

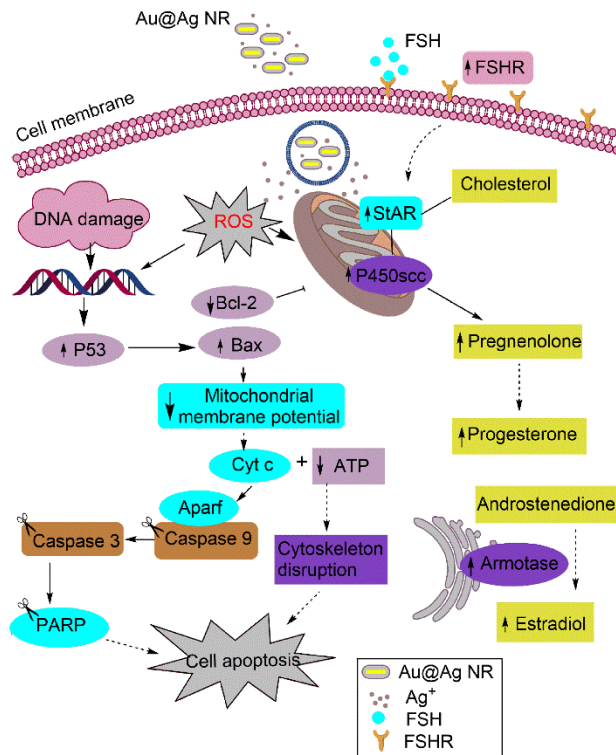
## INTERFERENCE OF STEROIDOGENESIS BY GOLD NANOROD CORE/SILVER SHELL NANOSTRUCTURE: IMPLICATIONS FOR REPRODUCTIVE TOXICITY OF SILVER NANOMATERIALS

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Silver nanomaterials are widely used in personal care products. Recent studies have indicated that these nanomaterials may penetrate the blood-placental barrier and gain access to the ovaries. It is largely unknown how silver nanomaterials influence ovarian physiology and functions such as hormone production. This study examines the *in vitro* toxicology of silver nanomaterials, focusing especially on cytotoxicity and steroidogenesis while exploring their underlying mechanisms. In this study, primary rat granulosa cells were exposed to gold nanorod core/silver shell nanostructures (Au@Ag NRs), which were compared to cells exposed to gold nanorods only. The Au@Ag NRs generated more reactive oxygen species (ROS), reduced mitochondrial membrane potential, and decreased production of adenosine triphosphate. Au@Ag NRs promoted steroidogenesis, including progesterone and estradiol, in a time and dose-dependent manner. Chemical reactivity and transformation of Au@Ag NRs were then studied by electron spin resonance spectroscopy (ESR) and X-ray absorption near edge structure, which identified the generation of free radicals and intracellular silver species. These results suggested that both particle-specific activity and intracellular silver ion release of Au@Ag NR contribute to the toxic response of granulosa cells.



*Figure 1, Schematic representation of the proposed mechanism for cell apoptosis and steroidogenesis interference induced by Au@Ag NRs exposure.*