The “Leuven” or entheseal stress hypothesis

- Restoration of homeostasis
- Enthesal stress
  - Acute inflammation
    - TNF
    - BMP
    - WNT
  - Triggering of progenitor cells
    - DISH
    - FOP
  - New bone formation and ankylosis
- Chronic inflammation

- HLA-B27
- Infection
- ERAP1
- IL23R
- IL1
- IBD