Lecture no. 11

EICOSANOIDS

- 1. *Eicosanoids* are a large group of molecules derived from **20 carbon** polyunsaturated fattyacids.
- 2. The principal groups of hormones of this class are **prostaglandins** (PG), prostacyclins, leukotrienes (LT), thromboxanes (TX) and lipoxins (LX).
- 3. Arachidonic acid is the most abundant precursor for these hormones.
- 4. Stores of arachadonic acid are present in membrane lipids and released through the action of various lipases.
- 5. The specific eicosanoids synthesized by a particular cell are dictated by the group of processing enzymes expressed in that cell.
- 6. Phospholipase A₂ releases arachidonic acid from membrane phospholipids.
- 7. The released arachidonic acid is converted to prostaglandin H_2 (PGH₂) by cyclic oxygenase 1 (Cox 1) and cyclic oxygenase 2 (Cox 2) enzymes.
- 8. PGH₂is converted to prostacyclins, thromboxanes and prostaglandins by isomerase enzymes present in tissues.
- 9. ThromboxaneA₂ is synthesized by platelets and promotes vasoconstriction and platelet aggregation.
- 10.Prostacyclinis produced in the lungs and vascular endothelium, inhibits platelet aggregation and produces vasodilatation.
- 11.Leukotrienes mediate allergic response and inflammation.

Prostaglandins(PG)

1. Von Euler in 1934 isolated and named a substance isolated from seminal fluid as prostaglandins.

- 2. Unlike the other hormones, prostaglandins are not localized in any particular tissue.
- 3. Most prostaglandins act locally at the site of their production on a cell interaction—paracrine effect and also act by autocrine mechanism.
- 4. Prostaglandins have varied functions particularly important in female reproduction, parturition, cardiovascular system.
- 5. They play a prominent role in causing pain, inflammation and fever.
- 6. PGs may be considered to regulate several physiological functions such as contraction of smooth muscles in reproductive organs, and functions in erection, ejaculation, sperm transport, ovulation and also luteolysis.
- 7. They also regulate blood pressure, lipolysis, gastric secretion and blood clotting.
- 8. High levels of estrogen promote oxytocin receptors in the endometrium stimulatePGF2 alpha synthesis and its secretion from the endometrium.
- 9. PGF₂ alpha potentiates the contractions of the uterus by estrogen during parturition.
- 10. In females PGF2 α is involved in the follicular rupture and ovulation, also aids sperm transport in males. From endometrium PGF passes directly through the walls of utero-ovarian vein into artery by counter current mechanism and causes vasoconstriction of the blood vessels supplying the CL, the hypoxia in CL leading to *luteolysis*.