## Math and Medicine: Homework Assignment 7 Due on October 20

- 1. Consider a multistage carcinoma model where the mutation rate of a tissue from stage i (with i "hits") to stage i + 1 is  $(i + 1)\mu$  (that is, the mutation rate from stage 1 to stage 2 is double that of stage 0 to stage 1, that from stage 2 to stage 3 is triple that of stage 0 to stage 1, and so forth).
  - **a.** Write the equations for a *k*-hit cancer.
  - **b.** Find how incidence I(a) depends on age for a two-hit cancer. Does the graph of  $\ln(I(a))$  against  $\ln(a)$  fall approximately along a line? If so, what is the slope of that line?
  - c. Extra Credit: How well does this work for a k-hit cancer?
- 2. Some cancer-associated viruses, like hepatitis C, do not directly cause cancer, but just increase cell turnover rates. Suppose infection increases the cell turnover rate by a factor of  $\theta$ .
  - **a.** Write differential equations for the populations of cells  $S_i$  with *i* hits, assuming that all cells begin in stage 0, and that the mutation rate from stage *i* to i + 1 is  $\mu$ .
  - **b.** Is the incidence of cancer increased by a factor of  $\theta$  for a two-hit cancer?
  - c. Is the incidence increased by a factor of  $\theta$  or  $\theta^{k-1}$  for a k-hit cancer?