

An Agent-Based Model of COVID-19 Diffusion to Plan and Evaluate Intervention Policies

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Abstract A model of interacting agents, following plausible behavioral rules into a world where the Covid-19 epidemic is affecting the actions of everyone. The model works with (i) infected agents categorized as symptomatic or asymptomatic and (ii) the places of contagion specified in a detailed way. The infection transmission is related to three factors: the characteristics of both the infected person and the susceptible one, plus those of the space in which contact occurs. The model includes the structural data of Piedmont, an Italian region, but we can easily calibrate it for other areas. The micro-based structure of the model allows factual, counterfactual, and conditional simulations to investigate both the spontaneous or controlled development of the epidemic.

The model is generative of complex epidemic dynamics emerging from the consequences of agents' actions and interactions, with high variability in outcomes, but frequently with a stunning realistic reproduction of the successive contagion waves in the reference region. There is also an inverse generative side of the model, coming

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from constructing a meta-agent optimizing the vaccine distribution among people groups—characterized by age, fragility, work conditions—to minimize the number of symptomatic people, using genetic algorithms.

1 A quick introduction to our agent-based epidemic model

The starting point is a compartmental model with Susceptible, Infected, and Recovered people (S.I.R.), but adding both a more detailed breakdown of the subjects involved in the contagion process [1] and a multi-scale framework to account for the interaction at different dimensional, and spatial levels [2]. From the virus micro-level, we move to individuals and up to the collective behavior of the population.

Following [3], we know that the analysis based on the assumption of heterogeneity strongly differs from S.I.R. compartmental structures modeled by differential equations. Their work ponders when it is better to use agent-based models and when it would be better to use differential equation models. Differential equation models assume homogeneity and perfect mixing of characteristics within compartments, while agent-based models can capture heterogeneity in agent attributes and the structure of their interactions. We follow the second approach.

- The model works with:
 - i infected agents categorized as symptomatic or asymptomatic and
 - ii the places of contagion specified in a detailed way, thanks to agent-based modeling capabilities.
- The infection transmission is related to three factors: the infected person's characteristics and those of the susceptible one, plus those of the space in which a contact occurs.

Finally, we subscribe the call of [4] to «cover the full behavioural and social complexity of societies under pandemic crisis» and we work arguing that «the study of collective behavior must rise to a “crisis disciplin” just as medicine, conservation, and climate science have, with a focus on providing actionable insight to policymakers and regulators for the stewardship of social systems», as in [5].

Now a look at the structure of the whole presentation.

In Section 1.1, we discuss models and specifically agent-based models; in Section 1.2, the biochemical support to agents' intrinsic susceptibility construction; in Section 1.3, the structure of the model, with the daily sequence of the agents' actions. Section 2 introduces a detailed description of the internal model mechanisms, with conditional actions in Section 2.1, parameters in Section 2.2 and agents' interaction in Section 2.3.

A technique for contagion representation is reported in Section 3. Then we explore simulation cases in Section 4, building several batches of runs and comparing extreme situations in Sections 4.1.

Section 4.2 introduces the actual epidemic data of the reference region. With those data, we verify factual and counterfactual analyses in Section 5. Considering the possibility of calculating infection indicators without delays (Section 5.4), we experiment with the effect of adopting the control measure with 20 days of anticipation (Section 5.5). In Section 5.6 we verify another counterfactual policy, that of concentrating the efforts uniquely in defense of fragile persons. Section recap 6.1 summarizes the results.

The final application of the model is dedicated to a planning exercise on vaccination campaigns (Section 7). We introduce an analysis of the vaccine mechanism in the perspective of our model (Section 7.1) and using genetic algorithms to optimize the behavior of a meta-agent deciding the sequence of the vaccinations (Section 7.7). Finally, a specific, realistic case is deeply analyzed in Section 7.3.

1.1 Why model? Why agents? Why another model?

Why another model, and most of all, why model? With [6]:

The choice (...) is not whether to build models; it's whether to build explicit ones. In explicit models, assumptions are laid out in detail, so we can study exactly what they entail. On these assumptions, this sort of thing happens. When you alter the assumptions that is what happens. By writing explicit models, you let others replicate your results.

With even more strength:

I am always amused when these same people challenge me with the question, "Can you validate your model?" The appropriate retort, of course, is, "Can you validate yours?" At least I can write mine down so that it can, in principle, be calibrated to data, if that is what you mean by "validate" a term I assiduously avoid.

To reply to "why agents?", with [7] we define in short what an agent-based model is:

An agent-based model consists of individual agents, commonly implemented in software as objects. Agent objects have states and rules of behavior. Running such a model simply amounts to instantiating an agent population, letting the agents interact, and monitoring what happens. That is, executing the model—spinning it forward in time—is all that is necessary in order to "solve" it.

More in detail:

There are, ostensibly, several advantages of agent-based computational modeling over conventional mathematical theorizing. First, [...] it is easy to limit agent rationality in agent-based computational models. Second, even if one wishes to use completely rational agents, it is a trivial matter to make agents heterogeneous in agent-based models. One simply instantiates a population having some distribution of initial states, e.g., preferences. That is, there is no need to appeal to representative agents. [...] Finally, in most social processes either physical space or social networks matter. These are difficult to account for mathematically except in highly stylized ways. However, in agent-based models it is usually quite easy to have the agent interactions mediated by space or networks or both.

And now, "why another?" As a commitment to our creativity, using our knowledge to understand what is happening. Indeed, with arbitrariness: it is up to others and time to judge. Modeling the Covid-19 pandemic requires a scenario and the actors.

As any model, also this one is based on assumptions: time will tell whether these were reasonable hypotheses. Modeling the Covid-19 pandemic requires a scenario and the actors. As in a theater play, the author defines the roles of the actors and the environment. The characters are not real, they are prebuilt by the author, and they act according to their peculiar constraints. If the play is successful, it will run for a long time, even centuries. If not, we will rapidly forget it. Shakespeare's Hamlet is still playing after centuries, even if the characters and the plot are entirely imaginary. The same holds for our simulations: we are the authors, we arbitrarily define the characters, we force them to act again and again in different scenarios. However, in our model, the micro-micro assumptions are not arbitrary but based on scientific hypotheses at the molecular level, the micro agents' behaviors are modeled in an explicit and realistic way. In both plays and simulations, we compress the time: a whole life to two or three hours on the stage. In a few seconds, we run the Covid-19 pandemic spread in a given regional area.

1.2 Here a Gianpiero's section on the biochemical of the contagion

1.3 Our model

With our model, we move from a macro compartmental vision to a meso and micro-analysis capability. Its main characteristics are:

- scalability: we take in account the interactions between virus and molecules inside the host, the interactions between individuals in more or less restricted contexts, the movement between different environments (home, school, workplace, open spaces, shops); the movements occur in different parts of the daily life, as in [8];
in detail, the scales are:
 - *micro*, with the internal biochemical mechanism involved in reacting to the virus, as in [9], from where we derive the critical importance assigned to an individual attribute of intrinsic susceptibility related to the age and previous morbidity episodes; the model indeed incorporates the medical insights and consistent perspectives of one of its co-authors, former full professor of clinical biochemistry, signing also the quoted article; a comment on Lancet [10] consistently signals the syndemic character of the current event: «Two categories of disease are interacting within specific populations—infection with severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and an array of non-communicable diseases (NCDs)»;
 - *meso*, with the open and closed contexts where the agents behave, as reported above;

- *macro*, with the emergent effects of the actions of the agents;
- *granularity*: at any level, the interactions are partially random and therefore the final results will always reflect the sum of the randomness at the different levels; changing the constraints at different levels and running multiple simulations should allow the identification of the most critical points, where to focus the intervention.

Summing up, S.I.s.a.R. [11] is an agent-based model designed to reproduce the diffusion of the COVID-19 using agent-based modeling in NetLogo [12]. We have Susceptible, Infected, symptomatic, asymptomatic, and Recovered people: hence the name S.I.s.a.R. The model works on the structural data of Piedmont, an Italian region, but we can quite easily calibrate it for other areas. It can reproduce the events following a realistic calendar (national or local government decisions, as in Section 2.2), via its script interpreter. The model is online at <https://terna.to.it/simul/SIsaR.html>, from where it is also possible to run the code without installation. Into the *Info* sheet of the model, we have more than 20 pages of Supporting Information about both the structure and the calibration of the model.

The micro-based structure of the model allows factual, counterfactual, and conditional simulations to investigate both the spontaneous or controlled development of the epidemic. Examples of counterfactual situations are those considering:

- i different timing in the adoption of the non-pharmaceutical containment measures;
- ii alternative strategies focusing exclusively on the defense of fragile people.

The model generates complex epidemic dynamics, emerging from the consequences of agents' actions and interactions, with high variability in outcomes, but frequently with a stunning realistic reproduction of the contagion waves that occurred in the reference region.

We take charge of the variability of the epidemic paths within the simulation, running batches of executions with 10,000 occurrences for each experiment.

Following [13], the AI and inverse generative side of the model comes from constructing a meta-agent optimizing the vaccine distribution among people groups—characterized by age, fragility, work conditions—to minimize the number of symptomatic people (deceased persons come from there).

We can characterize the action of the planner both:

- i introducing ex-ante rules following “plain” or “wise” strategies that we imagine as observers or
- ii evolving those strategies via the application of a genetic algorithm.

The genome is a matrix of vaccination quotas by people groups, with their time range of adoption.

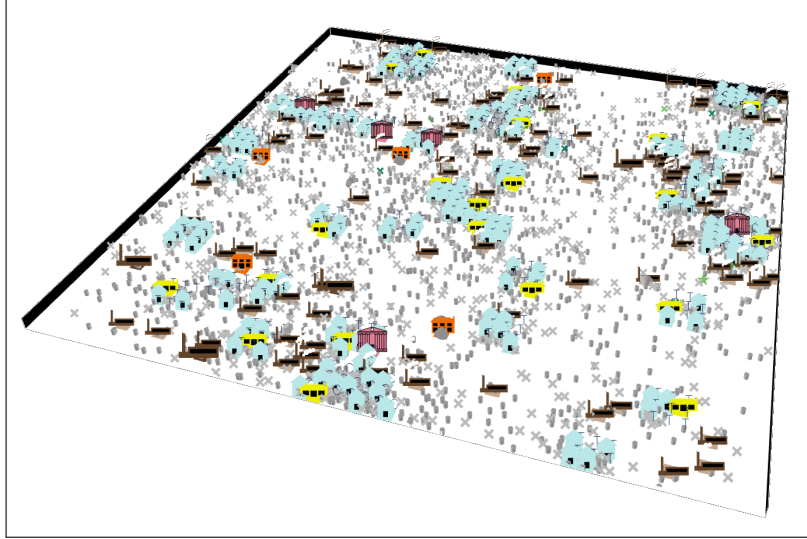


Fig. 1: A 3D representation of the model world

2 How S.I.s.a.R. works

We have two initial infected individuals in a population of 4350 individuals, on a scale of 1:1000 with Piedmont. The size of the initial infected group is out of scale: it is the smallest number, ensuring the epidemic's activation in a substantial number of cases. Initial infected people bypass the incubation period. For implausibility reasons, we never choose initial infected people among persons in nursing homes or hospitals. The presence of agents in close spaces—such as classrooms, factories, homes, hospitals, nursing homes—is made with realistic numbers, not to be read in scale: e.g., a classroom contains 25 students, a home two persons, large factories up to 150 employees, small ones up to 15, etc.; the movements occur in different parts of the daily life, as in [8].

In Fig. 1 we have a 3D representation of the model world—as one of the possible random maps that the simulation generates— with persons in gray, houses in cyan, nursing homes in orange, hospitals in orange, schools in yellow, factories (with shops and offices) in brown. Persons have a cylinder as shape, if regular or robust (young); a capital X if fragile; temporary their colors can be: red, if symptomatic; violet, if asymptomatic; turquoise, if symptomatic recovered; green, if asymptomatic recovered.

We use random maps to have a neutral representation of the structure of the space when doing the batches of 10,000 repetitions of the simulation.

We can set:

- min and max duration of the individual infection;
- the length of the incubation interval;

- the critical distance, i.e., the radius of the possibility of infection in open air, with a given probability;
- the correction of that probability, due to the personal characteristics of both active and the passive agents; passive agents, as receivers, can be robust (young), regular, fragile, and extra fragile.

We have two main types of contagion: (a) within a radius, for people moving around, also if only temporary present in a house/factory/nursing home/hospital (in schools we only have students and teachers); (b) in a given space (room or apartment) for people resident in their home or in a hospital or in a nursing home or being in school or in a working environment.

People in hospitals and nursing homes can be infected in two ways: (a) and (b). Instead, while people are at school, they can only receive the disease from people in the same classroom, where only teachers and students are present, so this is a third infection mechanism (c). In all cases, the personal characteristics of the recipients are decisive.

We remark that workplaces are open to all persons, as clients, vendors, suppliers, external workers can go there. In contrast, schools are mainly reserved for students and school operators and are less affected by contact with other types of agents.

All agents have their home, inside a city, or a town. The agents also have usual places (UPs) where they act and interact, moving around. These positions can be interpreted as free time elective places. When we activate the schools, students and teachers have both UPs and schools; healthcare operators have both UPs and hospitals or nursing homes; finally, workers have both UPs and working places. In each day (or tick of the model), we simulated realistic sequences of actions.

Fig. 2 describes what happens during every *day* in our simulated world, with each day as a sequence of several actions.

2.1 Conditional actions

Agents' movements in space, to go to work, school, and other UPs are subject to two interrelated general conditions.

- I Symptomatic persons are at home or in a hospital or a nursing home and do not move.
- II People not constrained by *condition I* can move if (primary rule) there are no general limitations (e.g., lockdown) *OR* one of the following specific sub-conditions applies:
 - a. agents who are hospital healthcare operators or nursing home healthcare operators;
 - b. all people, according to the probability of moving of the whole non-symptomatic agents (2.2, iii);
 - c. regular people, according to the probability of moving of the regular non-symptomatic agents (2.2, iv);

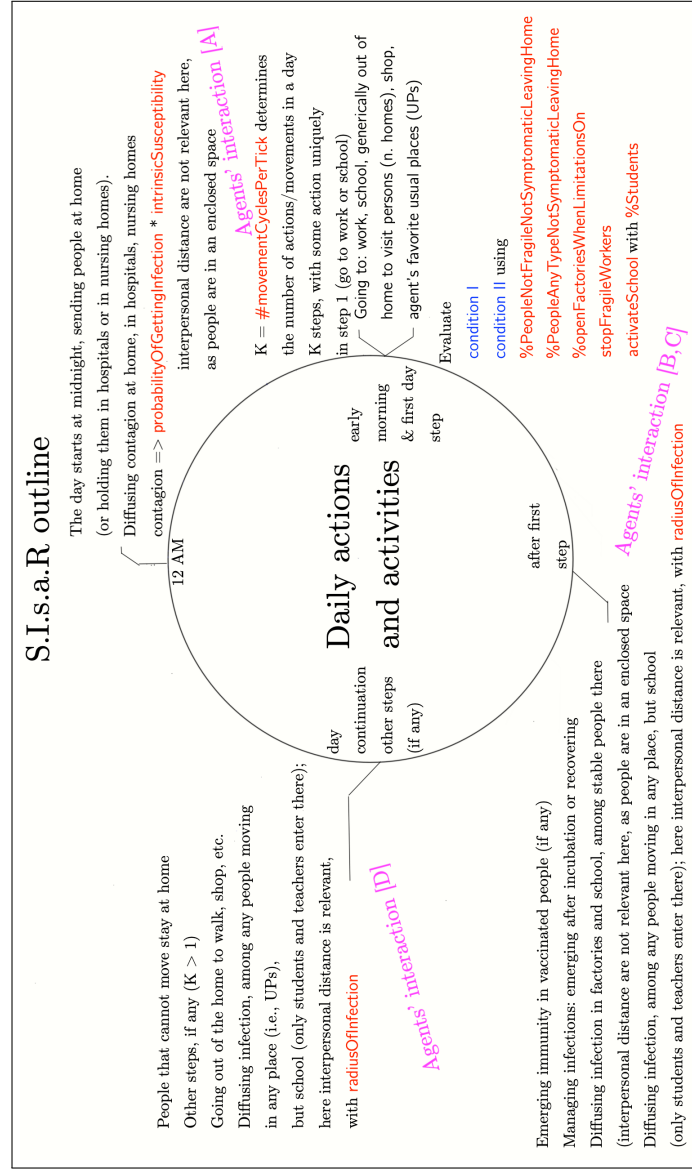


Fig. 2: A day in the simulation, with N repetitions where N is the duration of a specific outbreak; look at: Section 2.1 for the rules of the conditional actions; Section 2.2 for the parameter definitions; and Section 2.3 for details on the agent interactions

- d. workers, if all the factories are open or it is open their workplace (2.2, v);
- e. teachers, if the schools are open (2.2, vi);

- f. students, if the schools are open, but with a possible quota limitation (2.2, vii).

2.2 Parameter definition

We define here the parameters of Fig. 2, also with the short names used in program scripts, in round brackets. The values of the parameters are reported in detail in Appendix 1 - Parameter values (Section 9).

- i *probabilityOfGettingInfection* (**prob**) is the base probability of getting infected, to be multiplied by the *intrinsicSusceptibility* factor (ii); it is activated if the subject is within a circle of radius (viii) with an infected person; values at (9, i);
- ii the *intrinsicSusceptibility* is based on *intrinsicSusceptibilityFactor* set to 5 in Eq. 1

$$\text{intrinsicSusceptibility} = \text{intrinsicSusceptibilityFactor}^{\text{groupFragility}} \quad (1)$$

with *groupFragility* exponent set to:

- 1 for extra-fragile persons,
- 0 for fragile persons,
- 1 for regular persons,
- 2 young people from 0 to 24 years old;

- iii *%PeopleAnyTypeNotSymptomaticLeavingHome* (**%PeopleAny**) determines, in a probabilistic way, the number of people of any kind going around in case of limitations/lockdown; the limitations operate only if the lockdown is on (into our simulated world, from day 20); values at (9, iii);
 - iv *%PeopleNotFragileNotSymptomaticLeavingHome* (**%PeopleNot**) determines, in a probabilistic way, the number of regular people going around in case of limitations/lockdown; as above, the limitations operate only if the lockdown is on (into our simulated world, from day 20); values at (9, iv);
- the parameters iii and iv in some time interval change very frequently, reproducing into the model the uncertainty of the decisions that were happening in the real world in the same periods;

NB, the parameters iii and iv produce independent effects, as in the following examples: (a) the activation of *%PeopleAny* at 31, 0 and, simultaneously, of *%PeopleNot* at 31, 80, means that people had to stay home on that day, but people specifically not fragile could go out in 80% of the cases; (b) *%PeopleAny* at 339, 80 and, simultaneously, *%PeopleNot* at 339, 100 means that fragile and not fragile persons cannot always go around, but only in the 80% of the cases; instead, considering uniquely non-fragile persons they are free to go out; the construction is an attempt to reproduce a fuzzy situation; in future versions of the model, we will define the quotas more straightforwardly:

- %FragilePeopleNotSymptomaticLeavingHome;
 - %NotFragilePeopleNotSymptomaticLeavingHome;
- v *%openFactoriesWhenLimitationsOn* (%Fac) determines, in a probabilistic way, what factories (small and large industries, commercial surfaces, private and government offices) are open when limitations are on; if the factory of a worker is open, the subject can go to work, avoiding restrictions (but uniquely in the first step of activity of each day); values at (9, v);
- vi *stopFragileWorkers* (sFW) is *off* (set to 0) by default; if *on* (set to 1), fragile workers (i.e., people fragile due to prior illnesses) can move out of their homes following the iii and iv parameters, but cannot go to work; the regular case is that the workers (fragile or regular) can go to their factory (if open) also when limitations are on; values at (9, vi);
alternatively, we also have the *fragileWorkersAtHome* parameter; if *on* (set to 1) the total of the workers is unchanged, but the workers are all regular; we can activate this counterfactual operation uniquely at the beginning of the simulation;
- vii when *activateSchools* (aSch) is *on* (set to 1), teachers and students go to school avoiding restrictions (but uniquely in the first step of activity of each day); *%Students* (%St) limits to its value the quota of the students moving to school; the residual part is following the lessons from home; values at (9, vii);
- viii following *radiusOfInfection* (radius), the effect of the contagion—outside enclosed spaces, or there, but for temporary presences—is possible within that distance; values at (9, viii).

2.3 Agents' interaction

We underline that our simulation tool is not based on micro-simulation sequences, calculating the contagion agent by agent, on the base of their characteristics and ex-ante probabilities. It implements a true agent-based simulation, with the agents acting and, most of all, interacting, thus generating continuously contagion situations.

Each run creates a population with expected characteristics, but also with random specifications, to assure the heterogeneity in agents. The daily choices of the agents are partially randomized, to reproduce real-life variability.

Contagions arise from agents' interactions, in four situations, as specified in Fig. 2:

- A - in houses (at night), hospitals, nursing homes;
- B - in schools, workplaces in general, among people stable there;
- C - in the places above (excluding schools) by people temporary there and in open spaces (UPs above);
- D - interactions mainly in open spaces (UPs above).

3 Contagion representation

We introduce a tool that allows analyzing the contagions' sequences in simulated epidemics and identifying the places where they occur.

- We represent each infected agent as a horizontal segment (from the starting date to the final date of the infection) with vertical connections to other agents if they receive the disease.
- We represent the new infected agents via further segments at an upper level.
- We display multiple information using three elements.
 - Colors in horizontal segments (areas of the infections): black for unknown places, gray for open spaces, cyan for houses, orange for nursing homes, pink for hospitals, yellow for schools, brown for factories, with shops and offices.
 - Vertical connecting segments keep the same color of the horizontal generating one.
 - Line thickness; proportional to fragility.
 - Styles: dotted lines for incubation, dashed lines for asymptomatic subjects, solid lines of symptomatic ones.
- This graphical presentation enables understanding at a glance how an epidemic episode is developing. In this way, it is easier to reason about countermeasures and, thus, to develop intervention policies.

At <https://github.com/terna/contagionSequence> we have the calculation tool, *sequentialRecords.ipynb*, generating these sequences.

Fig. 3 is useful as an example. We start with two agents from the outside, with black as the color code (unknown place). The first one is young—as reported by the thickness of the segment, starting at day 0 and finishing at day 22—and asymptomatic (dashed line); it infects no one. The second one—regular, as reported by the thickness of the segment, starting at day 0 and finishing at day 15—is asymptomatic (dashed line) and infects four agents on day 2. All the four infected agents received the infection at work (brown color) and turn to be asymptomatic after the days of incubation (dotted line); the first and the fourth are regular agents; the second and the third are fragile ones.

Continuing the analysis: on day 3, the second agent infects three other agents (at home, at work, at work) [...]; on day 13, agent number five infects seven regular agents at work and an extra-fragile one in a nursing home (orange color), etc.

If a vertical segment changes its color, we have an agent in an upper infecting someone on the same day of the infection transmitted by an agent in a lower row, so we lose some information.

In Fig. 4 we see an example of an epidemic where non-pharmaceutical containment measures are in adoption: a first wave shows an interlaced effect of contagions at home, in nursing homes, and at work. After a phase in which contagions develop mainly at home, a skinny bridge connects the first wave to a second one, restarting from workplaces. The thickness of the *snake* of the contagions measures the stock

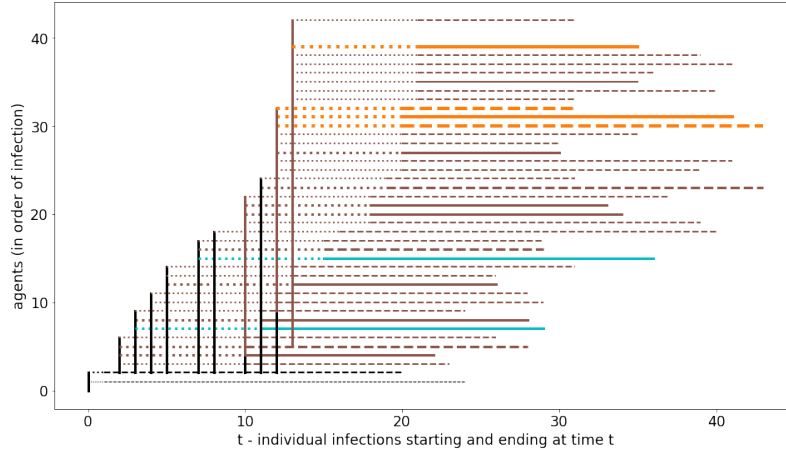


Fig. 3: A case with containment measures, first 40 infections: workplaces (brown) and nursing homes (orange) interweaving

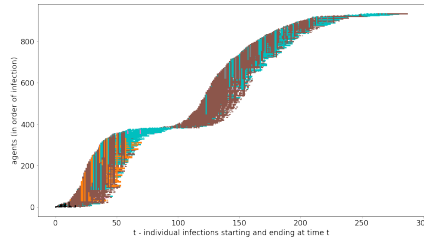


Fig. 4: A Case with containment measures, the whole epidemics: workplaces (brown) and nursing homes (orange) and then houses (cyan), with a bridge connecting two waves

on infects agents on a given date; the slope reports the speediness of the epidemic development; the vertical coordinate reports the cumulative number of infected people.

In Appendix 2 - A gallery of contagion sequences (Section 10) we have several examples of contagion sequences.

4 Exploring scenarios with simulation batches

The sequences described in Section 3 offer mainly suggestions and interpretations. To explore systematically the introduction of factual, counterfactual, and prospective interventions to control the spread of the contagions, we need to analyze batches

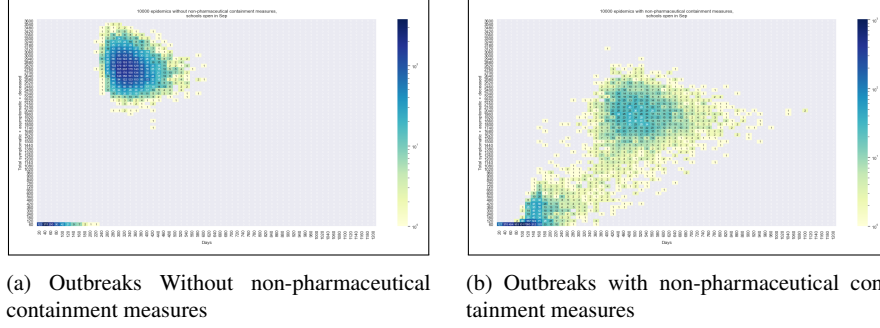


Fig. 5: Starting our analyses: 10,000 epidemics in Piedmont

of simulations. In this perspective, each simulation run—whose length coincides with the disappearance of symptomatic or asymptomatic contagion cases—is a datum in a scenario of time and outcome variability. Consequently, we need to represent compactly the results emerging from batches of repetitions to compare the consequences of each batch’s basic assumptions.

For this purpose, we used blocks of ten thousand repetitions. Besides summarizing the results with the usual statistical indicators, we adopt the technique of the heat-maps. With [14], we develop a search for comparative analyzes, not for forecasting. This consideration is consistent with the enormous standard deviation values that are intrinsic to the specific reality.

At https://github.com/terna/readIsaR_BatchResults we have the codes producing the maps of the batches. A heat-map is a double histogram: in our application, it displays each simulated epidemic’s duration in the x axis and the count of the symptomatic, asymptomatic, and deceased agents in the y axis (on a scale of 1:1000). The cells contain the number of epidemics of each specific duration and outcome, reported via a logarithmic color scale.

4.1 Epidemics without and with control

As a starting point, we compare the situations represented in Figs. 5a and 5b. In Fig. 5a the heat-map reports the distribution in duration and infection causation of 10,000 simulated outbreaks left to spread without any control; coherently, with the school always open. The numbers in Table 1 are scary. The concentration of the cases in the heat-map shows that, except a few instances spontaneously concluding in a short period (left bottom corner), produces a heavy *cloud* of cases lasting one year or one year and a half, hitting (as symptomatic, asymptomatic and deceased) from 2,000 to 3,500 persons on a total of 4,350 in the region (scale of 1:1000).

In Fig. 5b and the related Table 2 we report a similar simulation batch of 10,000 runs of the model, but with the adoption of the basic non-pharmaceutical containment

	(000) symptomatic	totalInfected&Deceased	duration
mean	969.46	2500.45	303.10
std	308.80	802.88	93.50

Table 1: Mean values and standard deviations in Fig. 5a cases

	(000) symptomatic	totalInfected&Deceased	duration
mean	344.22	851.64	277.93
std	368.49	916.41	213.48

Table 2: Mean values and standard deviations in Fig. 5b cases

measures, registered in the values of the parameters in Appendix 1 - Parameter values (Section 9). A calendar is at <https://terna.to.it/simul/calendario092.pdf>, and the model—version 0.9.6— is updated until April 2021. The results are dramatically different, showing the efficacy of the containment measures.

4.2 Actual data

The critical points for our experiments with the simulation model of the epidemic in Piedmont are Summer and Fall 2020, as in Fig. 6a, where we have the time series of the first part of Piedmont’s actual epidemic. The blue line represents the cumulative number of infected persons. Initially, only symptomatic cases were accounted for, but after the 2020 Summer, with more generalized tests also asymptomatic patients are included.

Explanation:

- from <http://www.protezionecivile.it/web/guest/department>, the Italian Civil Protection Department web site, we find at <https://github.com/pcm-dpc/COVID-1>, i.e., the repository of regional data;
- initially, only symptomatic cases were accounted for, but after the 2020 Summer, with more generalized tests, asymptomatic patients are included;
- we had data about symptomatic infected people in the first wave, but from October 2020, data are mixed: in the above *git* repository, in October and November, we had “Positive cases emerged from clinical activity”, unfortunately then reported as “No longer populated” (from the end of November, our observation) and “Positive cases emerging from surveys and tests, planned at national or regional level”, again “No longer populated” (from the end of November, our observation);
- as a consequence, the subdivision between symptomatic and asymptomatic cases is no longer possible.

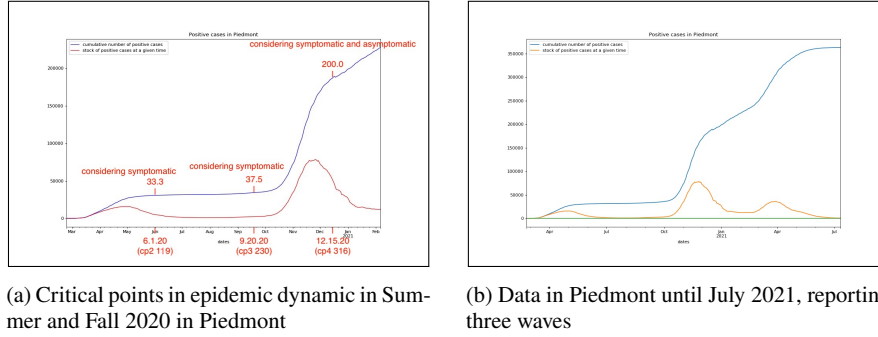


Fig. 6: Actual data

Considering the dynamic of the data in Fig. 6a, we search within the simulation batch for cases with both:

- i numbers of infected persons quite similar at cp2 and at cp3; besides, numbers not too different from those of the figure; (with *cp*, we indicate the internal check points of the simulation program. In Fig. 6a we also report the number of days from the beginning of the epidemic for each check point);
- ii the number of infected persons at cp4 has to be significantly greater than those at the previous check point.

In a lot of cases, epidemics satisfying condition (i) fail to match condition (ii); both the situations happen only in less than the 1.5% of the instances in a batch of ten thousand epidemic. We can conclude that the second wave registered in Piedmont after the Summer stability is due to new infected agents coming from outside and restarting the contagion process.

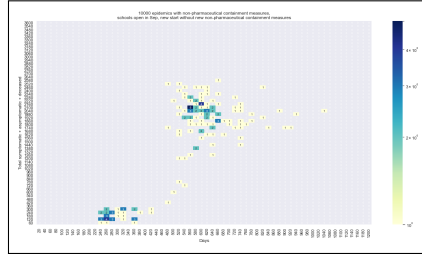
Other critical points in our analysis are the day on which the vaccination campaign starts, 373 of the simulation (Feb. 12th, 2021), and the day of the effectiveness of the initial vaccinations, 40 days later, day 413 (Mar. 22nd, 2021). At those dates, within the simulations, we can find either the presence of many infected agents or of few ones, as effectively was the situation in Piedmont.

As documentation, in Fig. 6b the time series covering the whole period.

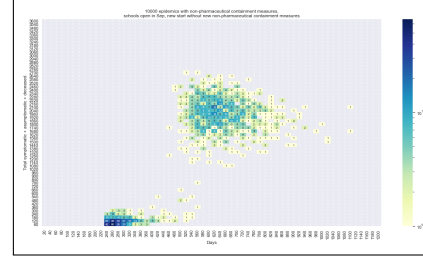
5 Factual and counterfactual analyses

In Fig. 7 we collect the heat-maps of the experiments:

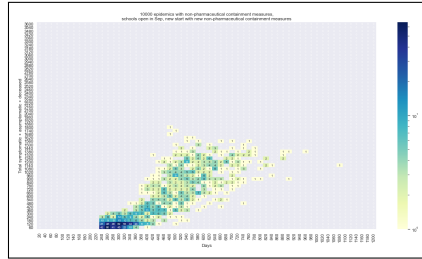
- observing the emergence of spontaneous second waves, in the absence of specific control measures (Sections 5.1);
- causing the emergence of the second wave through infections from outside, again in the absence of specific control measures (Section 5.2);



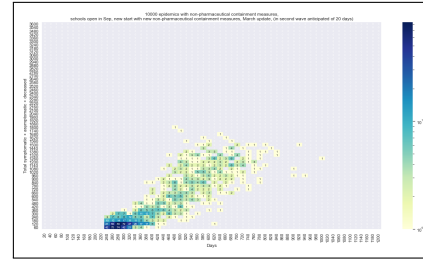
(a) First wave with non-pharmaceutical containment measures, spontaneous second wave, without specific measures



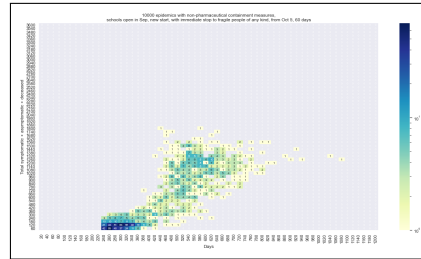
(b) First wave with non-pharmaceutical containment measures, forcing the second wave, without specific measures



(c) First wave with non-pharmaceutical containment measures, forcing the second wave, with new specific non-ph. containment measures



(d) First wave with non-pharmaceutical containment measures, forcing the second wave, with new specific non-ph. containment measures, acting 20 days in advance



(e) First wave with non-ph. containment measures, forcing the second wave; in second wave, uniquely stopping fragile people, including fragile workers

Fig. 7: Heat-maps of the factual and counterfactual analyses

- causing the emergence of the second wave through infections from outside, in the presence of specific control measures (Section 5.3);
- reproducing the case of Section 5.3, anticipating by twenty days the start and end of all control measures (Section 5.5);
- reproducing the case of Section 5.3, limiting the control measures the fragile workers and fragile people in general (Section 5.6).

(1000)	Jun 1, 20		Sep 20, 20		Dec 15, 20		Feb 1, 21		May 1, 21		Dec 15, 20 to end		
cum. v.	sym.	all	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	days
count	170.0	170.0	170.0	170.0	140.0	140.0	131.0	131.0	128.0	128.0	140.0	140.0	140.0
mean	37.9	100.2	60.4	159.3	248.4	648.7	432.2	1109.5	656.3	1655.5	701.1	1757.9	594.2
std	16.4	61.0	19.6	71.7	167.4	424.3	220.4	538.4	215.4	513.3	246.4	599.7	118.9

Table 3: Spontaneous second wave, without specific measures

5.1 Spontaneous second wave, without specific containment measures

In an initial plain batch of 10.000 runs of the Piedmont related model, we count only 140 cases of epidemics with both the absence of new contagions in Summer 2020 and their explosion in Fall, as in Fig. 6a.

We select, first of all, 170 cases in the following way. Considering June 1st, we choose the epidemics that have a number of symptomatic agents in the (10, 70] interval (their mean: 37.9) and, on September 20th, a number of symptomatic agents in the (20, 90] interval (their mean: 60.4). Due to the lack of data described in Section 4.2, to compare December 15th and September 20th situations, we use symptomatic plus asymptomatic agents' count. We observe the existence of 140 outbreaks with the required characteristics; the December mean of the infected agents is 648.7, sensibly larger than the actual value: ≈ 200.0 . We overestimate the reality being the long-lasting simulated outbreaks, the larger ones, and, most of all, having no containment measures operating in the simulations.

Fig. 7a and Table 3 show the outbreaks with similar cumulative numbers before and after the Summer 2020 pause (170 cases), with the second wave of 140 cases developing in the absence of containment measures.

The adverse cases are 140 out of 10,000, a very light ratio as the second wave occurred in the Fall. The transition to the third wave that we see in Fig. 6b is easy to explain, as the second wave never completely ended.

5.2 Second wave, new infections from outside, without specific containment measures

To generate a framework consistent with the presence of a second wave after a period of substantial inactivity of the epidemic, we introduced the occurrence of two cases of infected persons coming back from outside after Summer vacancies, conventionally on September 1st, 2020.

Now we have 1407 cases of epidemics with: (i) on June 1st, a number of symptomatic agents in the (10, 70] interval (their mean: 35.6) and (ii), on September 20th, a number of symptomatic agents in the (20, 90] interval (their mean: 40.0). As in Section 5.1, to compare December 15th and September 20th situations we use symptomatic plus asymptomatic agents' count. We observe the existence of 1044 outbreaks with the required characteristics; the December mean of the infected

(1000)	Jun 1, 20		Sep 20, 20		Dec 15, 20		Feb 1, 21		May 1, 21		Dec 15, 20 to end		
cum. v.	sym.	all	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	days
count	1407.0	1407.0	1407.0	1407.0	1044.0	1044.0	1005.0	1005.0	980.0	980.0	1044.0	1044.0	1044.0
mean	35.6	72.7	40.0	84.1	180.4	462.1	354.1	900.4	623.8	1563.3	726.6	1810.9	620.9
std	14.1	42.6	16.7	52.8	134.6	354.6	213.8	535.4	217.9	527.0	221.9	544.0	110.8

Table 4: Second wave, new infections from outside, without specific measures

(1000)	Jun 1, 20		Sep 20, 20		Dec 15, 20		Feb 1, 21		May 1, 21		Dec 15, 20 to end		
cum. v.	sym.	all	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	days
count	1407.0	1407.0	1407.0	1407.0	874.0	874.0	719.0	719.0	523.0	523.0	874.0	874.0	874.0
mean	35.6	72.7	40.0	84.1	<i>130.0</i>	<i>340.6</i>	<i>194.4</i>	<i>512.8</i>	295.7	791.2	252.7	666.4	494.1
std	14.1	42.6	16.7	52.8	83.9	232.6	104.1	276.9	119.1	300.6	156.8	416.4	122.7

Table 5: Second wave, new infections from outside, with new specific measures

agents is 462.1, again sensibly larger than the actual value: ≈ 200.0 . We overestimate the reality being the simulations run without the adoption of containment measures.

Both Fig. 7b and Table 4 show the outbreaks with similar cumulative numbers before and after the Summer 2020 pause (1407 cases), with the second wave of 1044 cases producing, in the absence of containment measures, a heavy cloud as that of Fig. 5a, with infected people of any kind in a range approximately from 1,500 to 2,800 realizations, with an equivalence, to the Piedmont scale, to 1,5-2,8 millions of subjects.

The number of cases is now sufficient to evaluate the effects of factual (Section 5.3 and counterfactual (Sections 5.5) and 5.6) simulation experiments.

5.3 Second wave, new infections from outside, with new specific containment measures

As in Section 5.2 we have 1407 cases of epidemics with: (i) on June 1st, a number of symptomatic agents in the (10, 70] interval (their mean: 35.6) and (ii), on September 20th, a number of symptomatic agents in the (20, 90] interval (their mean: 40.0). We use symptomatic plus asymptomatic agents' count December 15th and September 20th situations, recording 874 outbreaks with the required characteristics. The December mean of the infected agents is 340.6, closer to the actual value (≈ 200.0) due to the introduction into the simulation of specific control measures for the second wave. We always overestimate the reality because the surviving epidemics are the larger ones.

In Fig. 7c we see that the heavy cloud of the previous figure dissolved, and in Table 5 the figures in *italic* emphasize the effect of the containment interventions on the cases of epidemic continuation, in any case, reduced for the same motivation.

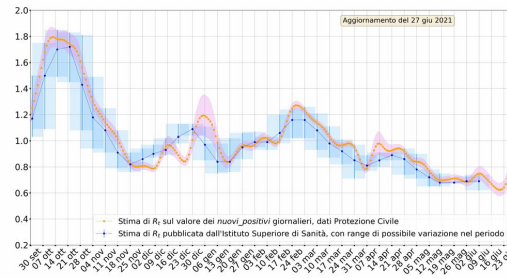


Fig. 8: In blue, the R_t values as reported by the Istituto Superiore di Sanità and in red the anticipated calculation published regularly, from the end of November 2020, at <https://mondoeconomico.eu> by Stefano Terna

5.4 Here a Stefano's section on calculating the infection indicator without delays

See Fig. 8 about . . .

Methodology:

https://github.com/tomorrowdata/COVID-19/blob/main/notebooks/Rt_on_italian_national_data.ipynb

5.5 Second wave, new infections from outside, with new specific containment measures, introduced 20 days in advance

The counterfactual situation is related to the start and end dates of the actions of containment, both occurring 20 days in advance, with a natural barrier set on October 5th, 2020. Before that date, the conditions to start second wave new policies were missing.

As in the last two sections, we have 1407 cases of epidemics alive at the critical dates of June 1st and September 20th, after a Summer interval characterized by a quiet phase. Considering December 15th and September 20th situations, the second wave epidemics are 769, again decreasing because the anticipated actions have eliminated some other cases. The December mean of the infected agents is 294.2, higher than the actual value (≈ 200.0). We always overestimate the mean of the epidemic effects, being the surviving epidemics the larger ones.

Comparing Fig. 7d and Fig. 7c the difference is not evident; instead, the italic figures, and must of all, the red bold ones, in Table 6 reports clearly the comparative advantage of this counterfactual experiment with respect to the values of Table 5.

(1000)	Jun 1, 20		Sep 20, 20		Dec 15, 20		Feb 1, 21		May 1, 21		Dec 15, 20 to end		
cum. v.	sym.	all	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	days
count	1407.0	1407.0	1407.0	1407.0	769.0	769.0	637.0	637.0	471.0	471.0	769.0	769.0	769.0
mean	35.6	72.7	40.0	84.1	112.2	294.2	172.0	467.9	276.5	748.6	248.9	663.4	499.3
std	14.1	42.6	16.7	52.8	66.8	188.4	91.5	251.3	112.9	286.9	158.0	417.5	124.1

Table 6: Second wave, new infections from outside, with new specific measure anticipation of -20 days

(1000)	Jun 1, 20		Sep 20, 20		Dec 15, 20		Feb 1, 21		May 1, 21		Dec 15, 20		
cum. v.	sym.	all	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	sympt.	totalInf.	to end sympt.	totalInf.	days
count	1407.0	1407.0	1407.0	1407.0	886.0	886.0	761.0	761.0	637.0	637.0	886.0	886.0	886.0
mean	35.6	72.7	40.0	84.1	128.1	326.3	211.0	555.1	323.3	862.1	301.1	792.3	515.5
std	14.1	42.6	16.7	52.8	89.6	234.2	118.1	306.7	126.4	315.9	170.7	450.2	116.9

Table 7: Second wave, new infections from outs., stop fragile people. 60 days from Oct. 5

5.6 Second wave, new infections from outside, with a unique intervention measure: stopping fragile people for 60 days

This second counterfactual experiment is based on an immediate stop to the circulation of fragile persons and specifically of fragile workers, plus isolating nursing homes and hospitals. Schools are always open in this experiment. The decision is activated on October 5th, 2020, as the second wave is becoming evident. In [15] we have important consideration suggesting the importance of taking into account fragility in a long-term fighting perspective against this kind of epidemics.

As in the last three sections, we have 1407 cases of epidemics alive at the critical dates of June 1st and September 20th, after a Summer interval characterized by a quiet phase. Considering December 15th and September 20th situations, the second wave epidemics are 886, lightly above the values of Section 5.3 and Section 5.5, but without locking the economic and the society as a whole. The December mean of the infected agents is 326.3, higher than the actual value (≈ 200.0) for the explained overestimation bias.

Comparing Fig. 7e and Fig. 7c the difference is not evident; instead, the italic figures, and must of all, the violet bold ones, in Table 7 reports the similarity of the effects of this counterfactual experiment with respect to the values of Table 5.

6 Cost-benefit analysis of the of interventions [Beppe's contribution]

draft, draft, draft, draft, draft

Scenarios		Dec 15, 20		Dec 15, 20 to end		days
		sympt.	totalInf.	sympt.	totalInf.	
no containment	count	140.0	140.0	140.0	140.0	140.0
in spontaneous	mean	248.4	648.7	701.1	1757.9	594.2
second wave	std	167.4	424.3	246.4	599.7	118.9
no containment	count	1044.0	1044.0	1044.0	1044.0	1044.0
in forced	mean	180.4	462.1	726.6	1810.9	620.9
second wave	std	134.6	354.6	221.9	544.0	110.8
basic containment	count	874.0	874.0	874.0	874.0	874.0
in forced	mean	130.0	340.6	252.7	666.4	494.1
second wave	std	83.9	232.6	156.8	416.4	122.7
-20 days cont.	count	769.0	769.0	769.0	769.0	769.0
in forced	mean	112.2	294.2	248.9	663.4	499.3
second wave	std	66.8	188.4	158.0	417.5	124.1
frag. subj. & workers control	count	886.0	886.0	886.0	886.0	886.0
in forced	mean	128.1	326.3	301.1	792.3	515.5
second wave	std	89.6	234.2	170.7	450.2	116.9

Table 8: Report of the key results, with count, mean, and std

Considering the different interventions of Section 5.3 and 5.6, we can evaluate

...

First of all, we can evaluate the costs for the attribution of sick pay to workers who are compulsorily absent from work, if they are not able to carry out remotely their work activities ...

We have also relevant social benefits, e.g., schooling, and economic benefits, as activities do not stop ...

An interesting reference is [16], where ... WORKS IN PROGRESS.

6.1 To recap

Write a nice global comment here

All waves, in second wave and at the end, with day count

Fig. 8

7 Planning vaccination campaigns

7.1 How vaccines work, (Alberto's section)

Vaccines are biological preparations made from killed or attenuated microorganisms, from viruses or from some of their components (antigens), or from substances they produce made safe by chemical (e.g.: formaldehyde) or heat treatment, while maintaining their immunogenic properties. (CITATION). Today, vaccines can be composed of proteins obtained by recombinant DNA techniques (CITATION) using genetic engineering techniques. They usually contain, in addition to the antigenic fraction, sterile water (or a saline-based physiological solution), as well as adjuvants, preservatives and stabilisers. Adjuvants are included in the vaccine in order to enhance the immune system response; preservatives are added to prevent contamination of the prpreparate by bacteria; stabilisers are introduced to increase the shelf life of the product and to maintain the properties of the vaccine during storage.

7.2 Planning a vaccination campaign using genetic algorithms (GAs), with non-pharmaceutical containment measures in action

We compare the effect of choosing the vaccination quotas via GAs with two predetermined strategies, considering three hypotheses (vaccinated people: still spread the contagion; do not spread the contagion; do it in the 50% of the case); we show here only the first case results.

Key dates:

- in the internal calendar of the model, day 373 is Feb. 12th, 2021, which is effectively the starting point of the vaccinations in the region;
- the day of the effectiveness of the initial vaccinations, 40 days later, is day 413 (Mar. 22nd, 2021).

(Technical detail. We simulate the vaccination campaigns with the GAs BehaviorSearch program, with a maximum step number of 865, related to the actual duration of case I. In the case of "y" use, the simulation never ends; when applying the GAs quotas ex-post, the simulation stops when no more infected people exist. In any case, if necessary, we have a hard finish button.)

7.2.1 Vaccination groups

We take into consideration seven groups in order of decreasing fragility but also considering the exposure to contagion:

g1 extra fragile people with three components;

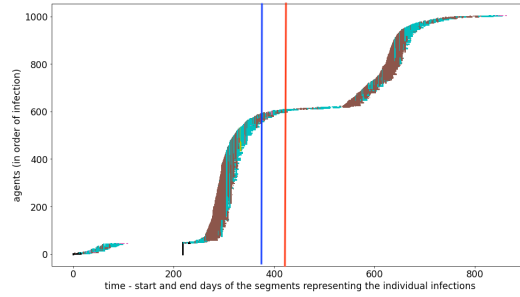


Fig. 9: Crucial dates: blue line for the starting point of the vaccination campaign and red line for the start of the effectiveness of the initial vaccinations

- due to intrinsic characteristics: people in nursing homes;
- due to risk exposure:
 - nursing homes operators;
 - healthcare operators;

g2 teachers;

g3 workers with medical fragility;

g4 regular workers;

g5 fragile people without special characteristics;

g6 regular people, not young, not worker, and not teacher;

g7 young people excluding special activity cases (a limited number in *g1*).

7.3 A specific realistic case

We analyze the case of Fig. 9 to . . .

7.4 Vaccination quotas, *plain* strategy

Considering the *plain* option adopted in Table 9 and remembering that the time-sequence in daily actions is the priority, we will primarily vaccinate the left column groups to move gradually to other columns: (*g1*) extra fragile people, (*g2*) teachers, (*g3*) fragile workers, (*g4*) regular workers, (*g5*) fragile people, (*g6*) regular people, (*g7*) young people.

In Tables 9 and 10 . . .

From day	Q. of vaccines (000)	<i>g1</i>	<i>g2</i>	<i>g3</i>	<i>g4</i>	<i>g5</i>	<i>g6</i>	<i>g7</i>
373	5	0.1	0.1	0.1	0.1	0.1	0.1	0.1
433	10	0.1	0.1	0.1	0.1	0.1	0.1	0.1
493	10	0.1	0.1	0.1	0.1	0.1	0.1	0.1
553	10	0.1	0.1	0.1	0.1	0.1	0.1	0.1
613	20	0.1	0.1	0.1	0.1	0.1	0.1	0.1
738	end							

Table 9: From the day of the first column, considering the quantity of the second column (000), the vaccination of each group follows the quota of the related columns

(000)	<i>g1</i>	<i>g2</i>	<i>g3</i>	<i>g4</i>	<i>g5</i>	<i>g6</i>	<i>g7</i>
Susc. at t = 0	133	84	240	1560	1179	254	900

Table 10: Susceptible persons at the beginning of the simulation

From day	Q. of vaccines (000)	<i>g1</i>	<i>g2</i>	<i>g3</i>	<i>g4</i>	<i>g5</i>	<i>g6</i>	<i>g7</i>
373	5	0.1	0.1	0.1	0.0	0.1	0.0	0.0
433	10	0.1	0.1	0.1	0.0	0.1	0.0	0.0
493	10	0.1	0.1	0.1	0.1	0.1	0.1	0.1
553	10	0.1	0.1	0.1	0.1	0.1	0.1	0.1
613	20	0.1	0.1	0.1	0.1	0.1	0.1	0.1
738	end							

Table 11: From the day of the first column, considering the quantity of the second column (000), the vaccination of each group follows the quota of the related columns

7.5 Vaccination quotas, *wise* strategy

In Tables 11 and 10 . . .

7.6 GAs quotas in the experiment, with vac. people spreading or not spreading the infection

In Tables 13 and 12 . . .

Groups: (*g1*) extra fragile people, (*g2*) teachers, (*g3*) fragile workers, (*g4*) regular workers, (*g5*) fragile people, (*g6*) regular people, (*g7*) young people.

(000)	<i>g1</i>	<i>g2</i>	<i>g3</i>	<i>g4</i>	<i>g5</i>	<i>g6</i>	<i>g7</i>
Susc. at t = 0	133	84	240	1560	1179	254	900
Susc. when vacc. starts	124	81	162	1234	1032	245	891

Table 12: Susceptible persons at the beginning of the simulation and when the vaccination campaign starts, day 373, Feb. 12th, 2021

From day	Q. of vaccines (000)	<i>g1</i>	<i>g2</i>	<i>g3</i>	<i>g4</i>	<i>g5</i>	<i>g6</i>	<i>g7</i>
373	5	0.01	0	0	0.79	0.18	0.38	0.19
433	10	0.94	0.06	0.32	0.54	0.19	0.83	0.5
493	10	0.97	0.97	0.74	0.79	0.2	0.14	0.52
553	10	0.98	0.83	0.02	0.39	0.99	0.04	0.48
613	20	0.52	0.01	0.83	0.6	1	0.27	0.9
738	end							

Table 13: GAs best strategy in case I, with *vaccinated people still spreading the infection*: from the day of the first column, considering the quantity of the second column, the vaccination of each group follows the quota of the related columns

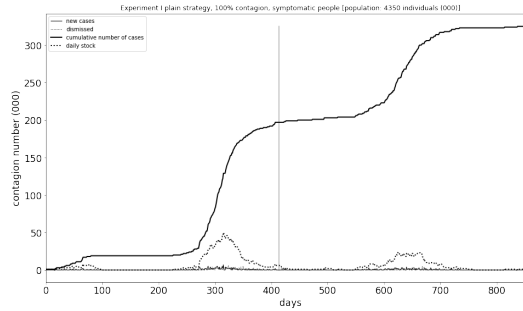
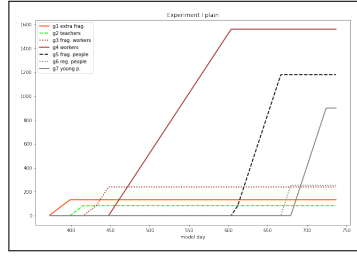


Fig. 10: Base symptomatic series; the vertical line is at day 413 is not relevant here

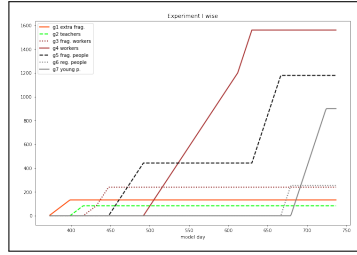
7.7 An experiment with GA

7.7.1 Time dynamics without vaccinations

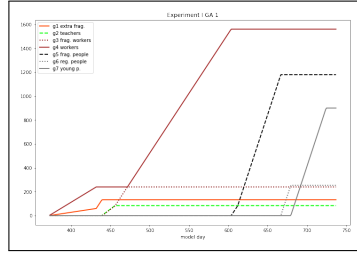
In Fig. 10 ...



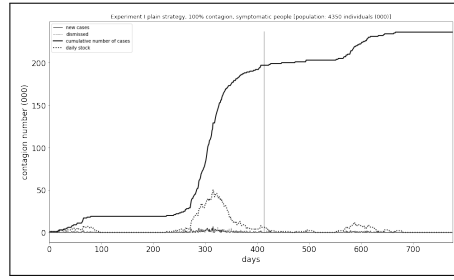
(a) “Plain” vacc. sequence; on the y axis the number of vaccinated subjects of each group (if vaccination is complete, the line is horizontal)



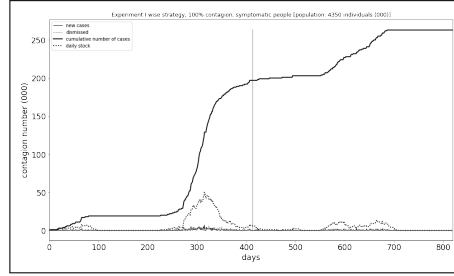
(c) “Wise” vacc. sequence; on the y axis the number of vaccinated subjects of each group (if vaccination is complete, the line is horizontal)



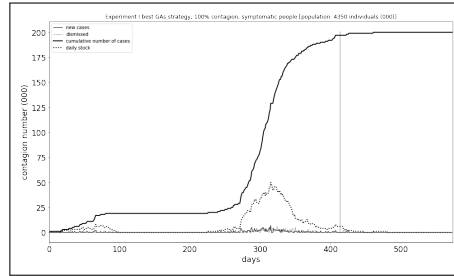
(e) GA vacc. sequence; on the y axis the number of vaccinated subjects of each group (if vaccination is complete, the line is horizontal)



(b) “Plain” vaccination symptomatic series; the vertical line is at day 413, when the effectiveness of first vaccination starts



(d) “Wise” vaccination symptomatic series; the vertical line is at day 413, when the effectiveness of first vaccination starts



(f) GAs vaccination symptomatic series; the vertical line is at day 413, when the effectiveness of first vaccination starts

Fig. 11: Vaccination sequences and time series

7.7.2 Time dynamics with *plain* vac. strategy, vac. people still spreading the infection

In Figs. 11a and 11b ...

7.7.3 Time dynamics with *wise* vac. strategy, vac. people still spreading the infection

In Figs. 11c and 11d . . .

7.7.4 Time dynamics with best GAs strategy, vac, people still spreading the infection

In Figs. 11e and 11f . . .

8 A new model and future developments

9 Appendix 1—Parameter values

We report here the values of parameters of Fig. 2, also with the short names used in program scripts, in round brackets. Look at Section 2.2 for the definition. Day numbering is related to actual dates via the Table 14. Day 1 is fixed at Feb 4th, 2020.

The values adopted in the experiments reported in this work are the following.

- i The values of *probabilityOfGettingInfection* (prob) are: 0.05 (starting phase); 0.02 at day 49 (adoption of non-pharmaceutical measures); 0.035 at day 149 (some relaxation in compliance); 0.02 at day 266 (again, compliance to rules).
- ii *intrinsicSusceptibility* is set discussing Eq. 1 in (2.2, ii).
- iii The values of *%PeopleAnyTypeNotSymptomaticLeavingHome* (%PeopleAny) are: at (day) 20, 90; at 28, 80; at 31, 0; at 106, 80; at 110, 95; at 112, 85; at 117, 95; at 121, 90; at 259, 90; at 266, 80; at 277, 50; at 302, 70; at 320, 90; at 325, 50; at 329, 80; at 332, 50; at 336, 80; at 337, 50; at 339, 80;
- iv The values of *%PeopleNotFragileNotSymptomaticLeavingHome* (%PeopleNot) are: at (day) 31, 80; at 35, 70; at 36, 65; at 38, 15; at 42, 25; at 84, 30; at 106, 0; at 302, 90; at 325, 50; at 332, 50; at 337, 50; at 339, 100; at 349, 90;
- v The values of *%openFactoriesWhenLimitationsOn* (%Fac) are: at (day) 38, value4 0; at 49, 20; at 84, 70; at 106, 100; at 266, 90; at 277, 70; at 302, 80; at 320, 90; at 325, 30; at 329, 90; at 332, 30; 336, 90; at 337, 30; at 339, 100;
- vi *stopFragileWorkers* (sFW): by default, 0; in one of the experiments we used sFW with set to 1 (on) at day 245 and to 0 (off) at day 275;
- vii The values of *activateSchools* (aSch) are: at (day) 1, on; at 17, off; at 225, on; at 325, off; at 339, on;
the values of *%Students* (%St) are: at (day) 0, 100; at 277, 50; at 339, 50; at 350, 50 (repeated values are not relevant for the model, but for the use of the programmer-author);
- viii The value of *radiusOfInfection* (radius) is 0.2; in the model, space is missing of a scale, but forcing the area to be in the scale of a region as Piedmont, 0.2 is

Day	Date	Day	Date	Day	Date	Day	Date
25	28- 2-2020	200	21- 8-2020	375	12- 2-2021	550	6- 8-2021
50	24- 3-2020	225	15- 9-2020	400	9- 3-2021	575	31- 8-2021
75	18- 4-2020	250	10-10-2020	425	3- 4-2021	600	25- 9-2021
100	13- 5-2020	275	4-11-2020	450	28- 4-2021	625	20-10-2021
125	7- 6-2020	300	29-11-2020	475	23- 5-2021	650	14-11-2021
150	2- 7-2020	325	24-12-2020	500	17- 6-2021	675	9-12-2021
175	27- 7-2020	350	18- 1-2021	525	12- 7-2021	700	3- 1-2022

Table 14: The days of the simulation and their equivalent dates in the calendar

equivalent to 20 meters; we have to better calibrate this measure with movements and probabilities; this is a critical step in future developments of the model.

10 Appendix 2—A gallery of contagion sequences

The gallery of contagion sequences, reported in Table 15, shows the vast variety of situations generated by the agent-based simulations, producing actions and interactions of the agents.

- 1a An outbreak without containment measures, with a unique wave, but very heavy: contagions are in nursing homes (orange), workplaces (brown), homes (cyan), hospitals (pink).
- 1b This is the previous epidemic without containment measures, considering the first 200 infections, with the main contribution of nursing homes (orange) and workplaces (brown).
- 1c Another outbreak, always without containment measures: nursing homes (orange) as a starter.
- 2a The [1c] epidemic, without containment measures, first 200 infections: nursing homes (orange) as a starter; around day 70, a unique contagion at home makes the epidemic continue.
- 2b Another case without containment measures showing the initial action of contagions in workplaces (brown) and homes (cyan).
- 2c Here we see the first 200 infections showing that the initial profound effects of contagions in workplaces (brown) and homes are due, in the beginning, to the fragile persons, also asymptomatic,
- 3a An outbreak with containment measures, where we see another influential contribution of workplaces (brown) and homes (cyan) to the epidemic diffusion.
- 3b Here the first 200 infections: after day 100, we observe many significant cases of fragile workers diffusing the infection.

- 3c In this outbreak, with containment measures, the infections arise from workplaces (brown), nursing homes (orange), and homes (cyan), but also hospitals (pink).
- 4a Here we explore the first 200 infections of [3c]: in the beginning, workplaces (brown), hospitals (pink), nursing homes (orange), and homes (cyan) are interweaving.
- 4b An outbreak with containment measures where the effect of the contagions in workplaces (brown), nursing homes (orange), and homes (cyan) is evident.
- 4c In the first 200 infections of [4b], workplaces (brown) and nursing homes (orange) are strictly interweaving.
- 5a An outbreak with containment measures where the effect of nursing homes (orange) is prevalent.
- 5b An outbreak with containment measures with a highly significant effect from workplaces (brown).
- 5c Stopping fragile workers at day 20 in the previous case, we obtain a beneficial effect, but home contagions (cyan) keep alive the pandemic, which explodes again in workplaces (brown).
- 6a Exploring the first 200 infections of the case [5c], we have evidence of the event around day 110 with the new phase due to a unique asymptomatic worker.
- 6b Finally, the same epidemic stopping fragile workers and any fragility at day 15 case and isolating nursing homes.
- 6c An outbreak with containment measures spontaneously stopping in a short period.

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References

1. A. Scala, A. Flori, A. Spelta, E. Brugnoli, M. Cinelli, W. Quattrociocchi, F. Pammolli, *Scientific Reports* **10**(1), 13764 (2020). DOI 10.1038/s41598-020-70631-9. URL <https://doi.org/10.1038/s41598-020-70631-9>
2. N. Bellomo, R. Bingham, M.A.J. Chaplain, G. Dosi, G. Forni, D.A. Knopoff, J. Lowengrub, R. Twarock, M.E. Virgillito, *Mathematical Models and Methods in Applied Sciences* **30**(08), 1591 (2020). URL <https://doi.org/10.1142/S0218202520500323>
3. H. Rahmandad, J. Sterman, *Management Science* **54**(5), 998 (2008)

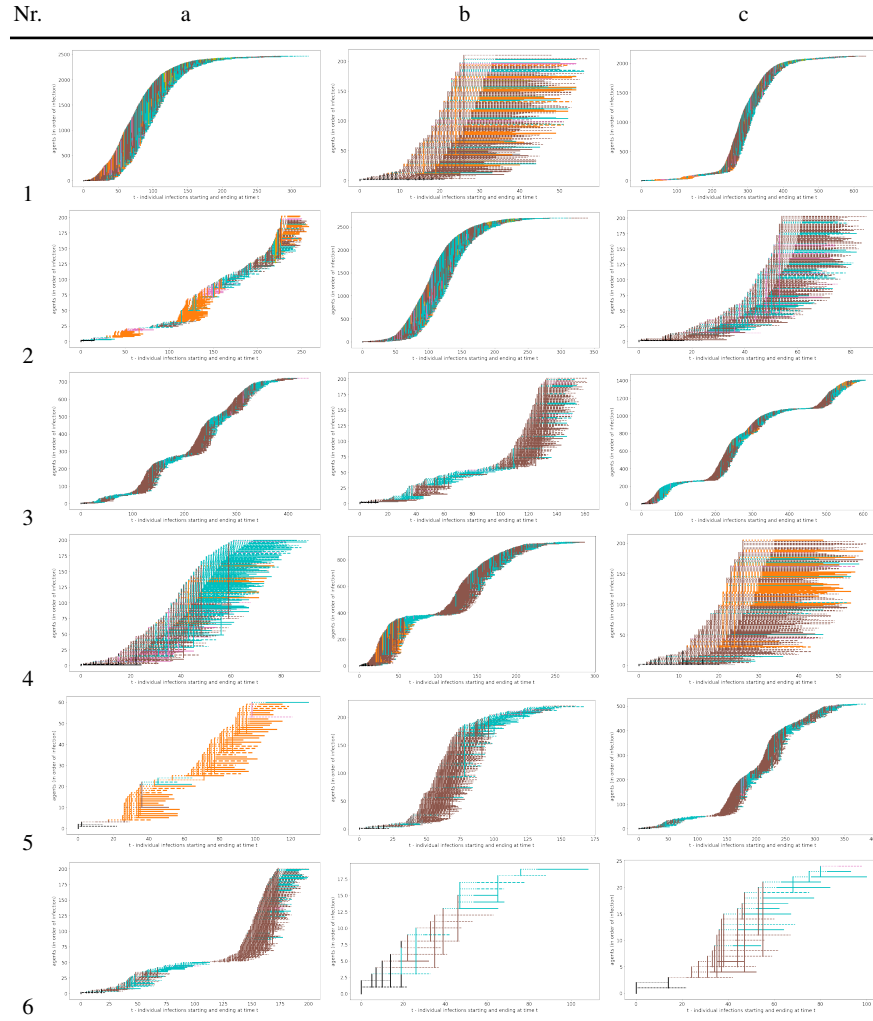


Table 15: Gallery of sequences, symptomatic and asymptomatic agents

4. F. Squazzoni, J.G. Polhill, B. Edmonds, P. Ahrweiler, P. Antosz, G. Scholz, E. Chappin, M. Borit, H. Verhagen, F. Giardini, N. Gilbert, *Journal of Artificial Societies and Social Simulation* **23**(2), 10 (2020). DOI 10.18564/jasss.4298. URL <http://jasss.soc.surrey.ac.uk/23/2/10.html>
5. J.B. Bak-Coleman, M. Alfano, W. Barfuss, C.T. Bergstrom, M.A. Centeno, I.D. Couzin, J.F. Donges, M. Galesic, A.S. Gersick, J. Jacquet, A.B. Kao, R.E. Moran, P. Romanczuk, D.I. Rubenstein, K.J. Tombak, J.J. Van Bavel, E.U. Weber, *Proceedings of the National Academy of Sciences* **118**(27) (2021). DOI 10.1073/pnas.2025764118. URL <https://www.pnas.org/content/118/27/e2025764118>

6. J. Epstein, *Journal of Artificial Societies and Social Simulation* **11**(4), 12 (2008). URL <http://jasss.soc.surrey.ac.uk/11/4/12.html>
7. R. Axtell, *Why agents? On the varied motivations for agent computing in the social sciences*. Tech. rep., Center on Social and Economic Dynamics Brookings Institution (2000)
8. A. Ghorbani, F. Lorig, B. de Bruin, P. Davidsson, F. Dignum, V. Dignum, M. van der Hurk, M. Jensen, C. Kammler, K. Kreulen, L.G. Ludescher, A. Melchior, R. Mellema, C. Păstrăv, L. Vanhée, H. Verhagen, *Review of Artificial Societies and Social Simulation* (2020). URL <https://rofasss.org/2020/04/25/the-assocc-simulation-model/>
9. F. Silvagno, A. Vernone, G.P. Pescarmona, *Antioxidants* **9**(7), 624 (2020). DOI 10.3390/antiox9070624. URL <http://dx.doi.org/10.3390/antiox9070624>
10. R. Horton, *Lancet* (London, England) **396**(10255), 874 (2020). URL <https://www.thelancet.com/action/showPdf?pii=S0140-6736%2820%2932000-6>
11. P. Terna, G. Pescarmona, A. Acquadro, P. Pescarmona, G. Russo, S. Terna. *SIsaR, An Agent-Based Model of the Diffusion of Covid-19 Using NetLogo, with Susceptible, Infected, symptomatic, asymptomatic, and Recovered People* (2020). URL <https://terna.to.it/simul/SIsaR.html>
12. U. Wilensky. *Netlogo* (1999). URL <http://ccl.northwestern.edu/netlogo/>
13. T.M. Vu, C. Probst, J.M. Epstein, A. Brennan, M. Strong, R.C. Purshouse, *Genetic and Evolutionary Computation Conference : [proceedings]*. *Genetic and Evolutionary Computation Conference* **2019**, 1356 (2019). DOI 10.1145/3321707.3321840. URL <https://pubmed.ncbi.nlm.nih.gov/33083795>
14. P. Steinmann, J.R. Wang, G.A. van Voorn, J.H. Kwakkel, *Review of Artificial Societies and Social Simulation* **17** (2020). URL <https://rofasss.org/2020/04/17/deep-uncertainty/>
15. N. Phillips, *Nature* **590**(7846), 382 (2021). URL <https://www.nature.com/articles/d41586-021-00396-2>
16. D. Miles, M. Stedman, A.H. Heald, *International Journal of Clinical Practice* **n/a**(n/a), e13674 (2020). DOI 10.1111/ijcp.13674. URL <https://onlinelibrary.wiley.com/doi/abs/10.1111/ijcp.13674>