

# Effect of nanostructured titanium dioxide on photoinduced cancer treatment

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## Abstract

The multivariate condition of cancer disease has been approached by scientific community in many ways. Recent studies focus on individualized treatments, minimizing the undesirable consequences of the conventional treatment methods. There are still aspects of the research field of alternative cancer treatments that remains to be discovered.

TiO<sub>2</sub> has a variety of applications of daily routine and of advanced technology. Due to its biocompatibility, it has also a great number of biomedical applications. It is well established that photo-excited TiO<sub>2</sub> nanoparticles, induce generation of pairs of electrons and holes which react with water and oxygen to yield reactive oxygen species (ROS) which can damage cancer cells. Therefore, TiO<sub>2</sub> is a promising photosensitizer against cancer.

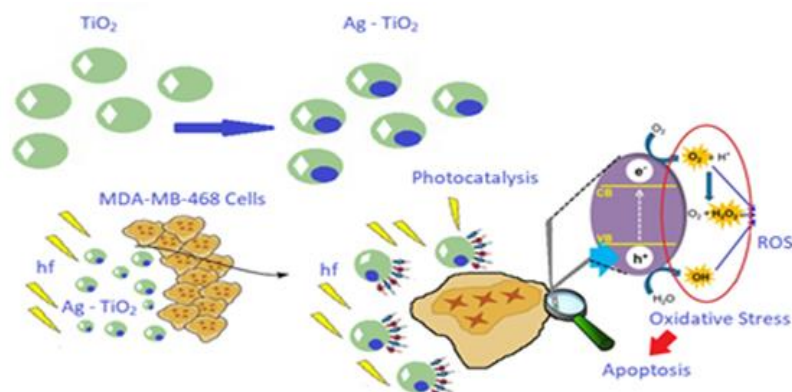
The photo-excitement of TiO<sub>2</sub> can be achieved by UV light with significant anticancer effect, but the aim of this study is the development of TiO<sub>2</sub> nanoparticles with the potential to photo-induce anticancer effect via the mechanism of oxidative stress upon irradiation with visible light. Surface modification by doping with metal ions generally improves TiO<sub>2</sub> photocatalytic activity, as this process can lead to reduction of electron-hole recombination, resulting in efficient separation and stronger photocatalytic reactions. Particularly, silver is an important dopant, which up-regulates TiO<sub>2</sub> biological activity. Thus, Ag-doping of TiO<sub>2</sub> was undergone, followed by detailed characterization (XRD, micro-Raman, SEM). Cultured highly malignant, breast cancer epithelial cells (MDA-MB-468) were irradiated, using visible light, in the presence of Ag-doped TiO<sub>2</sub> aqueous dispersion. Cell viability was estimated, by MTT colorimetric assay. Western blot analysis of protein expression, as well as DNA laddering assay were employed to investigate the existence of cell apoptosis.

We demonstrated that Ag-doped TiO<sub>2</sub> nanoparticles induced apoptosis in cancer cells. The molecular mechanism of TiO<sub>2</sub> nanoparticles cytotoxicity is associated with increased pro-apoptotic Bax expression and caspase-mediated poly (adenosine diphosphate (ADP)-ribose) polymerase (PARP) activation thus resulting in DNA fragmentation and programmed cell death.

Further studies are already in progress, focalizing at the development of visible-light-excited co-doped TiO<sub>2</sub> nanoparticles with silver and nitrogen, for targeted cancer therapy in parallel with series of experiments, focusing on the encapsulation of TiO<sub>2</sub> in polymers, in order to control the release of nanoparticles for drug delivery system development.

## References

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**Figure. Schematic representation of photokilling effect of Ag-doped TiO<sub>2</sub> on breast cancer epithelial cells.** Ag-doped TiO<sub>2</sub> nanoparticles induced apoptosis in highly malignant MDA-MB-468 cancer cells.