

UiO : Kjemisk institutt

Det matematisk-naturvitenskapelige fakultet

FYS4150, project 5

Sondre Torp - Sondrt@student.matnat.uio.no



1 Abstract

The goal of this project was to develop a Monte Carlo simulation of the spread of an infectious disease. This was done with the classical SIRS model of epidemiology, and it was studied and used to develop the necessary transition probabilities for the simulation.

A deterministic approach, the 4th order Runge-Kutta, was used in conjunction with the Monte Carlo method on the classical SIRS model. These were then tested and improved to suffice for different scenarios and diseases.

We found that the Monte Carlo is a solid and semi-realistic approach to fluctuations in diseases, given enough data.

This is far from a done analyzes; The model should be tested on real data, parameters should be added; incubation time could be one, and the code should go through a major makeover so that it is easier to expand the model and more intuitive to use. It should also be made a simple way of tracing individuals in the group.

Innhold

1 Abstract	1
2 Abstract	2
3 Introduction	3
4 Physical Theory	4
4.1 The SIRS	4
4.1.1 The steady state	5
4.2 Improvements to SIRS	5
4.2.1 Vital dynamics	5
4.2.2 Seasonal Variation	6
4.2.3 Vaccination	6
5 Computational Theory & implementation	6
5.1 4th order Runge-Kutta	6
5.2 Monte Carlo Simulation	8
5.3 Neural networks	8
6 Result & Discussion	10
6.1 SIRS modelling	10
6.1.1 4th order Runge-Kutta on the ODEs from SIRS, testing recovery rate.	10
6.1.2 Neural networks	11
6.1.3 Monte Carlo simulation of simple SIRS model, using different time.	11
6.1.4 Monte Carlo simulation of simple SIRS model, testing recovery rate	12
6.2 SIRS modelling with Improvements	14
6.2.1 Vital dynamics	14
6.2.2 Seasonal Variation	16
6.2.3 Vaccination	17
7 Conclusion	17
8 References:	18

2 Abstract

The goal of this project was to develop a Monte Carlo simulation of the spread of an infectious disease. This was done with the classical SIRS model of epidemiology, and it was studied and used to develop the necessary transition probabilities for the simulation.

A deterministic approach, the 4th order Runge-Kutta, was used in conjunction with the Monte Carlo method on the classical SIRS model. These were then tested and improved to suffice for different scenarios and diseases.

We found that the Monte Carlo is a solid and semi-realistic approach to fluctuations in diseases, given enough data.

This is far from a done analysis; The model should be tested on real data, parameters should be added; incubation time could be one, and the code should go through a major makeover so that it is easier to expand the model and more intuitive to use. It should also be made a simple way of tracing individuals in the group.

3 Physical Theory

3.1 The SIRS

The SIRS model is the one of the simplest compartmental models, and is very similar to the SIR model, with one exception, this exception stopes the SIRS model from being solvable analythically.

The **SIRS** model considers an isolated population N , that consists of three compartments:

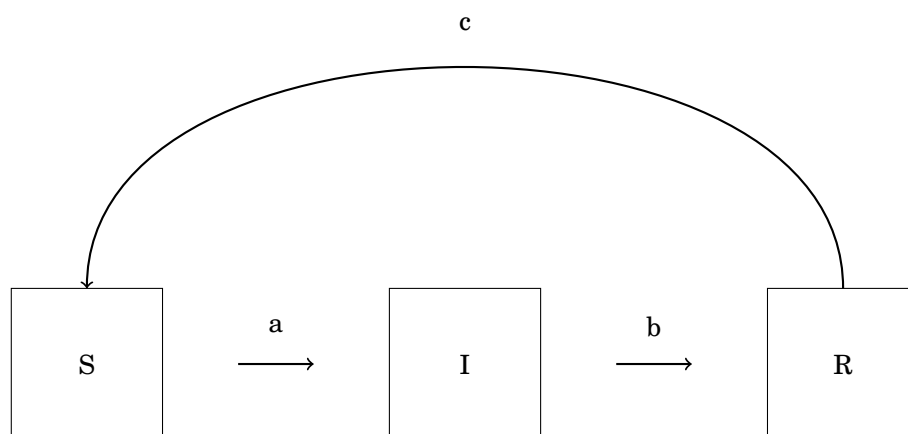
- **S**: The number of people without immunity to the disease.
- **I**: The number of people who are currently infected
- **R**: The number of past infected whom have developed an immunity to the disease.

These variables (**S**, **I**, **R**) represent the number of people in each compartment, and since the size of these are dependent on time, we make the precise number a function of time. ($\mathbf{S}(t)$, $\mathbf{I}(t)$, $\mathbf{R}(y)$). For a specific disease in a specific population, these functions can be worked out to predict possible outbreaks and perhaps bring them under control.

The model is dynamic in that the number in each compartment may change over time. This is dynamic aspect is most obvious in an endemic disease. The main difference between the SIRS model and the SIR model is the fact that in the SIR model a member of the population can never join the susceptible group, ($S(t)$), again after leaving, such as the disease measles. This type of disease can not break out again until the number of susceptible people has built back up. While in the SIRS model, you can lose"your immunity to the disease. A person can move from one group to another as indicated in the figure 1 below.

The rate of transmission a , the rate of recovery b and the rate of immunity lose c , helps describing the flow of people moving between the groups. The group is assumed to be mixed homogeneously and the total population remains constant.

$$N = S(t) + I(t) + R(t) \quad (1)$$



Figur 1: A flow diagram in which the boxes represent the different compartments and the arrows the transition between the compartments for a SIRS model.

If we assume that the time scale is much smaller than the average person's lifetime, than the effect of the birth and death rate of the population can be ignored. For these assumption we are given a set of coupled differential equations that we use to construct our model.

$$\frac{dS}{dt} = cR - \frac{aSI}{N} \quad (2)$$

$$\frac{dI}{dt} = \frac{aSI}{N} - bI \quad (3)$$

$$\frac{dR}{dt} = bI - cR \quad (4)$$

Though this set does not have a analytic solution, the equilibrium solutions are simple to obtain. The constrain ?? reduces the three dimensional system into a two dimensional one.

$$\frac{dS}{dT} = c(N - S - I) - \frac{aSI}{N} \quad (5)$$

$$\frac{dI}{dT} = \frac{aSI}{N} - bI \quad (6)$$

$$(7)$$

3.1.1 The steady state

The steady state is found by setting both of these equations equal to zero. The fraction of people in each group will then be:

$$s^* = \frac{b}{a} \quad (8)$$

$$i^* = \frac{1 - \frac{b}{a}}{1 + \frac{b}{c}} \quad (9)$$

$$r^* = \frac{b}{c} \frac{1 - \frac{b}{a}}{1 + \frac{b}{c}} \quad (10)$$

Each fraction must be between (0 – 1) and the three factions must sum up to 1. This also shows that $b < a$ for the disease to establish itself in the population.

3.2 Improvements to SIRS

The same principles as for the simple model are utilized to extend the model to include more details about the population and the disease.

3.2.1 Vital dynamics

Vital dynamics are added so that the model can describe the spread of the diseases which occur over longer stretches of time. If e is introduced as the brith rate, d and d_I as death rate, and death rate for the infected people due to the disease, then the differential equations, assuming all babies born are susceptible, are given by:

$$\frac{dS}{dT} = cR - \frac{aSI}{N} - dS + eN \quad (11)$$

$$\frac{dI}{dT} = \frac{aSI}{N} - bI - dI - d_I I \quad (12)$$

$$\frac{dR}{dT} = bI - cR - dR \quad (13)$$

3.2.2 Seasonal Variation

Seasonal variations can also be added to better represent diseases such as influenza, where the rate of transmission depends largely on the time of the year. During cold months individuals are more likely to spend time in closer proximity to one another, resulting in a higher rate of transmission. We change the transmission rate (a), so that it oscillates.

$$a(t) = A \cos(\omega t) + a_0 \quad (14)$$

where a_0 is the average transmission rate, A is the maximum deviation from a_0 , and ω is the frequency of oscillation.

3.2.3 Vaccination

Diseases with available vaccinations allow people to move directly from S to R , breaking the cyclic structure.[zaman2008stability] Here it is assumed that a susceptible individual's choice to become vaccinated does not depend on how many other susceptible are vaccinated or the anti vaccination movement. We can assume that the rate of vaccination, f , depends on time, since this rate may oscillate during the course of a year and/or increase as awareness and medical research increase. The differential equations become:

$$\frac{dS}{dT} = cR - \frac{aSI}{N} - f \quad (15)$$

$$\frac{dI}{dT} = \frac{aSI}{N} - bI \quad (16)$$

$$\frac{dR}{dT} = bI - cR + f \quad (17)$$

4 Computational Theory & implementation

The code used for all models of SIRS can be found here: [github](#).

4.1 4th order Runge-Kutta

The 4th order Runge-Kutta(RK4) is one of the classic methods for numerical integration of ODE models. For a brief introduction of RK4 refers to Wikipedia.

For this problem; consider the following initial value problem of ODE

$$\frac{dy}{dt} = f(t, y) \quad (18)$$

$$y(t_0) = y_0 \quad (19)$$

where $y(t)$ is the unknown function which we would like to approximate.

The iterative formula of RK4 method for solving ODE 19 is as follows:

$$y_{n+1} = y_n + \frac{\Delta t}{6} (k_1 + 2k_2 + 2k_3 + k_4) \quad (20)$$

$$\begin{aligned}
k_1 &= f(t_n, y_n) \\
k_2 &= f(t_n + \frac{\Delta t}{2}, y_n + \frac{k_1 \Delta t}{2}) \\
k_3 &= f(t_n + \frac{\Delta t}{2}, y_n + \frac{k_2 \Delta t}{2}) \\
k_4 &= f(t_n + \Delta t, y_n + k_3 \Delta t) \\
t_{n+1} &= t_n + \Delta t \\
n &= 0, 1, 2, 3, \dots
\end{aligned}$$

The SIRS model is defined as follows:

$$\begin{aligned}
\frac{dS}{dt} &= cR - \frac{aSI}{N} \\
\frac{dI}{dt} &= \frac{aSI}{N} - bI \\
\frac{dR}{dt} &= bI - cR
\end{aligned}$$

with a, b, c, S, I, R and N as previously defined.4.1

According to the general iterative formula 20, the iterative formulas for $S(t), I(t), R(t)$ of SIRS model can be written out:

$$\begin{aligned}
S_{n+1} &= S_n + \frac{\Delta t}{6}(k_1^S + 2k_2^S + 2k_3^S + k_4^S) \\
k_1 &= f(t_n, S_n, I_n, R_n) = cR - \frac{aSI}{N} \\
k_2 &= f(t_n + \frac{\Delta t}{2}, S_n + \frac{k_1^S \Delta t}{2}, I_n + \frac{k_1^I \Delta t}{2}, R_n + \frac{k_1^R \Delta t}{2}) = -\frac{a}{N}(S_n + \frac{k_1^S \Delta t}{2})(I_n + \frac{k_1^I \Delta t}{2}) + c(R_n + \frac{k_1^R \Delta t}{2}) \\
k_3 &= f(t_n + \frac{\Delta t}{2}, S_n + \frac{k_2^S \Delta t}{2}, I_n + \frac{k_2^I \Delta t}{2}, R_n + \frac{k_2^R \Delta t}{2}) = -\frac{a}{N}(S_n + \frac{k_2^S \Delta t}{2})(I_n + \frac{k_2^I \Delta t}{2}) + c(R_n + \frac{k_2^R \Delta t}{2}) \\
k_4 &= f(t_n + \Delta t, S_n + k_3^S \Delta t, I_n + k_3^I \Delta t, R_n + k_3^R \Delta t) = -\frac{a}{N}(S_n + k_3^S \Delta t)(I_n + k_3^I \Delta t) + c(R_n + k_3^R \Delta t)
\end{aligned}$$

$$\begin{aligned}
I_{n+1} &= I_n + \frac{\Delta t}{6}(k_1^I + 2k_2^I + 2k_3^I + k_4^I) \\
k_1^I &= \frac{aS_n I_n}{N} - bI_n \\
k_2^I &= \frac{a}{N}(S_n + \frac{k_1^S \Delta t}{2})(I_n + \frac{k_1^I \Delta t}{2}) - b(I_n + \frac{k_1^I \Delta t}{2}) \\
k_3^I &= \frac{a}{N}(S_n + \frac{k_2^S \Delta t}{2})(I_n + \frac{k_2^I \Delta t}{2}) - b(I_n + \frac{k_2^I \Delta t}{2}) \\
k_4^I &= \frac{a}{N}(S_n + k_3^S \Delta t)(I_n + k_3^I \Delta t) - b(I_n + k_3^I \Delta t)
\end{aligned}$$

$$\begin{aligned}
R_{n+1} &= R_n + \frac{\Delta t}{6}(k_1^R + 2k_2^R + 2k_3^R + k_4^R) \\
k_1^R &= bI_n - cR_n \\
k_2^R &= b(I_n + \frac{k_1^I \Delta t}{2}) - c(R_n + \frac{k_1^R \Delta t}{2}) \\
k_3^R &= b(I_n + \frac{k_2^I \Delta t}{2}) - c(R_n + \frac{k_2^R \Delta t}{2}) \\
k_4^R &= b(I_n + k_3^I \Delta t) - c(R_n + k_3^R \Delta t)
\end{aligned}$$

Pew, now note that since the population $N = S(t) + I(t) + R(t)$ is constant, they will have $0 = \frac{dS}{dt} + \frac{dI}{dt} + \frac{dR}{dt}$. Meaning that only two of the three ODEs are independent and are thus sufficient to solve the ODEs. In our code, we ignore this fact, so that we can utilize the full power of our cpu.

4.2 Monte Carlo Simulation

This project consists of different cases with a different degree of complexity. For the basis case it is enough to use the Monte Carlo method. This method utilizes the fact that a large number of experiments converges towards the expectation value. When the probability of doing something is added, the system utilizes the Metropolis Algorithm. The system has to do a random choice of acceptance or denial of the case. This random choice is done with the Random Number Generator from NumPy[NumPy]. A random number is created between $(0 - 1)$. If the number is less than the probability term, accept the new state. If the number is greater than the probability for change then discard the change. The procedure is outlined in four steps:

- Chose a transition state randomly.
- Find the probability for the transition.
- A RNG is used to chose a number between $(0 - 1)$. Now if the probability term is less then the RNG number, reject the transition. If not, accept the tranistion.
- Update the system with the transferred states.

4.3 Neural networks

First of all we wanted to make sure that we could train the neural network(NN) to reproduce the RK4 solution. This is certainly a prerequisite to the idea working. The NN will here output three values, one for each concentration. We also chose to use the Swish function due to this article 8. We later tested for OTHER.

We then trained our network to reproduce the solution, one layer network with 8 nodes to output all three concentrations pretty accurately. This confirmed that the solution could be represented by a neural network. (Se the section for results) REF.

The next major issue is how do we get the relevant derivatives. The solution method developed here relies on using optimization to find a set of weights that produces a NN whose derivatives are consistent with the ODE equation. The NN outputs three concentrations, and we need the time derivatives of them. In Autograd we found three options; grad, elementwise_grad and Jacobian. We cannot use grad because our function is not scalar. We cannot use elementwise_grad because that would give us the wrong shape. This left us with one option. This was a little weird because it gave an output that was 4-dimensional, as expected from the documentation, the first and third output where related to our time steps, then we used some fancy sorting to see if the data was comparable to the derivatives defined by the ODEs. This was inefficient, due to it requiring a lot of calculations to create the jacobian.

Finally we solved the system with our NN, firstly defining a time grid to solve it on, defining our objective function, taking care of initial conditions, running the optimization and getting the solution from the NN.

5 Result & Discussion

5.1 SIRS modelling

Here we confirmed that the recovery rate, b , should be smaller than the rate of transmission, a , to sustain a disease. We saw that there is a surprisingly good match between the RK4 and the MC model, but the MC model needs more time, approximately 3 years, to stabilize and find a steady-state in the population. Finally we figured that the MC simulation crashes if the number of infected people reaches zero. Details can be read below.

The equilibrium expectation values and corresponding standard deviation, after 10 years of time or 3650 days, were found to be: [Nobody06]

	S(t)	I(t)	R(t)
Mean	108.81	110.11	181.06
Stdev	39.48	32.06	28.24

5.1.1 4th order Runge-Kutta on the ODEs from SIRS, testing recovery rate.

The simplest form of SIRS models differential equations solved with 4th order Runge - Kutta, then plotted. All plots in figure: 2, have a black line that represent the change in total population. One can see there is not much change in population, as expected, at this moment. The equilibrium-states, discussed in section: 4.1.1, are plotted with stapled lines. These solutions are only plotted for one year, due to their obvious approach of the equilibrium-states. The difference between these figures are due to different recovery rate. One can see from these figures that after the initial burst of infection most develops an immunity after a short time and the disease reaches a steady-state. We can also confirm that if the transmission rate is larger than the recovery rate ($b < a$) the disease manage to get some kind of foothold, to sustain itself in the population.

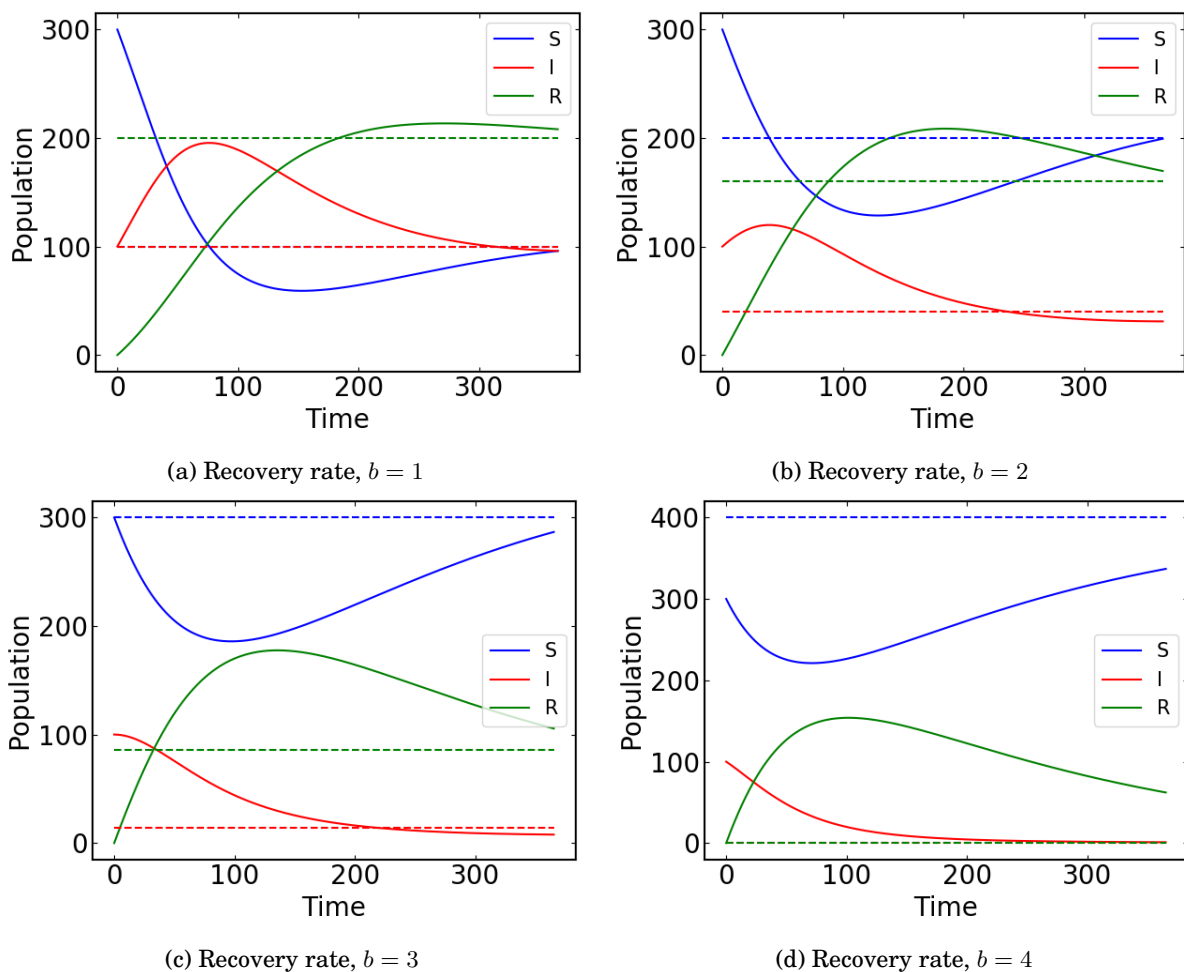
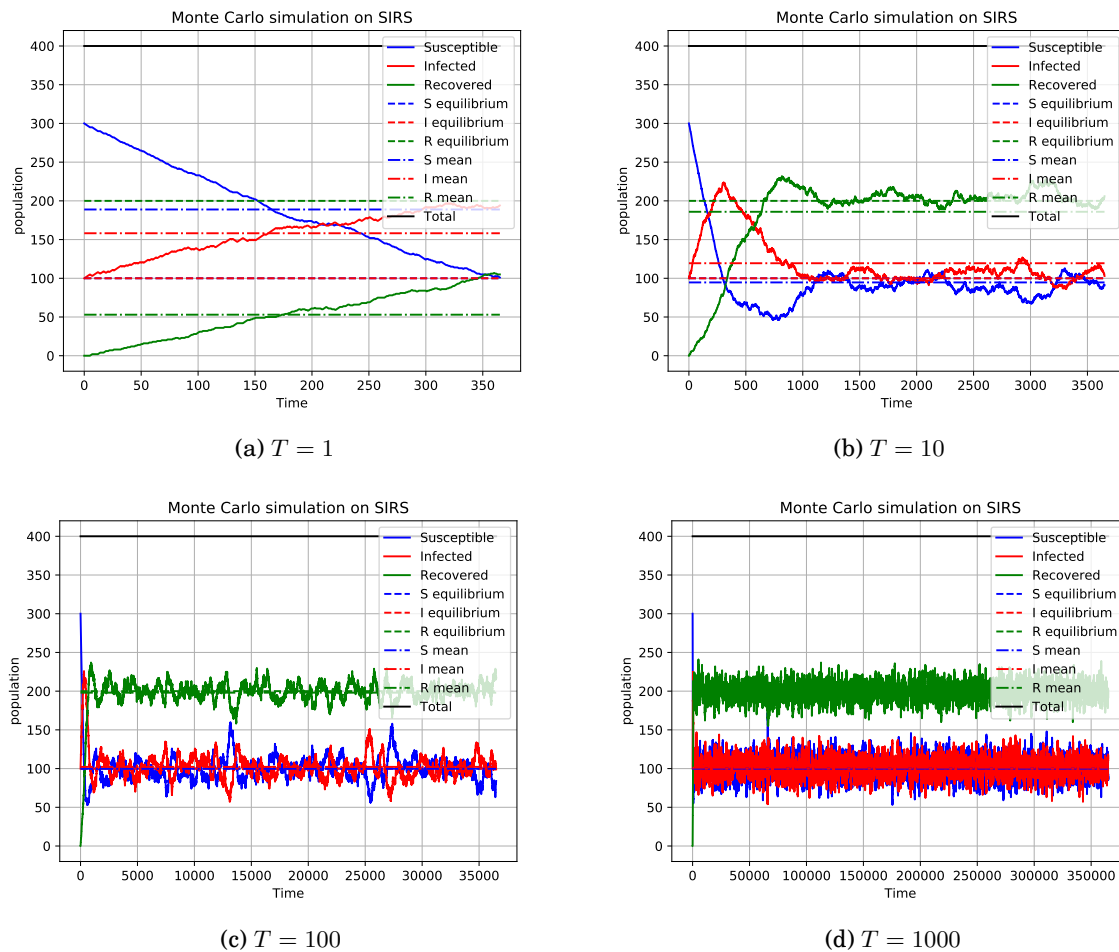


Figure 2: 4th order Runge-Kutta on ODEs given for SIRS model, where the only difference is the recovery rate, b , that is equal to 1, 2, 3, 4 respectively

5.1.2 Neural networks

5.1.3 Monte Carlo simulation of simple SIRS model, using different time.

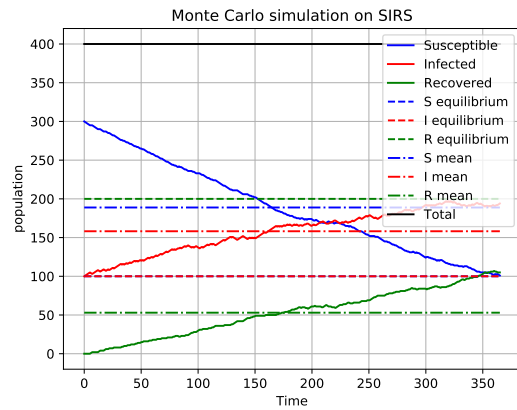
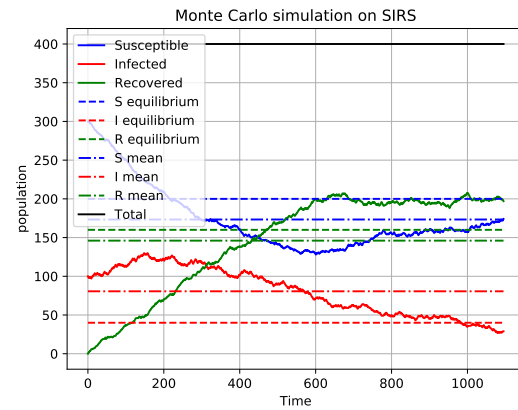
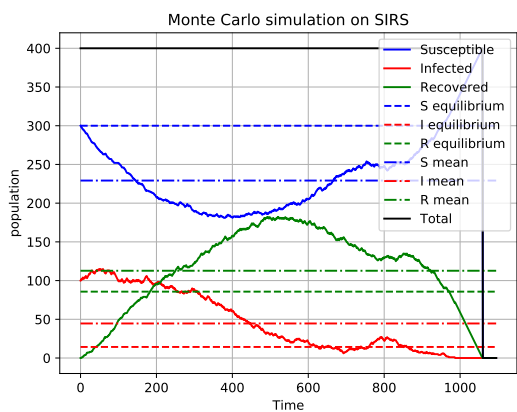
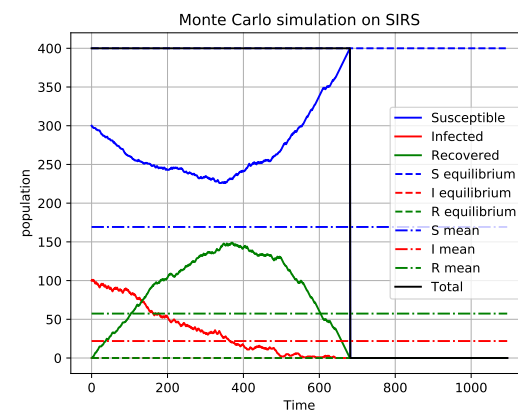
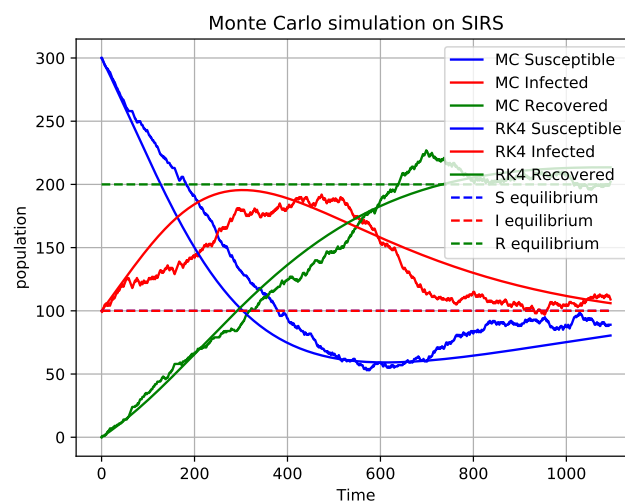
Firstly we quickly understood that one year was simply not enough time to get to a proper steady-state, so we look at how the simulation would look if we ran it for 1, 10, 100, 1000 years. And we quickly figured that 5 to 10 years should suffice, depending on what effects we want to observe. Here the both the equilibrium line, same as before, and the mean of the Monte Carlo simulation are plotted, and even after 1000 years, these lines were still off by 1 or 2.



Figur 3

5.1.4 Monte Carlo simulation of simple SIRS model, testing recovery rate

In these figures we notice something spectacular! The program crashes at higher recovery rate! This is due to the fact that there are no more infected people at some point in the program, and there is no reason to track an infection's development if there is no infection. It is a theoretical steady-state for the recovery rate, b , values that are smaller than the transmission rate, a , values, but due to the randomness of the Monte Carlo Simulations the infection tends to die out by itself, which makes a lot of sense.

(a) $T = 3$ and $b = 1$ (b) $T = 3$ and $b = 2$ (c) $T = 3$ and $b = 3$ (d) $T = 3$ and $b = 4$ (e) $T = 3$ and $b = 1$, both MC and RK4 plotted on top of each other. Removed the total population bar, due to space.

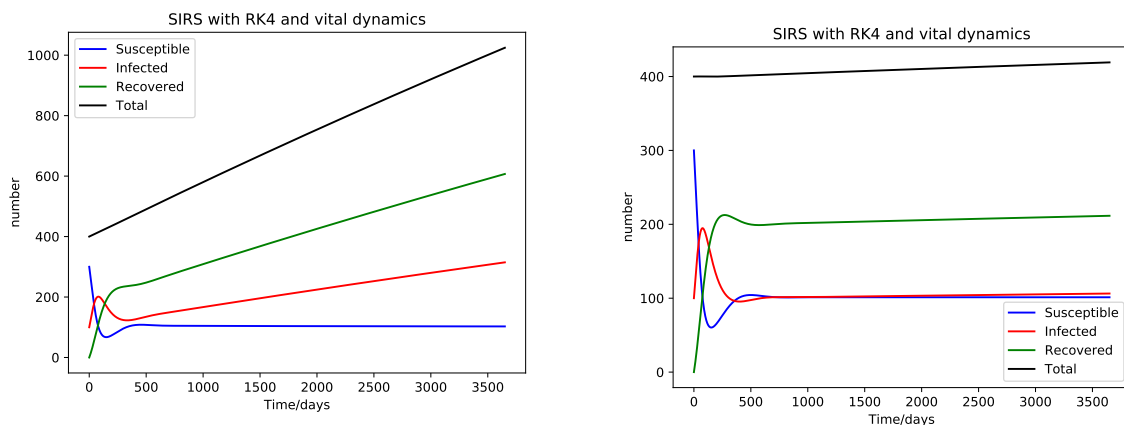
Figur 4

5.2 SIRS modelling with Improvements

Here we present the results from the different versions of SIRS, Vital dynamics, Seasonal variation and Vaccination, and some of these combined.

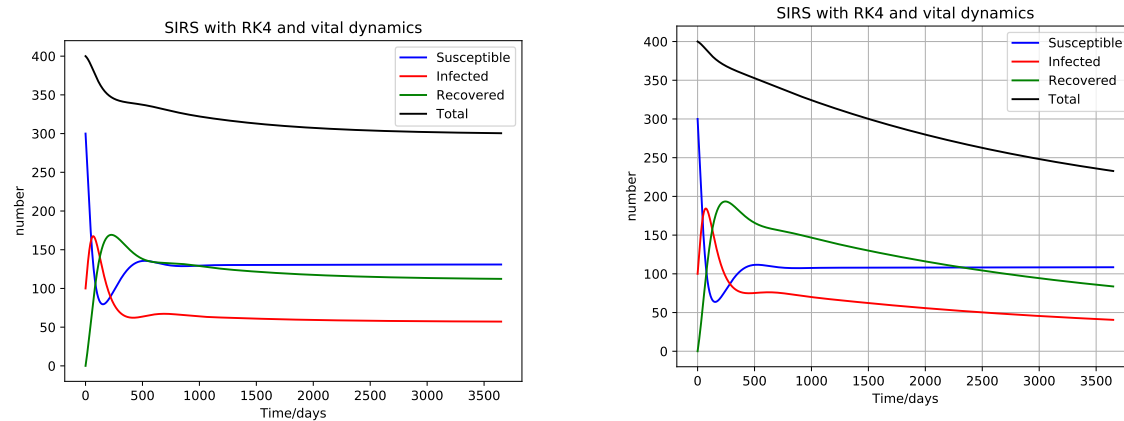
5.2.1 Vital dynamics

Vital dynamics are here included as presented in the theory section 4.2.1 for both RK4 and MC. There is some difference in the given death/birth - rates between RK4 and MC due to lack of normalization dt . It is easy to see that these numbers are vital for both the survival and death of the disease



(a) High birth-rate ($e = 0.05$), low death-rate, both in general, but also for the infected. ($d = 0.001, d_I = 0.01$)

(b) low birth-rate ($e = 0.005$), low death-rate, both in general, but also for the infected ($d = 0.001, d_I = 0.01$)



(c) high birth-rate ($e = 0.05$), high general death-rate and very high death-rate for the infected.

(d) low birth-rate ($e = 0.006$), low general death-rate ($d = 0.002$), high death-rate for infected ($d_I = 0.10$)

Figure 5: Looking at different birth-, death-, infected death - rates.

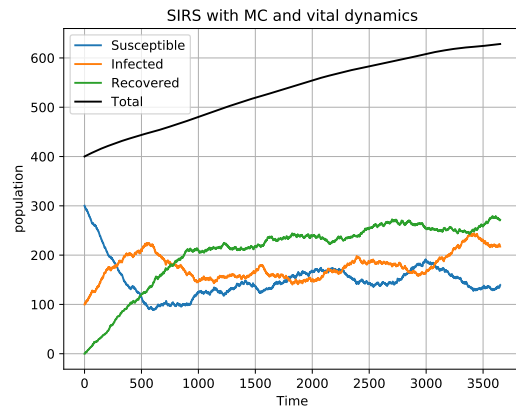
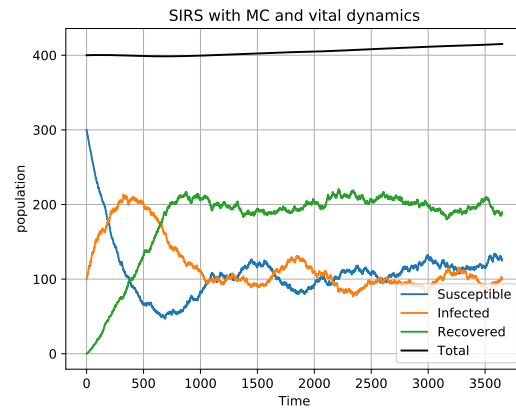
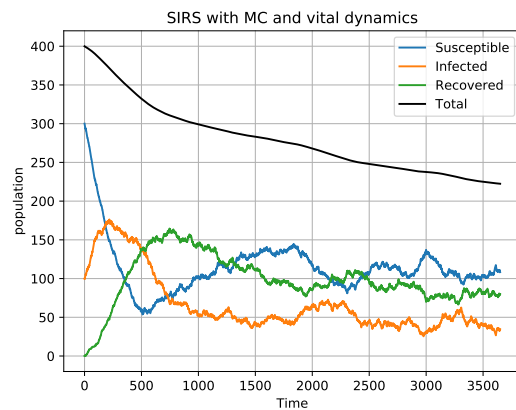
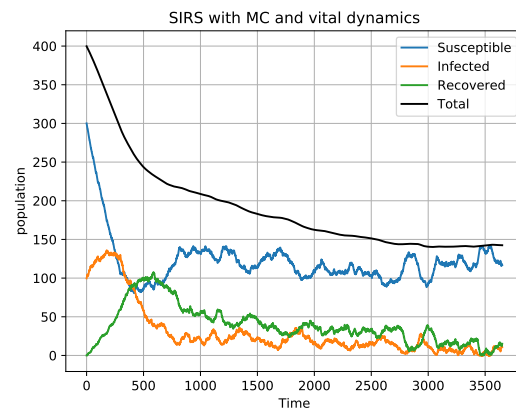
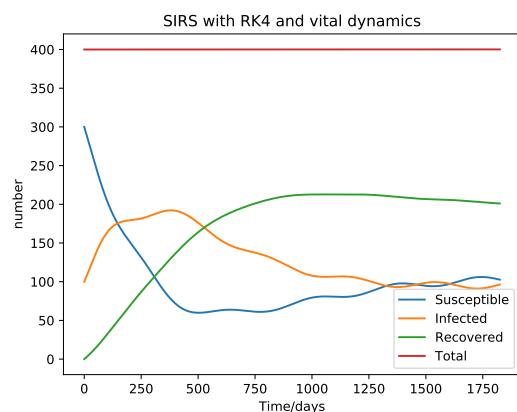
(a) $d = 0.0002, d_I = 0.0004, e = 0.0006$ (b) $d = 0.0002, d_I = 0.0001, e = 0.0006$ (c) $d = 0.0002, d_I = 0.001, e = 0.0006$ (d) $d = 0.0002, d_I = 0.003, e = 0.0006$

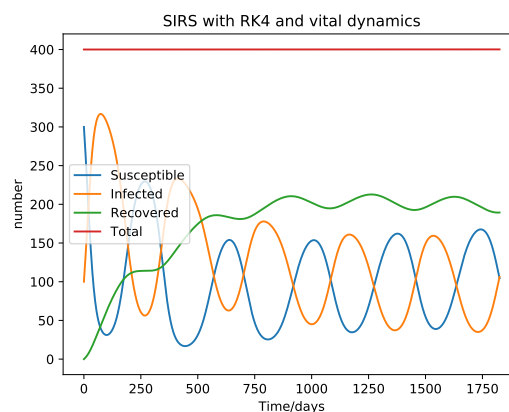
Figure 6: Main difference between the MC method and the RK4 is that dt was not normalized in the cod, so smaller numbers had a much bigger impact.

5.2.2 Seasonal Variation

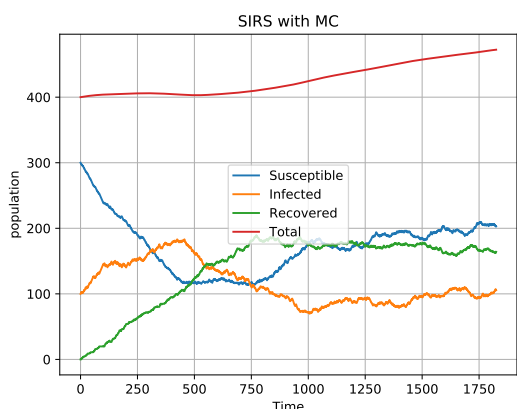
Here a Seasonal Variation is included 4.2.2, The value A represents the maximum deviation from the infection rate $a_0 = 4$.



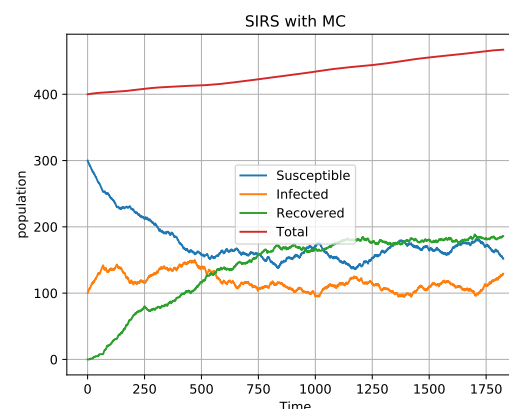
(a) RK4 $A = 1.2$



(b) RK4 $A = 2.0$



(c) MC $A = 1.2$



(d) MC $A = 2.0$

Figure 7: This figure shows the impact of seasonal variations on RK4 and MCS, notice how the seasonal variation impact is delayed in the MC scheme.

5.2.3 Vaccination

If we assume that vaccines work and the earth is round, than we can see what incredible tool vaccines are for managing diseases. These results shows how efficient vaccine in combination with seasonal variation can be, when you only vaccinate 10% of the population once per year.

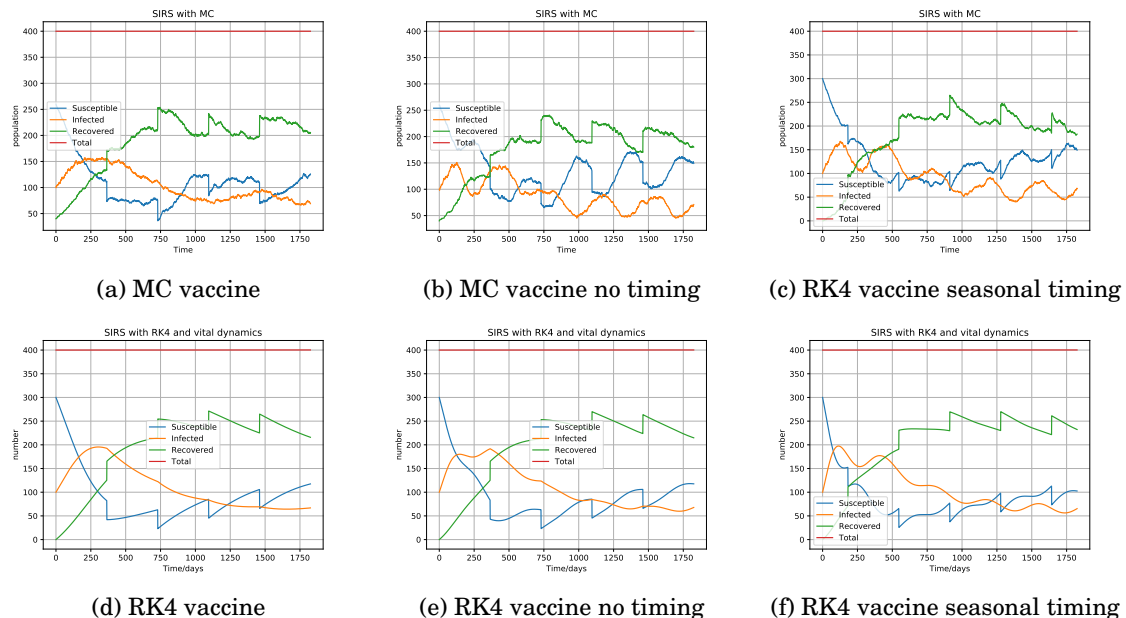


Figure 8: The SIRS model, figure a) and d) are clean SIRS models with the effect of a yearly vaccine, b) and e) include the seasonal variation of the disease, lastly in figure c) and f) we have timed the vaccine to be given right before the winter, where it is assumed to be at its worst. (at least in this model).

6 Conclusion

It seems that the SIRS model is a great system dynamics model for representing real diseases, in general a great foundation to build on for predicting disease, in this case, and how they establish themselves within the population. We have also seen that it is very versatile, with a few improvements tested out, and combined in this report. It was shown that a Monte Carlo simulation were a great way of approaching the differential equations in a more realistic manner, if given a large enough dataset.

7 References:

note: bibtex did not want to work with me on this project.

- Yang, Wan, Alicia Karspeck, and Jeffrey Shaman. "Comparison of filtering methods for the modeling and retrospective forecasting of influenza epidemics.PLoS computational biology 10.4 (2014): e1003583.
- Zaman, Gul, Yong Han Kang, and Il Hyo Jung. Stability analysis and optimal vaccination of an SIR epidemic model.BioSystems 93.3 (2008): 240-249.
- Ramachandran, Prajit, Barret Zoph, and Quoc V. Le. Searching for activation functions."(2018).