 chemicals had toxicity information. Of this subset, 43% were suggestive of reproductive toxicity. Although individual oil and gas chemicals have been linked to reproductive toxicity, few studies have assessed human exposure to individual chemicals. The objective of this review was to systematically evaluate original published research examining potential human reproductive health effects associated with exposure to both conventional and unconventional oil and gas extraction operations.

### MATERIALS AND METHODS Literature Search

Four literature search strategies were used to comprehensively identify published literature on this topic.

#### TABLE 1

## Terms in search 1 and search 2.

Terms in search I a	nu search 2.			
<b>Oil and gas terms</b> Coal bed methane Coal seam gas Coal bed methane	Crude oil Fracking Hydraulic fracturing	Hydrofracking Natural gas Oil and gas	Petroleum Shale Tight oil	Unconventional gas Unconventional oil
Reproduction term Abortion Acrosome Adnexa Adrenarch Androgen Andropause Anostrus Anogenital Anovulation Aspermia Asthenozoospermia	s Epidydymis Estrogen Estrous Estrus Fecundity Fertile Fertility Fertility Fertilization Fetal development Fetal movement Fetal organ	Luteinization Luteolysis Lymphnagiogenesis Maternal Maternal-ge Maternal-fetal exchange Menarche Menopause Menstrual Menstrual Menstruation Metestrus	Parturition Paternal age Penile Penile erection Peris Peripartum Placenta Placenta Placentation Postmenopause Postpartum	Sertoli Sertoli cell-only syndrome Sex Sex characteristics Sex determination Sex differentiation Sexual development Sexual maturation Sperm Sperm capacitation Sperm maturation
Birth Birth rate Breech Bulbourethral glands	Fetal organ maturity Fetal viability Fetal viability Fetal weight	Miscarriage Multiple pregnancy Musculoskeletal development Neurogenesis	Pregnancy Pregnancy loss Pregnancy maintenance Pregnancy outcome	Sperm transport Sperm-ovum interaction Spermatazoa Spermatic cord
Cementogenesis Cervical Cervix Climacteric Coitus Contraception Contraception behavior Corpus luteum	Follicular Follicular Folliculara atresia Gametogenesis Gastrulation Genitalia Germ cell Gestational age	Neurulation Obstetric Odontogenesis Oligospermia Oogenesis Organogenesis Orgasm Ovarian reserve	Pregnancy rate Premenopause Prenatal nutritional physiological phenomena Proestrus Progesterone Prostate Psuedopregnancy Puberty	Spermatogenesis Still birth Subfertility Superfetation Superovulation Term birth Testes Testis
Dentinogenesis Diestrus	Gonads Gravidity	Ovary Oviparity	Puberty Reproduction	Trimesters Uterus
Ectogenesis Ejaculation Ejaculatory ducts Embryonic and fetal development	Infertility Insemination Lactation Lacteal elimination	Oviposition Ovoviviparity Ovulation Ovulation inhibition	Reproductive Reproductive behavior Reproductive physiologic processes Reproductive physiological phenomena	Vagina Vas deferens Vitellogenesis Viviparity
Embryonic development Embryonic induction	Live birth Luteal	Ovum Parity	Scrotum Seminal vesicles	Vulva
Balise. Oil and gas and reproduction. Fertil Steril 2016.				

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