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% Supplemental code 1

clear all
close all
% rate of spacer acquisition  $dS/dt = r * Ni$ 
% cells grow at rate  $dNi/dt = \mu * Ni$ 
% cells grow from  $10^4$  cells to  $10^8$  cells and then are transferred

% general parameters
No =  $10^4$ ; % number of cells inoculated into each tube
Nf =  $10^8$ ; % number of cells at a cell transfer event
r = 8.16833E-05; % array expansion rate per cell per minute
mu = 0.02; % in 1/min
max_time = 14400; % maximum time in minutes
time_points = 2880; % number of time steps
dt = max_time / time_points;
max_array = 21; % maximum extension of the array - 1 (array length 1 = no extension)

% start at 24 h
t_min = 24*60/dt;

% parameters for modified model
mu_red = 0.995; % cell growth rate can be reduced for cells with expanded arrays
r_red = 1.00; % array expansion rate can be reduced for cells with expanded arrays

% colors and markers for plotting
C = {'ok', 'db', 'sr', 'xg', '+c'};
day_counter = 1;

% cells stores the number of cells at each array length
% cell(i) is information about the cells with array length i-1 (i=1 is
% unexpanded)
% cells.array is the array length in number of new spacers
% with 0 being the original length
% cells.num is the number of cells with the specified array length

% initialize population of cells
for j = 1:max_array
cells(j).array = j-1;
cells(j).num = 0;
cells(j).growth = mu*mu_red^(j-1);
cells(j).r = r*r_red^(j-1);
end

% at 24 h start with 6.5% +1 cells and 93.5% +0 cells
cells(1).num = No*0.978;
cells(2).num = No*0.022;

for t = t_min:time_points+1 % time in minutes

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% does array extension occur
for k = 1:length(cells)
    cell_flips = rand(floor(cells(k).num),1);
    number_ext = sum(cell_flips<(cells(k).r*dt));
    % if there is extension, move cells to new array length
    if number_ext > 0
        array_bin = cells(k).array+2;
        cells(array_bin).num = cells(array_bin).num+number_ext;
        cells(k).num = cells(k).num - number_ext;
    end

    % the cells grow
    cells(k).num = cells(k).num + cells(k).growth*cells(k).num*dt;
end

% transfer cells to a new tube if the population exceeds maximum
tot_cells = floor(sum([cells.num]));

if tot_cells>Nf

% calculate the fraction of cells at each array length
for m = 1:length(cells)
    cells(m).fract = cells(m).num/tot_cells;
end
fraction = [0, [cells.fract]];

% cells are partitioned by picking No random numbers, each random number
% assigns 1 cells to a specific array length
    cell_partition = rand(No,1);
    for m = 1:length(cells)
% find how many random numbers bewteen certain values
        lower_end = cell_partition>sum(fraction(1:m));
        upper_end = cell_partition <= sum(fraction(1:m+1));
        combined = and(lower_end, upper_end);
        transf_cells = sum(combined);
        cells(m).num = transf_cells;
    end

end

% show make a few plots after every 24 h
if mod((t-1)*dt,1440)==0

tot_cells = floor(sum([cells.num]));

% recalculate the fraction of cells at each array length
for m = 1:length(cells)
    cells(m).fract = cells(m).num/tot_cells;
    band(m) = cells(m).fract;
end

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% next line calculates the average array length
av_array_length = sum([0:1:max_array-1] * band');

% plot the average array length over time
figure(1)
hold on
plot(t*dt/(24*60), av_array_length, 'ko')
hold off
xlabel('time / day')
ylabel('average array length')

% plot the expanded fraction over time
figure(2)
hold on
plot(t*dt/(24*60), sum(band(2:end)), 'bo')
hold off
xlabel('time / day')
ylabel('percent expanded')

% plot the fraction of cells at each length
figure(3)
hold on
for j = 1:5
plot(t*dt/(24*60), cells(j).fract, C{j})
end
legend('+0', '+1', '+2', '+3', '+4')
hold off
xlabel('time / day')
ylabel('fraction of population')

% save the data for each 24 h period
data_out(day_counter).time = (t-1)*dt/(24*60);
data_out(day_counter).av_array_length = av_array_length;
data_out(day_counter).per_exp = sum(band(2:end));
data_out(day_counter).per_zeros = cells(1).fract;
data_out(day_counter).per_ones = cells(2).fract;
data_out(day_counter).per_twos = cells(3).fract;
data_out(day_counter).per_three = cells(4).fract;
data_out(day_counter).per_four = cells(5).fract;

% the day counter indexes data_out
day_counter = day_counter+1;

end

end

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