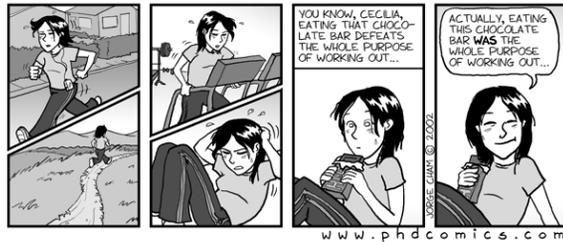


10 - basics constitutive equations - volume growth - growing tumors



10 - basic constitutive equations

1

balance equations

balance equations $['bæl.əns\ r'kwel.ɜəns]$ of mass, momentum, angular momentum and energy, supplemented with an entropy inequality constitute the set of conservation laws. the law of **conservation of mass**/matter states that the **mass of a closed system** of substances will remain **constant**, regardless of the processes acting inside the system. the principle of conservation of momentum states that the total momentum of a closed system of objects is constant.



balance equations

3

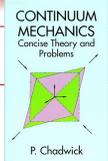
| day | date | topic |
|-----|--------|--|
| tue | jan 07 | motivation - everything grows! |
| thu | jan 09 | basics maths - notation and tensors |
| tue | jan 14 | project example - growing skin |
| thu | jan 16 | kinematics - growing brains |
| tue | jan 21 | basic kinematics - large deformation and growth |
| thu | jan 23 | kinematics - growing hearts |
| tue | jan 28 | kinematics - growing leaflets |
| thu | jan 30 | basic balance equations - closed and open systems |
| tue | feb 04 | basic constitutive equations - growing muscle |
| thu | feb 06 | basic constitutive equations - growing tumors |
| tue | feb 11 | volume growth - finite elements for growth - theory |
| thu | feb 13 | volume growth - finite elements for growth - matlab |
| tue | feb 18 | basic constitutive equations - growing bones |
| thu | feb 20 | density growth - finite elements for growth |
| tue | feb 25 | density growth - growing bones |
| thu | feb 27 | everything grows! - midterm summary |
| tue | mar 04 | midterm |
| thu | mar 06 | remodeling - remodeling arteries and tendons |
| tue | mar 11 | class project - discussion, presentation, evaluation |
| thu | mar 13 | class project - discussion, presentation, evaluation |
| thu | mar 14 | written part of final projects due |

where are we???

balance equations

balance equations $['bæl.əns\ r'kwel.ɜəns]$ of mass, linear momentum, angular momentum and energy **apply to all material bodies**. each one gives rise to a field equation, holding on the configurations of a body in a sufficiently smooth motion and a jump condition on surfaces of discontinuity. like position, time and body, the concepts of mass, force, heating and internal energy which enter into the formulation of the balance equations are regarded as having primitive status in continuum mechanics.

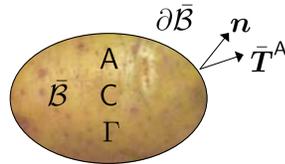
chadwick "continuum mechanics" [1976]



balance equations

4

generic balance equation



general format

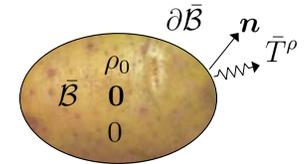
- A ... balance quantity
- B** ... flux $\mathbf{B} \cdot \mathbf{n} = \bar{T}^A$
- C ... source
- Γ ... production

$$D_t A = \text{Div}(\mathbf{B}) + C + \Gamma$$

balance equations - closed systems

5

balance of mass



balance of mass

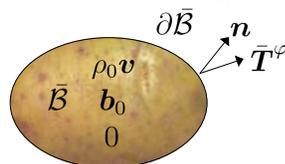
- ρ_0 ... density
- 0** ... no mass flux $\bar{T}^\rho = 0$
- 0 ... no mass source
- 0 ... no mass production

$$\text{continuity equation } D_t \rho_0 = 0$$

balance equations - closed systems

6

balance of (linear) momentum



balance of momentum

- $\rho_0 \mathbf{v}$... linear momentum density
- P** ... momentum flux - stress $\mathbf{P} \cdot \mathbf{n} = \bar{T}^\rho$
- \mathbf{b}_0 ... momentum source - force
- 0 ... no momentum production

$$\text{equilibrium equation } D_t(\rho_0 \mathbf{v}) = \text{Div}(\mathbf{P}) + \mathbf{b}_0$$

balance equations - closed systems

7

compare



NEWTON'S
THREE LAWS OF
GRADUATION

First published in 1679, Isaac Newton's "*Procrastinare Unmaturalis Principia Mathematica*" is often considered one of the most important single works in the history of science. Its Second Law is the most powerful of the three, allowing mathematical calculation of the duration of a doctoral degree.

SECOND LAW

"The age, \mathbf{a} , of a doctoral process is directly proportional to the flexibility, \mathbf{f} , given by the advisor and inversely proportional to the student's motivation, \mathbf{m} "

Mathematically, this postulate translates to:

$$age_{\text{PhD}} = \frac{\text{flexibility}}{\text{motivation}}$$

$$\mathbf{a} = \mathbf{F} / \mathbf{m}$$

$$\therefore \mathbf{F} = \mathbf{m} \mathbf{a}$$

This Law is a quantitative description of the effect of the forces experienced by a grad student. A highly motivated student may still remain in grad school given enough flexibility. As motivation goes to zero, the duration of the PhD goes to infinity.

PH.D. STANFORD.EDU
JORGE CHAM @THE STANFORD DAILY

$$D_t(\rho_0 \mathbf{v}) = \text{Div}(\mathbf{P}) + \mathbf{b}_0 \quad \text{mass point} \quad m D_t \mathbf{v} = m \mathbf{a} = \mathbf{F}$$

balance equations - closed systems

8

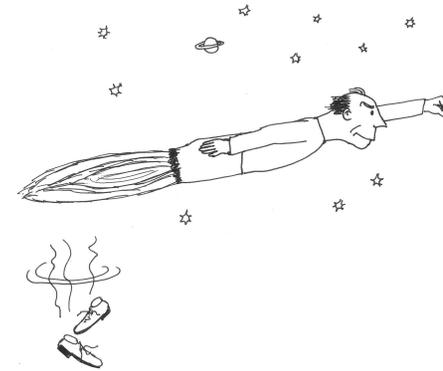
thermodynamic systems

open system ['ou.pən 'sɪs.təm] thermodynamic system which is allowed to exchange mechanical work, heat and mass, typically $\mathbf{P} = \mathbf{P}(\nabla\varphi, \dots)$, $\mathbf{Q} = \mathbf{Q}(\nabla\theta, \dots)$ and $\mathbf{R} = \mathbf{R}(\nabla\rho, \dots)$ with its environment. enclosed by a deformable, diathermal, permeable membrane. characterized through its state of deformation φ , temperature θ and density ρ .

balance equations - open systems

9

open system thermodynamics



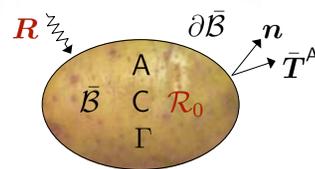
"...thermodynamics recognizes no special role of the biological..."

bridgman, 'the nature of thermodynamics', [1941]

balance equations - open systems

10

balance equations of growth



general format

A ... balance quantity
 B ... flux $\mathbf{B} \cdot \mathbf{n} = \bar{\mathbf{T}}^A$
 C ... source
 Γ ... production

$$D_t(\rho_0 A) = \text{Div}(\mathbf{B} + \mathbf{A} \otimes \mathbf{R}) + [\mathbf{C} + \mathbf{A} \mathbf{R}_0 - \nabla_x \mathbf{A} \cdot \mathbf{R} + \Gamma]$$

balance equations - open systems

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balance equations of growth

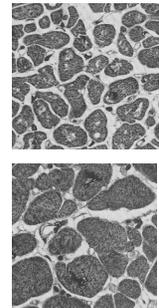
$$D_t \rho_0 = \text{Div}(\mathbf{R}) + \mathcal{R}_0$$

mass flux \mathbf{R}

- cell movement (migration)

mass source \mathcal{R}_0

- cell growth (proliferation)
- cell division (hyperplasia)
- cell enlargement (hypertrophy)



biological equilibrium

cowin & hegedus [1976], beaupré, orr & carter [1990], harrigan & hamilton [1992], jacobs, levenston, beaupré, simo & carter [1995], huiskes [2000], carter & beaupré [2001]

balance equations - open systems

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balance of (linear) momentum

- volume specific version

$$D_t(\rho_0 \mathbf{v}) = \text{Div}(\mathbf{P} + \mathbf{v} \otimes \mathbf{R}) + [\mathbf{b}_0 + \mathbf{v} \mathcal{R}_0 - \nabla_X \mathbf{v} \cdot \mathbf{R}]$$

- subtract weighted balance of mass

$$\mathbf{v} D_t \rho_0 = \text{Div}(\mathbf{v} \otimes \mathbf{R}) + \mathbf{v} \mathcal{R}_0 - \nabla_X \mathbf{v} \cdot \mathbf{R}$$

- mass specific version

$$\rho_0 D_t \mathbf{v} = \text{Div}(\mathbf{P}) + \mathbf{b}_0$$

mechanical equilibrium

balance equations - open systems

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example of open systems - rocket propulsion



balance of mass

$$D_t m = \mathcal{R} \quad \text{with} \quad \mathcal{R} \leq 0 \quad \text{ejection}$$

balance of momentum - volume specific

$$D_t[m\mathbf{v}] = m D_t \mathbf{v} + D_t m \mathbf{v} = \mathbf{f} + \mathcal{R} \mathbf{v}$$

balance of momentum - mass specific

$$m D_t \mathbf{v} = \mathbf{f} \quad \text{with} \quad \mathbf{f} = \mathbf{f}^{\text{closed}} + \mathbf{f}^{\text{open}}$$

balance of momentum - rocket head-ejection

$$D_t[m\mathbf{v}] - \mathcal{R} \bar{\mathbf{v}} = \mathbf{f}^{\text{closed}}$$

propulsive force

$$\mathbf{f}^{\text{open}} = [\bar{\mathbf{v}} - \mathbf{v}] \mathcal{R} \quad \text{velocity of ejection } \bar{\mathbf{v}}$$

example - rocket propulsion

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example of open systems - rocket propulsion



a saturn v rocket like the one that took men to the moon has a **mass of 2.500.000 kg** at liftoff. it goes straight up vertically and burns fuel at a uniform **rate of 16.000 kg/s** for a duration of **2 minutes**. the exhaust speed of gas from the saturn **v is 3.0 km/s**

what is the speed of the rocket immediately after the combustion ceases? you should include the effect of gravity near the surface of the earth, but you can neglect air resistance.

plot the **burnout velocity as a function of time** over the range of 0 to 120 seconds to see the increase in speed of the rocket with time.

example - rocket propulsion

15

example of open systems - rocket propulsion



$$m D_t \mathbf{v} = \mathbf{f}^{\text{closed}} + \mathbf{f}^{\text{open}}$$

$$\text{with } \mathbf{f}^{\text{closed}} = -m \mathbf{g} \quad \text{gravity}$$

$$\mathbf{f}^{\text{open}} = [\bar{\mathbf{v}} - \mathbf{v}] \mathcal{R} = w D_t m$$

$$m D_t \mathbf{v} = -m \mathbf{g} - D_t m w \quad || : m$$

$$D_t \mathbf{v} = -\mathbf{g} - \frac{1}{m} D_t m w$$

integration

$$v(t) = -g t - w \int_{m(0)}^{m(t)} \frac{1}{m} dm$$

$$m(t) = m(0) + \mathcal{R} t$$

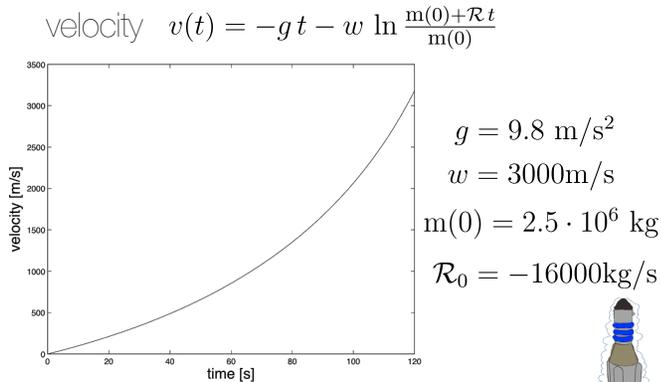
velocity

$$v(t) = -g t - w \ln \frac{m(0) + \mathcal{R} t}{m(0)}$$

example - rocket propulsion

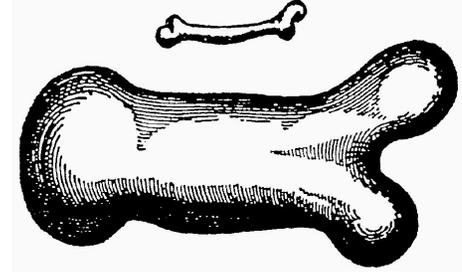
16

example of open systems - rocket propulsion



velocity $v(t = 120\text{s}) = 3207.1 \text{ m/s}$

example of open systems - the galileo giant



„...dal che e manifesto, che chi volesse mantener in un vastissimo gigante le proporzioni, che hanno le membra in un huomo ordinario, bisognerebbe o trouar materia molto piu dura, e resistente per formame l'ossa o vero ammettere, che la robustezza sua fusse a proporzione assai piu fiacca, che negli huomini de statura mediocre; altrimenti crescendogli a smisurata altezza si vedrebbero dal proprio peso opprimere, e cadere...”

galileo, "discorsi e dimostrazioni matematiche", [1638]

example of open systems - the galileo giant

the tallest man in medical history is robert pershing wadlow. he was born at alton, illinois, on february 22, 1918. he was **2.72m** / 8ft 11.1", tall.

his weight was **222.71kg**. his shoe size was 47cm / 18.5", and his hands measured 32.4cm / 12.75". his arm span was 2.88m / 9 ft 5.75", and his peak daily food consumption was 8000 calories.



guinness world records [2010]

example of open systems - the galileo giant

consider and compare the two cases:

closed system. calculate the vertical displacement and the energy for a giant with a **constant bone mineral density along the height**

open system. calculate the vertical displacement and the bone mineral density for a giant with a **constant energy along the height**



guinness world records [2010]

example of closed systems - the galileo giant

- balance equation closed systems $\rho_0 D_t v = \text{Div}(P) + b_0$
- neo hookean free energy $\psi_0 = \frac{1}{4} E_0 [F^2 - 1 - 2 \ln(F)]$
- stress from dissipation inequality $P = D_F \psi_0 = \frac{1}{4} E_0 [2F - 2\frac{1}{F}]$
- quasi-static case $\rho_0 D_t v = 0$
- constant gravity load $b_0 = \text{const along the height } h$
- from balance eqn linear stress $P = [X - h] b_0$
...linear along the height h
- closed system, constant density $\rho = \rho_0^* \quad [\rho - \rho_0^*] / \rho_0^* = 0$

example - the galileo giant

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example of open systems - the galileo giant

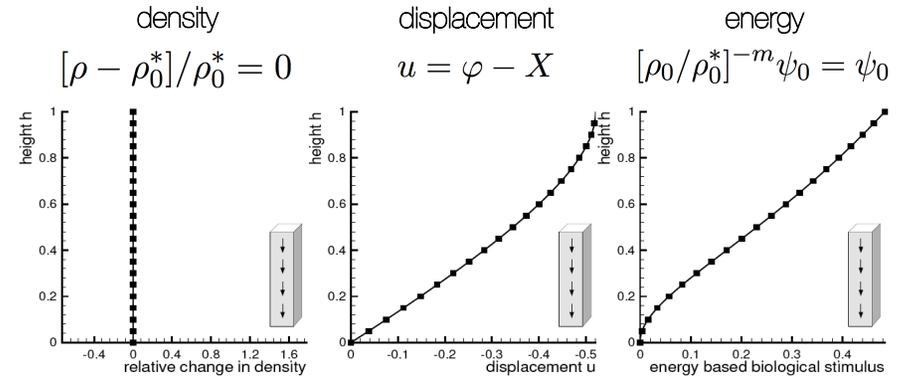


Figure 7.1: One-dimensional model problem - Closed system - Homogeneous density

example - the galileo giant

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example of open systems - the galileo giant

- balance equations open systems $D_t \rho_0 = \text{Div}(R) + \mathcal{R}_0$
 $\rho_0 D_t v = \text{Div}(P) + b_0$
- free energy $\psi_0 = [\rho_0 / \rho_0^*]^n \psi_0^{\text{neo}}$ $\psi_0^{\text{neo}} = \frac{1}{4} E_0 [F^2 - 1 - 2 \ln(F)]$
- stress from dissipation inequality $P = D_F \psi_0 = \frac{1}{4} E_0 [2F - 2\frac{1}{F}]$
- quasi-static case $\rho_0 D_t v = 0$
- constant gravity load $b_0 = \text{const along the height } h$
- from balance eqn linear stress $P = [X - h] b_0$
...linear along the height h
- open system, varying density $D_t \rho_0 = \mathcal{R}_0 = 0$
 $\mathcal{R}_0 = [\rho_0 / \rho_0^*]^{-m} \rho_0 - \rho_0^*$

example - the galileo giant

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example of open systems - the galileo giant

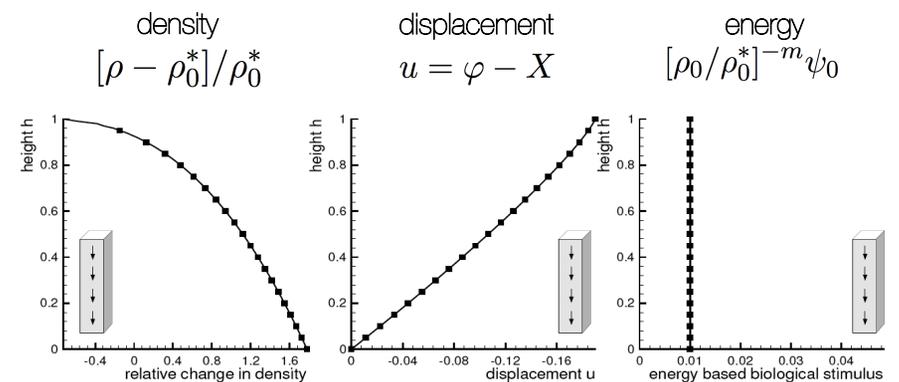


Figure 7.2: One-dimensional model problem - Open system - Homogeneous stimulus

example - the galileo giant

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constitutive equations

constitutive equations [kən'stɪ.tu.tɪv ɪ'kwer.ʒəns] in structural analysis, constitutive relations **connect applied stresses** or forces to **strains** or deformations. the constitutive relations for linear materials are linear. more generally, in physics, a constitutive equation is a relation between two physical quantities (often tensors) that is specific to a material, and does not follow directly from physical law. some constitutive equations are **simply phenomenological**; others are **derived from first principles**.



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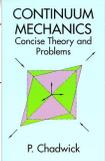
constitutive equations

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constitutive equations

constitutive equations [kən'stɪ.tu.tɪv ɪ'kwer.ʒəns] or equations of state bring in the **characterization of particular materials** within continuum mechanics. mathematically, the purpose of these relations is to supply connections between kinematic, mechanical and thermal fields. physically, constitutive equations represent the various forms of **idealized material response** which serve as **models** of the behavior of actual substances.

chadwick 'continuum mechanics' [1976]



constitutive equations

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neo hooke'ian elasticity

- free energy $\psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 \ln^2(\det(\mathbf{F})) + \frac{1}{2} \mu_0 [\mathbf{F}^t \cdot \mathbf{F} : \mathbf{I} - n^{\text{dim}} - 2 \ln(\det(\mathbf{F}))]$
- definition of stress $\mathbf{P}^{\text{neo}} = D_{\mathbf{F}} \psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 2 \ln(\det \mathbf{F}) \mathbf{F}^{-t} + \frac{1}{2} \mu_0 2 \mathbf{F} - \mu_0 \mathbf{F}^{-t} = \mu_0 \mathbf{F} + [\lambda_0 \ln(\det(\mathbf{F})) - \mu_0] \mathbf{F}^{-t}$
- definition of tangent operator $\mathbf{A}^{\text{neo}} = D_{\mathbf{F}\mathbf{F}} \psi_0^{\text{neo}} = D_{\mathbf{F}} \mathbf{P}^{\text{neo}} = \lambda_0 \mathbf{F}^{-t} \otimes \mathbf{F}^{-t} + \mu_0 \mathbf{I} \otimes \mathbf{I} + [\mu_0 - \lambda_0 \ln(\det(\mathbf{F}))] \mathbf{F}^{-t} \underline{\otimes} \mathbf{F}^{-1}$

constitutive equations

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tensor analysis - these derivatives might be handy

$$\begin{aligned} \{\bullet \otimes \circ\}_{ijkl} &= \{\bullet\}_{ij} \{\circ\}_{kl} \\ \{\bullet \bar{\otimes} \circ\}_{ijkl} &= \{\bullet\}_{ik} \{\circ\}_{jl} \\ \{\bullet \underline{\otimes} \circ\}_{ijkl} &= \{\bullet\}_{il} \{\circ\}_{jk} \end{aligned}$$

$$\begin{aligned} \frac{\partial \mathbf{F}}{\partial \mathbf{F}} &= \mathbf{I} \bar{\otimes} \mathbf{I} & \frac{\partial F_{ij}}{\partial F_{kl}} &= \delta_{ik} \delta_{jl} \\ \frac{\partial \mathbf{F}^{-1}}{\partial \mathbf{F}} &= -\mathbf{F}^{-1} \bar{\otimes} \mathbf{F}^{-t} & \frac{\partial F_{ij}^{-1}}{\partial F_{kl}} &= -F_{ik}^{-1} F_{lj}^{-1} \\ \frac{\partial \mathbf{F}^t}{\partial \mathbf{F}} &= \mathbf{I} \underline{\otimes} \mathbf{I} & \frac{\partial F_{ji}}{\partial F_{kl}} &= \delta_{il} \delta_{jk} \\ \frac{\partial \mathbf{F}^{-t}}{\partial \mathbf{F}} &= -\mathbf{F}^{-t} \underline{\otimes} \mathbf{F}^{-1} & \frac{\partial F_{ji}^{-1}}{\partial F_{kl}} &= -F_{li}^{-1} F_{jk}^{-1} \end{aligned}$$

constitutive equations

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tensor analysis - these derivatives might be handy

$$\frac{\partial \det(\mathbf{F})}{\partial \mathbf{F}} = \det(\mathbf{F}) \mathbf{F}^{-t} \quad \frac{\partial \ln(\det(\mathbf{F}))}{\partial \mathbf{F}} = \mathbf{F}^{-t}$$

$$\frac{\partial \det(\mathbf{F}^{-1})}{\partial \mathbf{F}} = -\frac{1}{\det(\mathbf{F})} \mathbf{F}^{-t} \quad \frac{\partial \ln(\det(\mathbf{F})^{-1})}{\partial \mathbf{F}} = -\mathbf{F}^{-t}$$

$$\frac{\partial \ln(\det(\mathbf{F}))}{\partial (\det(\mathbf{F}))} = \frac{1}{\det(\mathbf{F})} \quad \frac{\partial \ln^2(\det(\mathbf{F}))}{\partial \mathbf{F}} = 2 \ln(\det(\mathbf{F})) \mathbf{F}^{-t}$$

constitutive equations

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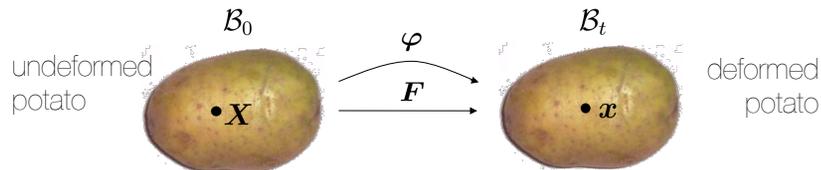
neo hooke'ian elasticity

- free energy $\psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 \ln^2(\det(F_{ij})) + \frac{1}{2} \mu_0 [F_{ij} F_{ij} - n^{\text{dim}} - 2 \ln(\det(F_{ij}))]$
- definition of stress $P_{ij}^{\text{neo}} = D_{F_{ij}} \psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 2 \ln(\det F_{ij}) F_{ji}^{-1} + \frac{1}{2} \mu_0 2 F_{ij} - \mu_0 F_{ji}^{-1} = \mu_0 F_{ij} + [\lambda_0 \ln(\det(F_{ij})) - \mu_0] F_{ji}^{-1}$
- definition of tangent operator $A_{ijkl}^{\text{neo}} = D_{F_{ij} F_{kl}} \psi_0^{\text{neo}} = D_{F_{kl}} P_{ij}^{\text{neo}} = \lambda_0 F_{ji}^{-1} F_{lk}^{-1} + \mu_0 I_{ik} I_{jl} + [\mu_0 - \lambda_0 \ln(\det(F_{ij}))] F_{li}^{-1} F_{jk}^{-1}$

constitutive equations

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neo hooke'ian elasticity

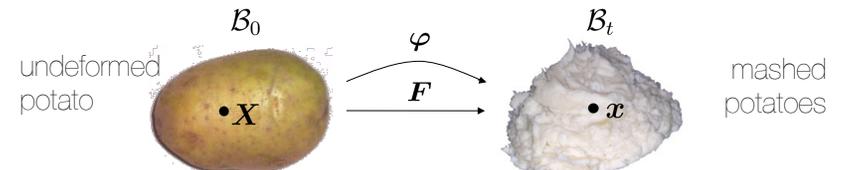


- free energy $\psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 \ln^2(\det(\mathbf{F})) + \frac{1}{2} \mu_0 [\mathbf{F}^t \cdot \mathbf{F} : \mathbf{I} - n^{\text{dim}} - 2 \ln(\det(\mathbf{F}))]$
- definition of stress $\mathbf{P}^{\text{neo}} = D_{\mathbf{F}} \psi_0^{\text{neo}} = \mu_0 \mathbf{F} + [\lambda_0 \ln(\det(\mathbf{F})) - \mu_0] \mathbf{F}^{-t}$

constitutive equations

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neo hooke'ian elasticity

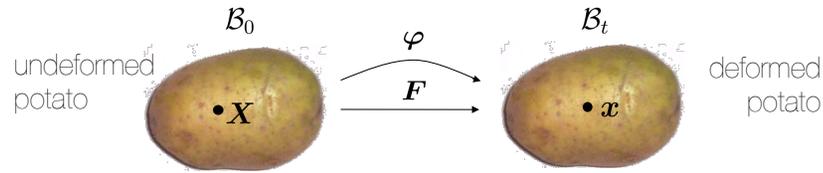


- free energy ~~$\psi^{\text{neo}} = \frac{1}{2} \lambda \ln^2(\det(\mathbf{F})) + \frac{1}{2} \mu [\mathbf{F}^t \cdot \mathbf{F} : \mathbf{I} - n^{\text{dim}} - 2 \ln(\det(\mathbf{F}))]$~~
- definition of stress ~~$\mathbf{P}^{\text{neo}} = \rho_0 D_{\mathbf{F}} \psi = \mu \mathbf{F} + [\lambda \ln(\det(\mathbf{F})) - \mu] \mathbf{F}^{-t}$~~
- remember! mashing potatoes is not an elastic process!

constitutive equations

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neo hooke'ian elasticity



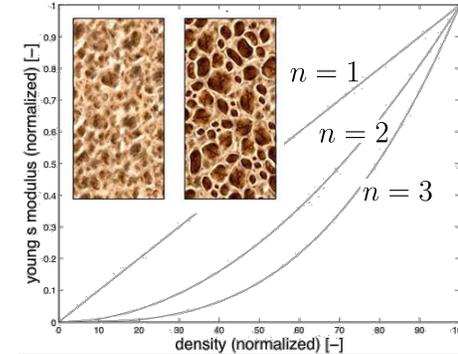
- free energy $\psi_0^{\text{neo}} = \frac{1}{2} \lambda_0 \ln^2(\det(\mathbf{F})) + \frac{1}{2} \mu_0 [\mathbf{F}^t \cdot \mathbf{F} : \mathbf{I} - n^{\text{dim}} - 2 \ln(\det(\mathbf{F}))]$
- large strain - lamé parameters and bulk modulus $\lambda = \frac{E\nu}{[1+\nu][1-2\nu]}$ $\mu = \frac{E}{2[1+\nu]}$ $\kappa = \frac{E}{3[1-2\nu]}$
- small strain – young's modulus and poisson's ratio $E = 3\kappa[1-2\nu]$ $\nu = \frac{3\kappa-2\mu}{2[3\kappa+\mu]}$

constitutive equations

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neo hooke'ian elasticity - cellular tissues

free energy $\psi_0 = \left[\frac{\rho_0}{\rho_0^*} \right]^n \psi_0^{\text{neo}}(\mathbf{F})$
 $E = 3.790 \rho_0^3 \text{ MPa}$ ρ_0 in g/cm^3

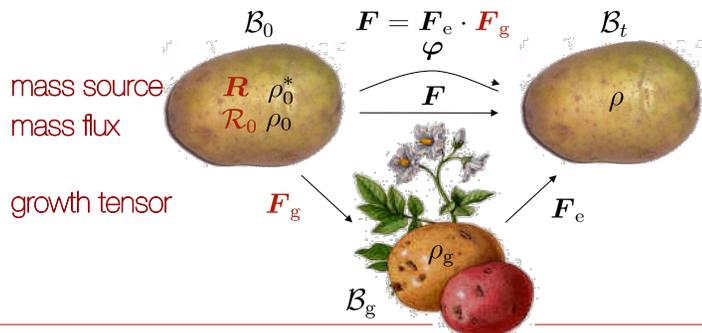


carter & hayes [1977]

constitutive equations

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additional constitutive equations for volume growth



multiplicative decomposition - but what is \mathbf{F}_g ?

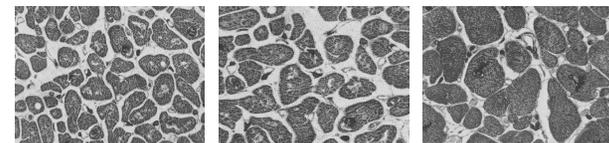
lee [1969], simo [1992], rodriguez, hoger & mc culloch [1994], epstein & maugin [2000], humphrey [2002], ambrosi & mollica [2002], himpel, kuhl, menzel & steinmann [2005]

constitutive equations

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volume growth at constant density

- free energy $\psi_0 = \psi_0^{\text{neo}}(\mathbf{F}_e)$
- stress $\mathbf{P}_e = \mathbf{P}_e^{\text{neo}}(\mathbf{F}_e)$
- growth tensor $\mathbf{F}_g = \vartheta \mathbf{I}$ $D_t \vartheta = k_\vartheta(\vartheta) \text{tr}(\mathbf{C}_e \cdot \mathbf{S}_e)$
- mass source $\mathcal{R}_0 = 3 \rho_0 \vartheta^2 D_t \vartheta$ growth function pressure increase in mass

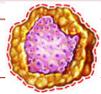


kinematic coupling of growth and deformation

rodriguez, hoger & mc culloch [1994], epstein & maugin [2000], humphrey [2002]

constitutive equations

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volume growth - cylindrical tumor

ductal carcinoma is a very common type of breast cancer in women. infiltrating ductal carcinoma refers to the development of invasive cancer cells within the milk ducts of the breast. it accounts for 80% of all types of breast cancer. on a mammography, it is usually visualized as a mass with fine spikes radiating from the edges, and small microcalcification may be seen as well. on physical examination, the lump usually feels hard or firm. on microscopic examination, the cancerous cells invade and replace the surrounding normal tissue inside the breast. tumors under 1 cm in diameter are unlikely to spread systemically. tumors under 4 cm in diameter are surgically removed. Additionally, the patient may be treated with chemotherapy, radiotherapy or hormonal therapy.



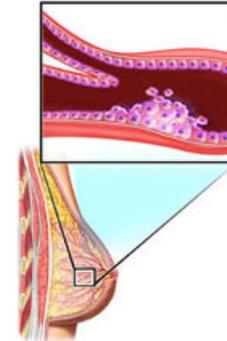
example - breast cancer

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volume growth - cylindrical tumor

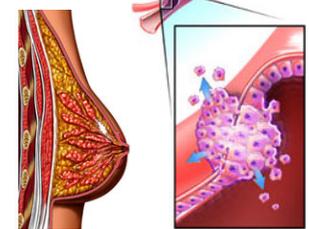
normal cells grow and multiply at a specific rate. cells that grow and multiply without stopping are called cancerous or malignant. however, they are not detectable when they first start growing.



invasive or infiltrating ductal carcinoma is the most common type of breast cancer. it occurs when the cells that line the milk duct become abnormal and spread into the surrounding breast tissue.

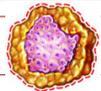


ductal carcinoma in situ is a non-invasive change in the cells that line the milk tubes that bring milk from the milk lobules to the nipple



example - breast cancer

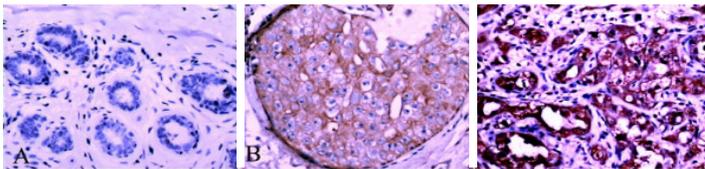
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volume growth - cylindrical tumor

model assumptions

- ductal carcinoma - tumor grows in breast duct for up to 10 cm
- model - homogeneous growth inside a rigid cylinder
- assumption - rotational symmetry
- strategy - solve for deformation that satisfies equilibrium and boundary conditions

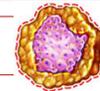


A normal ductal epithelial cells (negative) B tumor cells of ductal carcinoma (positive) C invasive ductal carcinoma cells (strongly positive)

kinematic coupling of growth and deformation

example - breast cancer

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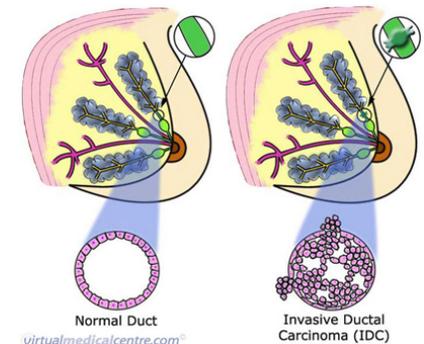
volume growth - cylindrical tumor

- homogeneous deformation inside a rigid cylinder

$$x = X \quad y = Y \quad z = \lambda Z$$

- deformation gradient

$$\mathbf{F} = \nabla_x \varphi \quad \mathbf{F} = \begin{bmatrix} 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & \lambda \end{bmatrix}$$



kinematic coupling of growth and deformation

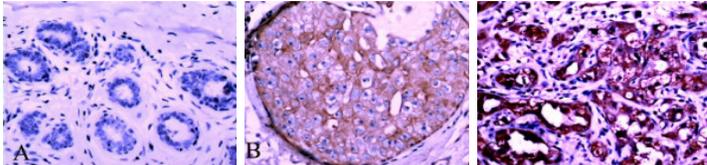
example - breast cancer

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volume growth - cylindrical tumor

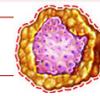
- free energy $\psi_0 = \psi_0^{\text{bko}}(\mathbf{F}_e) = \frac{1}{2}\mu[[I - 3] - \frac{2}{q}[[III]^{q/2} - 1]]$
- stress $\mathbf{P}_e = \mathbf{P}_e^{\text{bko}}(\mathbf{F}_e) = \frac{1}{J_e}\mu[-J_e^q \mathbf{F}_e^{-t} + \mathbf{F}_e] \cdot \mathbf{F}_e^{-t}$
- growth tensor $\mathbf{F}_g = \vartheta \mathbf{I} \quad \vartheta(t) = \exp(\alpha t/3)$
- deformation $x = X \quad y = Y \quad z = \lambda Z$



A normal ductal epithelial cells (negative) B tumor cells of ductal carcinoma (positive) C invasive ductal carcinoma cells (strongly positive)

kinematic coupling of growth and deformation

example - breast cancer



volume growth - cylindrical tumor

$\mathbf{F} = \nabla_X \varphi$

$$\mathbf{F} = \begin{bmatrix} 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & \lambda \end{bmatrix}$$

$\mathcal{B}_0 \xrightarrow{\mathbf{F}} \mathcal{B}_t$

$\mathbf{F}_g = \vartheta \mathbf{I}$

$$\mathbf{F}_g = \begin{bmatrix} \vartheta & 0 & 0 \\ 0 & \vartheta & 0 \\ 0 & 0 & \vartheta \end{bmatrix}$$

$\mathbf{F}_e = \mathbf{F} \cdot \mathbf{F}_g^{-1}$

$$\mathbf{F}_e = \begin{bmatrix} \frac{1}{\vartheta} & 0 & 0 \\ 0 & \frac{1}{\vartheta} & 0 \\ 0 & 0 & \frac{\lambda}{\vartheta} \end{bmatrix}$$

\mathcal{B}_g

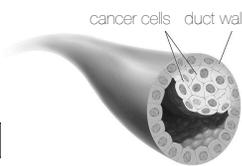
ambrosi & molica [2002]

example - breast cancer



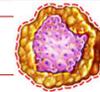
volume growth - cylindrical tumor

- stress $\boldsymbol{\sigma}_e = \begin{bmatrix} \sigma_{xx} & 0 & 0 \\ 0 & \sigma_{yy} & 0 \\ 0 & 0 & \sigma_{zz} \end{bmatrix}$
- $\sigma_{xx} = \sigma_{yy} = \mu \frac{\vartheta^3}{\lambda} \left[\frac{1}{\vartheta^2} - \left[\frac{\lambda}{\vartheta^3} \right]^q \right]$
- $\sigma_{zz} = \mu \frac{\vartheta^3}{\lambda} \left[\frac{\lambda^2}{\vartheta^2} - \left[\frac{\lambda}{\vartheta^3} \right]^q \right]$
- bc's $\sigma_{zz} = 0$
- axial displacement λ as a function of growth ϑ
- $\lambda = \vartheta^{[2-3q]/[2-q]}$
- growth induced stress σ_{xx} on tumor wall
- $\sigma_{xx} = \mu \left[\vartheta^{2q/[2-q]} - \vartheta^{[4-4q]/[2-q]} \right]$

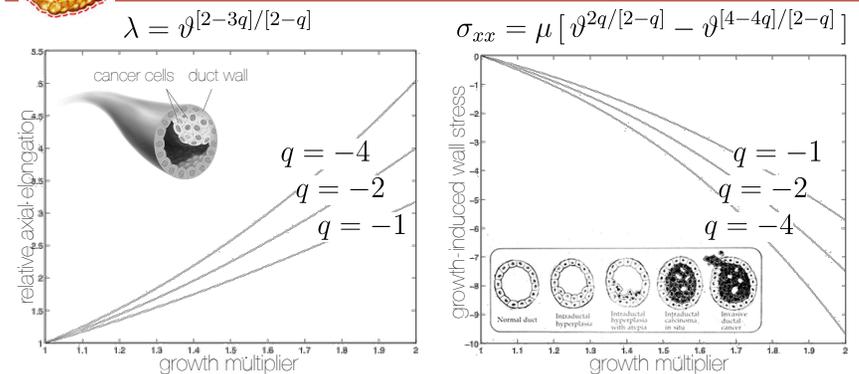


ambrosi & molica [2002]

example - breast cancer



volume growth - cylindrical tumor



tumor pressure on duct walls increases with growth

ambrosi & molica [2002]

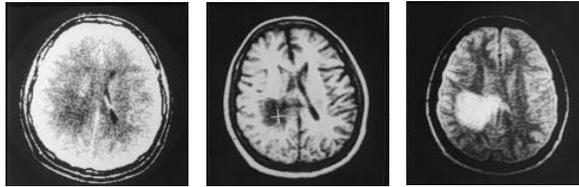
example - breast cancer



volume growth - spherical tumor

model assumptions

- tumor model - inhomogeneous growth of a sphere
- inhomogeneity - residual stress even in the absence of applied loads
- assumption - spherical symmetry
- strategy - solve for deformation that satisfies equilibrium and boundary conditions

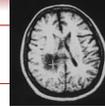


kinematic coupling of growth and deformation

ambrosi & mollica [2002]

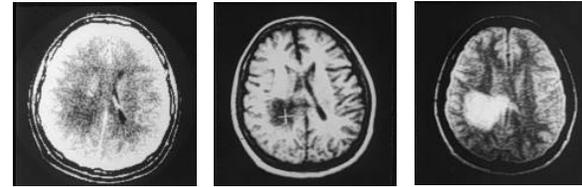
example - tumor growth

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volume growth - spherical tumor

- free energy $\psi_0 = \psi_0^{\text{bko}}(\mathbf{F}_e) = \frac{1}{2}\mu[[I - 3] - \frac{2}{q}[[III]^{q/2} - 1]]$
- stress $\mathbf{P}_e = \mathbf{P}_e^{\text{bko}}(\mathbf{F}_e) = \frac{1}{J_e}\mu[-J_e^q \mathbf{F}_e^{-t} + \mathbf{F}_e] \cdot \mathbf{F}_g^{-t}$
- growth tensor $\mathbf{F}_g = \vartheta \mathbf{I} \quad \vartheta(R) = \frac{\alpha}{R} \sinh(kR)$
- deformation $r = \lambda(R) \quad \theta = \Theta \quad \phi = \Phi$



kinematic coupling of growth and deformation

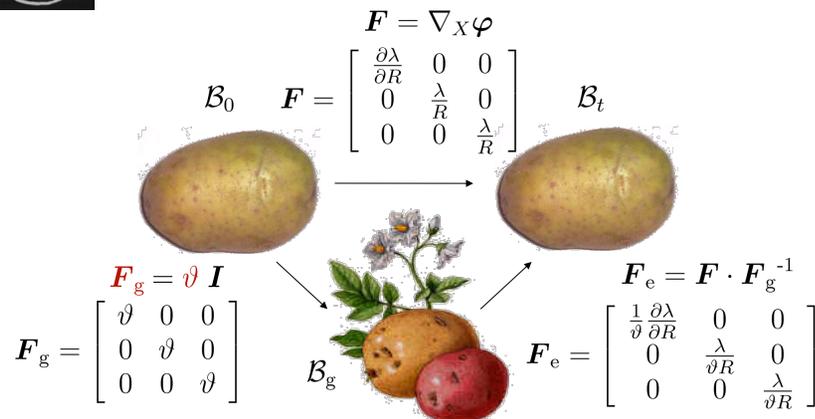
ambrosi & mollica [2002]

example - tumor growth

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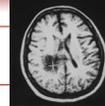
volume growth - spherical tumor



ambrosi & mollica [2002]

example - tumor growth

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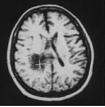
volume growth - spherical tumor

- stress $\mathbf{P}_e = \begin{bmatrix} P_{RR} & 0 & 0 \\ 0 & P_{\Theta\Theta} & 0 \\ 0 & 0 & P_{\Phi\Phi} \end{bmatrix}$
- $P_{RR} = \mu \vartheta \left[\frac{\partial \lambda}{\partial R} - J_e^q \vartheta^2 \left[\frac{\partial \lambda}{\partial R} \right]^{-1} \right]$
- $P_{\Theta\Theta} = P_{\Phi\Phi} = \mu \vartheta \left[\frac{\lambda}{R} - J_e^q \vartheta^2 \frac{R}{\lambda} \right]$
- $J_e = \frac{\partial \lambda}{\partial R} \frac{\lambda^2}{\vartheta^3 R^2}$
- balance of momentum $\frac{dP_{RR}}{dR} + \frac{2}{R}[P_{RR}R - P_{\Theta\Theta}] = 0$
- bc's $\lambda(R=0) = 0 \quad P_{RR}(R=\bar{R}) = 0$

ambrosi & mollica [2002]

example - tumor growth

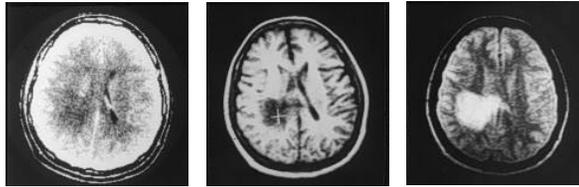
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volume growth - spherical tumor

ordinary differential eqn of 2nd order in λ 2B solved numerically

$$\frac{\partial^2 \lambda}{\partial R^2} = \frac{\frac{2}{R} \left[\frac{1}{R} + \frac{\vartheta^2 J_e^q [1-q]}{\lambda \partial \lambda / \partial R} \right] \left[\lambda - \frac{\partial \lambda}{\partial R} R \right] - \frac{1}{\vartheta} \left[\frac{\partial \lambda}{\partial R} - \frac{3 \vartheta^2 J_e^q [1-q]}{\partial \lambda / \partial R} \frac{\partial \vartheta}{\partial R} \right]}{1 + \frac{\vartheta^2 J_e [1-q]}{[\partial \lambda / \partial R]^2}}$$

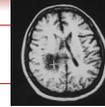


kinematic coupling of growth and deformation

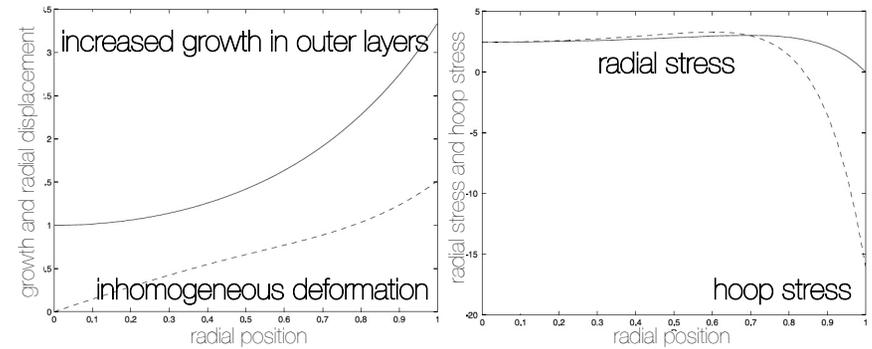
ambrosi & mollica [2002]

example - tumor growth

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volume growth - spherical tumor



hoop stress tensile inside / compressive outside

ambrosi & mollica [2002]

example - tumor growth

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