### **DEVELOPMENTAL BIOLOGY** TOPIC : Wnt AND TGF BETA FAMILIES

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### Wnt and TGF beta families

#### PARACRINE FACTORS

# Wnt Family

• The Wnt family The Wnts constitute a family of cysteine rich glycoproteins.

• There are at least 15 members of this family in vertebrates.

 Their name comes from fusing the name of the Drosophila segment polarity gene wingless with the name of one of its vertebrate homologues, integrated

- While Sonic hedgehog is important in patterning the ventral portion of the somites (causing the cells to become cartilage),
- Wnt1 appears to be active in inducing the dorsal cells of the somites to become muscle.

 Wnt proteins also are critical in establishing the polarity of insect and vertebrate limbs, and they are used in several steps of urogenital system development

## **TGF-**β superfamily

- There are over 30 structurally related members of the TGF-b superfamily, and they regulate some of the most important interactions in development.
- The proteins encoded by TGF-β superfamily genes are processed such that the carboxyterminal region contains the mature peptide.

 These peptides are dimerized into homodimers (with themselves) or heterodimers (with other TGF-β peptides) and are secreted from the cell.  The TGF-β superfamily includes the TGF-β family, the activin family, the bone morphogenetic proteins (BMPs), the Vg1 family, and other proteins.

# TGF-β family members

 TGF-β1, 2, 3, and 5 are important in regulating the formation of the extracellular matrix between cells and for regulating cell division (both positively and negatively).  TGF-β1 increases the amount of extracellular matrix epithelial cells make (both by stimulating collagen and fibronectin synthesis and by inhibiting matrix degradation).  TGF-βs may be critical in controlling where and when epithelia can branch to form the ducts of kidneys, lungs, and salivary glands.  The effects of the individual TGF-β family members are difficult to sort out, because members of the TGF-β family appear to function similarly and can compensate for losses of the others when expressed together.  Moreover, targeted deletions of the Tgf-β1 gene in mice are difficult to interpret, since the mother can supply this factor through the placenta and milk.  The members of the BMP family were originally discovered by their ability to induce bone formation; hence, they are the bone morphogenetic proteins.  Bone formation, however, is only one of their many functions, and they have been found to regulate cell division, apoptosis (programmed cell death), cell migration, and differentiation.  BMPs can be distinguished from other members of the TGF-β superfamily by their having seven, rather than nine, conserved cysteines in the mature polypeptide  The BMPs include proteins such as Nodal (responsible for left-right axis formation) and BMP4 (important in neural tube polarity, eye development, and cell death.

- As it turns out, BMP1 is not a member of the family; it is a protease.)
- The Drosophila Decapentaplegic protein is homologous to the vertebrate BMP4, and human BMP4 can replace the Drosophila homologue, rescuing those flies deficient in Dpp