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Intrinsic Control of the Follicle Cycle

Produced for the CRC for Premium Quality Wool undergraduate program by;
Dr. Graham Cam, CSIRO Animal Production.



Chalone Hypothesis

- The hair cycle is controlled by a locally active inhibitor that accumulates during anagen, causing the follicles to regress into catagen when a critical level of the inhibitor is reached.
 - this theory covers the autocrine and paracrine regulation of growth and differentiation

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Growth factors present during the follicle cycle and in adult sheep

- In the follicle cycle
 - Most of the TGF family do not change during the cycle
 - TGF β -3 and FGF-2 decrease during the cycle
 - EGF varied and displayed no obvious pattern

But

 - doses of EGF cause follicles to enter telogen
- In the adult sheep
 - EGF and FGF-2 are present in ORS
 - the dermal papilla does not contain EGF or FGF-2

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Molecules involved in the follicle cycle, initiation and development

- **BMP-2 & 4**
 - inhibits proliferation of bulb cells
- **KGF/FGF-7**
 - present in developing follicles and the papilla
 - less follicles and abnormal follicles develop if the FGF-7 receptor is altered
 - administration of FGF-7 increases fibre density
- **TGF- α**
 - deletion of TGF- α affects the shape of the fibres and follicles
 - present in IRS above the bulb
 - EGF receptor (EGFR; which TGF- α can use) is present in ORS
 - hair growth may be controlled by interactions with TGF- α and EGFR

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Future Directions

- Follicle culture
- *In-vivo* test system
- Mapping of genes
- Strategies for gene identification which involves identifying genes that:
 - are known to be critical to fibre production, such as the keratins and cell proliferation,
 - have been shown to affect skin and/or follicle function in other species,
 - have been implicated in developmental systems with similarities to the development of the wool or hair follicle.

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Summary

- There are many molecules that have been implicated in follicle initiation, development and the follicle cycle.
- It is likely that some molecules perform similar functions (e.g. multigene families, EGF/TGF α), so if one is missing, the follicle may still function.
- It appears that gene cascades can be recycled throughout development for the construction of different appendages (e.g. limbs and follicles)
- The interaction between molecules may modulate follicle function.

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