# Material Forces in the Context of Biological Tissue Remodelling

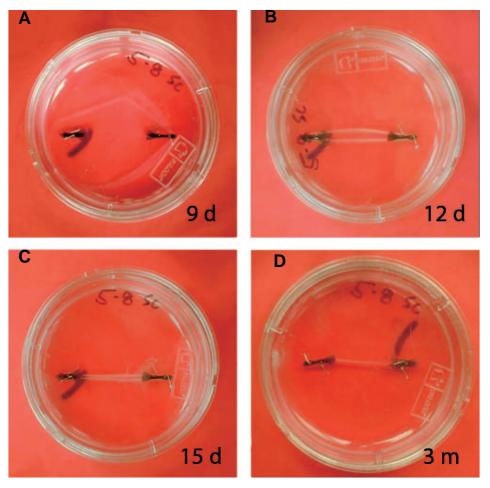
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#### **Growth and Remodelling**

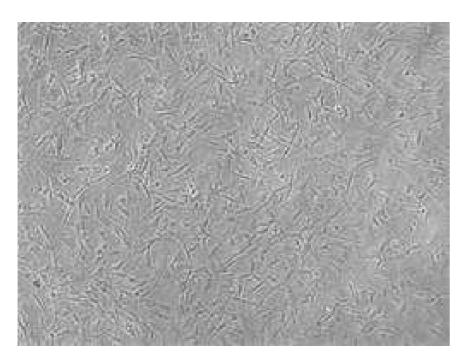
- Growth is a change in density due to mass transport (Epstein & Maugin [2000], Tao et al. [2001], Taber & Humphrey [2001], Humphrey & Rajagopal [2002], Lubarda & Hoger [2002], Kuhl & Steinmann [2002], KG et al. [2003])
  - Tissue is open with respect to mass
  - Multiple species, treated by mixture theory
- Remodelling is an evolution of the microstructure (Taber & Humphrey [2001], Ambrosi & Mollica [2002], Humphrey & Rajagopal [2002])
  - Local reconfiguration of material: self-assembly
  - Evolution of "reference" configuration: remodelled configuration

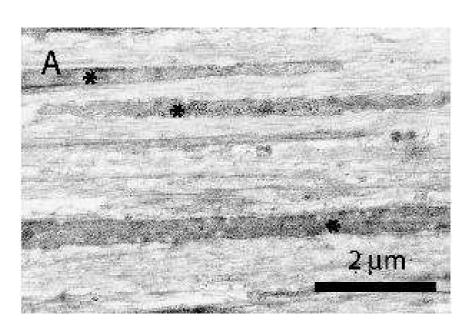
#### **Growth of tendon constructs**



Calve et al. 2003

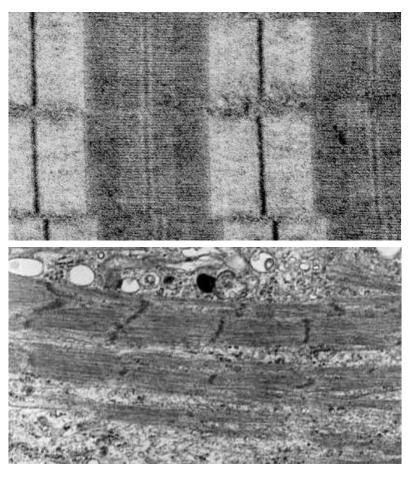
#### Remodelling of collagen during growth





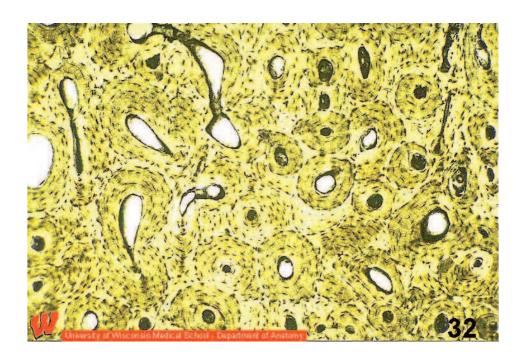
Calve et al. 2003

#### Remodelling during growth



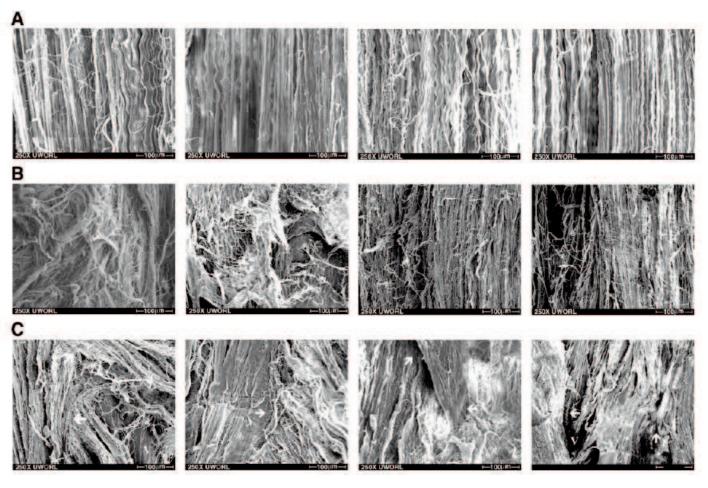
Hirsch et al. 1998

#### Remodelling of bone



- University of Wisconsin, Dept. of Anatomy
- The tissue reconfigures by changing its microstructure when stressed (Wolff [1892])

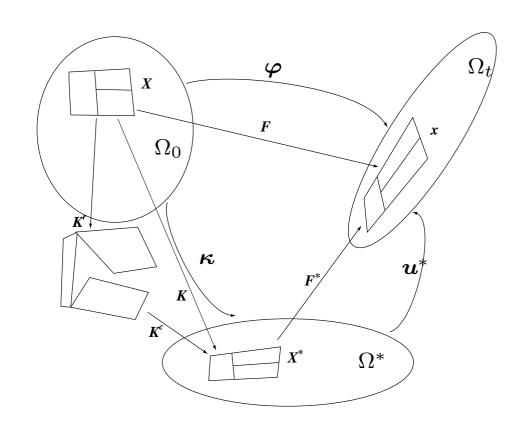
#### Remodelling of collagen due to load while healing



Provenzano et al. 2003

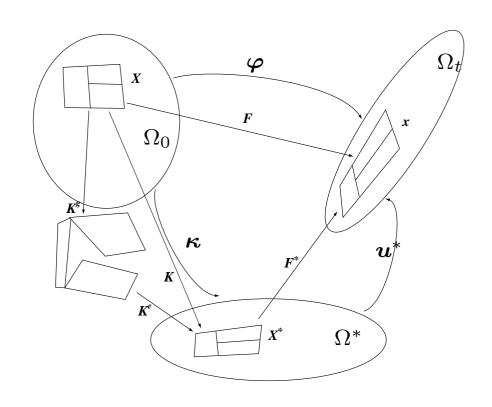
- Remodelling is the reconfiguration of the material
  - Stress-driven
  - "Preferred" configuration that varies pointwise and is in general incompatible. A further configurational change can occur, resulting in a compatible configuration.
- Biological tissue is capable of changes in configuration by motion of particles relative to ambient material
  - Motion in material space/Configurational change

### **Continuum Field Formulation**



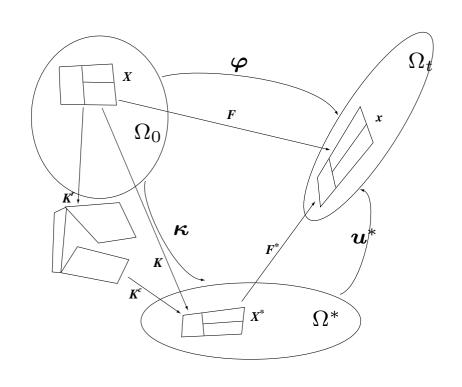
•  $K^{r}$  is given.  $\kappa(X, t) = ?$  (motion in material space)

### **Continuum Field Formulation**



 $m{\mathscr{E}}$  is a kinematic "growth" tensor,  $m{K}^{\mathrm{e}}$  and  $m{F}^*$  are elastic deformation gradients—internal stress problem

### **A Variational Method**



$$\Pi[\boldsymbol{u}^*, \boldsymbol{\kappa}] := \int_{\Omega^*} \hat{\psi}^*(\boldsymbol{F}^*, \boldsymbol{K}^c, \boldsymbol{X}^*) dV^* - \int_{\Omega^*} \boldsymbol{f}^* \cdot (\boldsymbol{u}^* + \boldsymbol{\kappa}) dV^* - \int_{\partial \Omega^*} \boldsymbol{t}^* \cdot (\boldsymbol{u}^* + \boldsymbol{\kappa}) dA^*$$

### **A Variational Method**

- Variation in spatial position:  $m{u}_{arepsilon}^* = m{u}^* + arepsilon \delta m{u}^*$
- ullet Equilibrium with respect to  $u^*$ :

$$\frac{\mathrm{d}}{\mathrm{d}\varepsilon} \Pi[\boldsymbol{u}_{\varepsilon}^*, \boldsymbol{\kappa}] \Big|_{\varepsilon=0} = 0$$

Euler-Lagrange equations:

$$\operatorname{Div}^* \boldsymbol{P}^* + \boldsymbol{f}^* = \boldsymbol{0}, \text{ in } \Omega^*; \quad \boldsymbol{P}^* \boldsymbol{N}^* = \boldsymbol{t}^* \text{ on } \partial \Omega^*; \text{ where } \boldsymbol{P}^* := \frac{\partial \psi^*}{\partial \boldsymbol{F}^*}$$

• Quasistatic balance of linear momentum in remodelled configuration,  $\Omega^*$ 

### **A Variational Method**

Equilibrium with respect to material motion:

$$\kappa_{\varepsilon} = \kappa + \varepsilon \delta \kappa$$

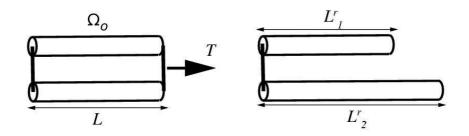
$$\frac{\mathrm{d}}{\mathrm{d}\varepsilon}\Pi[\boldsymbol{u}^*,\boldsymbol{\kappa}_{\varepsilon}]\Big|_{\varepsilon=0}=0$$

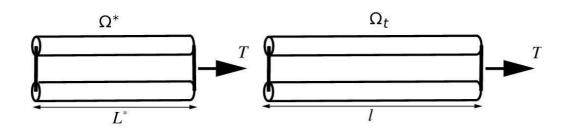
Euler-Lagrange equations:

$$-\mathrm{Div}^*(\psi^* \mathbf{1} - \boldsymbol{F}^{*\mathrm{T}} \boldsymbol{P}^* + \boldsymbol{\Sigma}^*) + \frac{\partial \psi^*}{\partial \boldsymbol{X}^*} = \mathbf{0} \text{ in } \boldsymbol{\Omega}^*,$$

$$-\left(\psi^* \mathbf{1} - \boldsymbol{F}^{*\mathrm{T}} \boldsymbol{P}^* + \boldsymbol{\Sigma}^*\right) \boldsymbol{N}^* = \mathbf{0} \text{ on } \partial \Omega^*$$
where  $\boldsymbol{\Sigma}^* := \frac{\partial \psi^*}{\partial \boldsymbol{K}^c} \boldsymbol{K}^{c\mathrm{T}}$ 

• Eshelby stress:  $\psi^* \mathbf{1} - {m F^*}^{\mathrm T} {m P}^*$ ; configurational stress:  ${m \Sigma}^*$ 

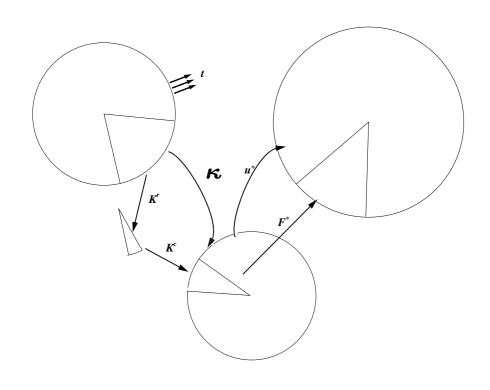




$$\kappa = L^* - L, \quad u^* = l - L^*$$

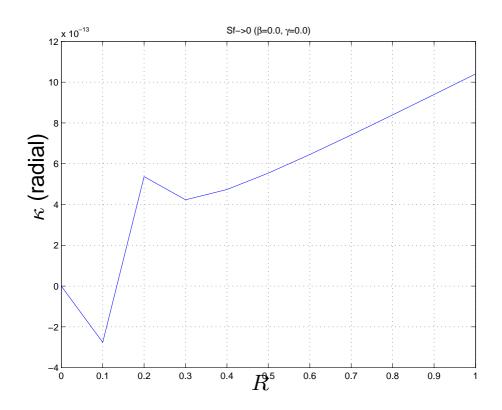
$$\Pi[u^*, \kappa] = \frac{1}{2} k^* (\kappa + L - L_1^{\mathrm{r}})^2 + \frac{1}{2} k^* (\kappa + L - L_2^{\mathrm{r}})^2 + 2 \cdot \frac{1}{2} k u^{*2} - T(u^* + \kappa)$$

$$\frac{\partial \Pi}{\partial u^*} = 0 \quad \Rightarrow \ 2ku^* = T; \qquad \frac{\partial \Pi}{\partial \kappa} = 0 \quad \Rightarrow \ \kappa = \frac{k}{k^*} u^* - \left(L - \frac{L_1^{\mathrm{r}} + L_2^{\mathrm{r}}}{2}\right)$$

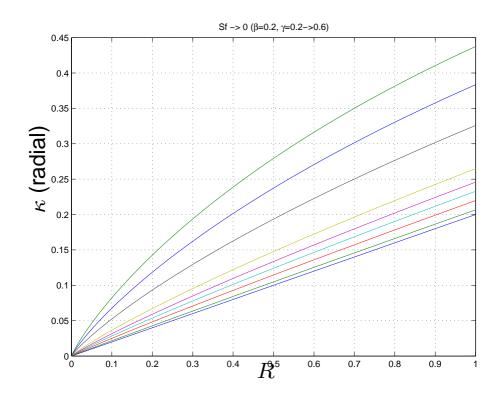


$$oldsymbol{K}^{ ext{r}} = \left[egin{array}{ccc} 1+eta & 0 & 0 \ 0 & 1+\gamma & 0 \ 0 & 0 & 1+\gamma \end{array}
ight], \quad oldsymbol{t}^* = \delta oldsymbol{e}_R$$

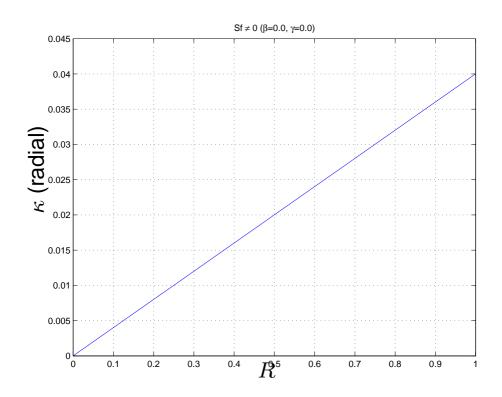
•  $\hat{\psi}^*(\pmb{F}^*, \pmb{K}^c, \pmb{X}^*) = \hat{\psi}_1^*(\pmb{F}^*) + \hat{\psi}_2^*(\pmb{K}^c)$ , (compressible neo-Hookean)



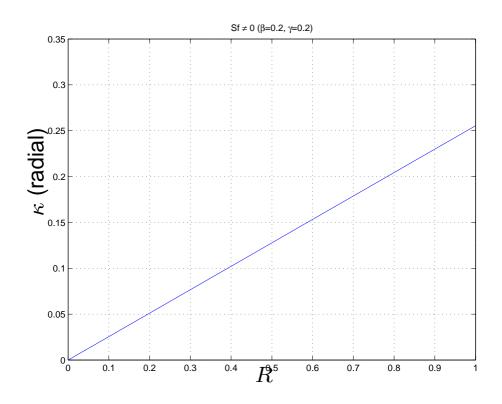
$$m{K}^{
m r} = \left[ egin{array}{ccc} 1 & 0 & 0 \ 0 & 1 & 0 \ 0 & 0 & 1 \end{array} 
ight], \quad m{t}^* = m{0} \; {
m Pa}$$



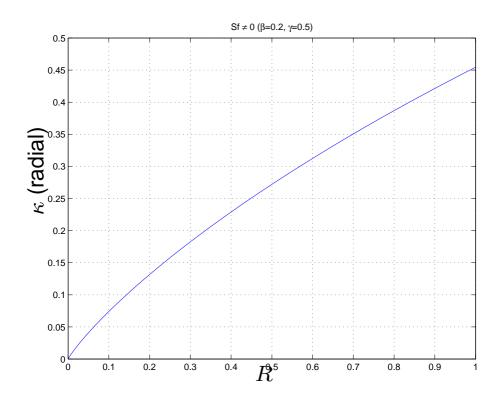
$$m{K}^{
m r} = \left[ egin{array}{cccc} 1 + eta & 0 & 0 \\ 0 & 1 + \gamma & 0 \\ 0 & 0 & 1 + \gamma \end{array} 
ight], eta = 0.2, \; \gamma = 0.2 - 0.6; \quad m{t}^* = m{0} \; {
m Pa}$$



$$m{K}^{\mathrm{r}} = \left[ egin{array}{ccc} 1 & 0 & 0 \ 0 & 1 & 0 \ 0 & 0 & 1 \end{array} 
ight], \quad m{t}^* pprox 10^9 m{e}_R \; \mathrm{Pa}$$



$$m{K}^{
m r} = \left[ egin{array}{ccc} 1.2 & 0 & 0 \ 0 & 1.2 & 0 \ 0 & 0 & 1.2 \end{array} 
ight], \quad m{t}^* pprox 10^9 m{e}_R \; {
m Pa}$$



$$m{K}^{
m r} = \left[ egin{array}{ccc} 1.2 & 0 & 0 \ 0 & 1.5 & 0 \ 0 & 0 & 1.5 \end{array} 
ight], \quad m{t}^* pprox 10^9 m{e}_R \; {
m Pa}$$

### Remarks

- Remodelling is coupled with growth—separate treatment for conceptual clarity
- The remodelled configuration,  $\kappa$  depends upon  $\hat{\psi}^*(\bullet, \mathbf{K}^c, \bullet)$
- Remodelled configuration is assumed to be an equilibrium state
  - Perturb conditions—new equilibrium
- Self-assembly processes in materials are similarly described by minimizing the Gibbs free energy of the systems with respect to the configuration