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Effects of bisphenol A on metabolism and evidences of a mode of action mediated through endocrine disruption

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**TITLE**

**Effects of Bisphenol A on metabolism and evidences of a mode of action mediated through endocrine disruption**

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**KEYWORDS**

Bisphenol A, BPA, endocrine disruption, obesity, type 2 diabetes, insulin

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**HIGHLIGHT**

- BPA decreases insulin synthesis and secretion, after prenatal or adult exposure
- BPA decreases insulin action, after prenatal, perinatal and adult exposure
- There is good evidence of BPA toxicity on pancreatic  $\beta$  -cells
- Estrogen receptor mechanism ( $\beta/\alpha$  or GPR30 types) may be involved
- Epigenetic mechanisms are also suggested

<sup>1</sup> BAT: brown adipose tissue; BMI: body mass index; bw: body weight; DES: diethylstilbestrol; DM: Diabete mellitus; ED: endocrine disruptor; eGFR: estimated glomerular filtration ; ERR: estrogen related receptor; ERR $\gamma$  or ERRgamma: Estrogen-related receptor gamma; ER $\alpha$  or; ERalpha: estrogen receptor  $\alpha$  ; ER $\beta$  or ERbeta: estrogen receptor beta ; FABP4: fatty acid binding protein 4; GD: gestation day; GLUT4: Glucose transporter type 4; GPER or GPR30: G protein-coupled estrogen receptor or G protein-coupled receptor 30, the membranous form of estrogen receptor; GR: glucocorticoid receptor (NR3C1); GSH:Glutathione ; GSIS: Glucose-Stimulated Insulin Secretion ; GTT: glucose tolerance test; HFD: High Fat Diet; IAPP: Islet Amyloid PolyPeptide ; ipGTT: intraperitoneal glucose tolerance tests; ipITT: intraperitoneal insulin tolerance tests; IR: insulin receptor; IRS-1: insulin receptor substrate 1; IS: insulin sensitivity; MDI: induction medium containing methylisobutylxanthine, dexamethasone, insulin; used in protocols to induce differentiation of 3T3-L1 cells into adipocyte-like cells; MoA: mode of action; MSCs: Mesenchymal Stromal Cells; NEFA: Non-Esterified Fatty Acids ; NHANES: US National Health and Nutrition Examination Survey; NHS: nurses' health study; NHSII: nurses' health study II; PI3K: phosphoinositide 3-kinase; PND: postnatal day; PPAR $\gamma$ : Peroxisome Proliferator Activated Receptor gamma ; PPRE: PPAR $\gamma$ ; Response Element ; ROSI: Rosiglitazone; SOD: superoxide dismutase; TBT: Tributyltin ; TD2M: Type-2 diabetes mellitus ; TDI: tolerable daily intake; TG: triglyceride; TO: T0070907, used as a PPAR $\gamma$  antagonist; WAT: White Adipose Tissue; WB: Western blott

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