Time, Space and Demography: Key Factors in the Exit Mechanisms for the Covid-19 Epidemics

Antonio Scala^{1,2}, Andrea Flori³, Alessandro Spelta^{4,5}, Emanuele Brugnoli¹, Matteo Cinelli¹, Walter Quattrociocchi^{6,1}, and Fabio Pammolli^{3,5}

¹Applico Lab, CNR-ISC

²Big Data in Health Society

³Impact, Department of Management, Economics and Industrial Engineering, Politecnico di Milano

⁴Univ. di Pavia

⁵Center for Analysis Decisions and Society, Human Technopole and Politecnico di Milano

⁶Univ. di Venezia 'Ca Foscari

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Abstract

We develop a minimalist compartmental model to analyze the impact of mobility restrictions in Italy during the Covid-19 outbreak. Our findings show that early lockdowns shifts the epidemic in time and that beyond a critical value of the lockdown strength, the epidemic restarts after lifting the restrictions. We investigate the effects of different lockdown scenarios and exit mechanisms by accounting for two fundamental sources of heterogeneity within the model: geography and demography. First, we consider Italian Regions as separate administrative entities, in which social interactions between age cohorts occur. Due to the sparsity of the mobility matrix, epidemics tend to develop independently in different regions. Second, we show how disregarding the specific structure of social contacts between age classes could lead to severe underestimation of a second wave of infection at the end of the lockdown period, while measures which take into account the age of the population can sustain the mitigation of rebound effects. Our model is general, and it highlights the effects of key parameters on non-pharmaceutical mitigation mechanisms for epidemics.

1 Introduction

Different epidemic models and approaches contribute to identify specific mechanisms relevant for policy design [1]. At present, although the World Health Organization (WHO) organizes regular calls for Covid-19 modelers to compare strategies and outcomes, policymakers barely handle the discrepancies between the proposed models¹.

To contain the Covid-19 epidemic, governments worldwide have adopted severe social distancing policies, ranging from partial to total population lockdown [2]. Restrictions have led to a sudden stop of economic activities in many sectors, while the majority of Covid-19 infections affect active population (i.e. the age cohort between 15–64 years) [3]. Overall, the impact of contagion and lockdown measures on health and on economic activities is substantial and pervasive.

Against this background, we introduce a model-based scenario analysis for Covid-19, highlighting the role of non-pharmaceutical variables in the epidemic spreading and how those dimensions are likely to impact on lockdown policies and exit strategies [4].

The general behavior of our model holds for the vast class of epidemic models where transmission rate is proportional to the number of susceptible people times the density of infected. We focus on the determinants of short-term interventions in response to an emerging epidemic when geographic and demographic compartiments are included in the model. Our goal is general in nature, since we focus on two relevant decomposability conditions, under which partial dynamics influence the overall configuration of the system (see, e.g., [5, 6, 7, 8]). We study how mobility restriction measures and the timing of the lockdown lift affect the total fraction of infected, the peak prevalence, and, possibly, the delay of the epidemic. Our analysis identifies two fundamental sources of heterogeneity in the diffusion process: regional boundaries and age cohorts [4]. We show how such dimensions can shape policy interventions aiming at containing the epidemic, irrespective of any detailed quantitative predictions. Then, we point out how disregarding such dimensions would affect exit scenarios.

With this paper, we contribute to the extant literature on trade-offs between mitigation, aimed to slow down the epidemic contagion, and suppression, consisting of temporarily lowering the risk of contagion [4, 9, 10]. Notwithstanding detailed quantitative information and relevant medical issues are not addressed here, our work still reveals how a simple and parsimonious compartimental model based on geographical and age classes uncovers relevant aspects which may help guiding decision makers in balancing the restrictions of the lockdown phase and the timing of its release. First, we show that early lockdowns shift the epidemic in time and that the delay is proportional to the anticipation time with an intensity which grows with the strength of the lockdown. Beyond a critical threshold, the epidemic would tend to fully recover its strength as soon as the lockdown is lifted. Second, we show how the sparsity of the matrix representing mobility flows across administrative regions accounts for the delays of

 $^{^1 \}rm See,~e.g.:~https://www.sciencemag.org/news/2020/03/mathematics-life-and-death-how-disease-models-shape-national-shutdowns-and-other$

the contagion and implies that once the epidemic has started, it then develops independently; the same consideration may apply to explain the independent dynamic of the epidemic spreading in different countries, as what observed in mainland China [11]. Finally, we consider social contact heterogeneity between age cohorts and we find that the structure of social contacts is of primarily importance to estimate post-lockdown effects so that age based strategies are a key ingredient to mitigate rebound effects.

2 Model

To analyze mobility-restriction policies, we introduce a minimalist compartmental model [10, 12]. Although many models, both mechanistic, statistic and stochastic [13], have been proposed for the Covid-19 infection, data collected from the national healthcare systems suffer from the lack of homogeneous procedures in medical testing, sampling and data collection [14]. Not to mention the difficulties in assessing the impact of variability in social habits during the epidemics [10, 15]. Moreover, especially in the early phases of the epidemic – i.e. the ones characterised by an exponential growth – different models sharing a given reproduction number R_0 can fit the data with equivalent accuracy (see SI discussion about fitting initial parameters). For these reasons, our aim is to focus on some fundamental qualitative scenarios and not on detailed predictions. We adapt the SIR model, the most basic epidemic model for flu-like epidemics, to the observed data available in the Italian case.

The model relies on four compartments, namely: S, I, O, R. Hence, S (usceptible) individuals can become I (nfective) when meeting another infective individual, I (nfectives) either become O (bserved) – i.e. present symptoms acute enough to be detected from the national health-care system – or are R (emoved) from the infection cycle by having recovered; also O (bserved) individuals are eventually R (emoved) from the infection cycle² (see SIs for a visual representation of the model workflow). Notice that, it is not still clear if there is an asymptomatic phase [16, 17]; we are implicitly assuming that asymptomatics are infective and their removal time is the same of the I class. The model is described by the following differential equations:

$$\partial_t S = -\beta S \frac{I}{N}$$

$$\partial_t I = \beta S \frac{I}{N} - \gamma I$$

$$\partial_t O = \rho \gamma I - hO$$

$$\partial_t R = (1 - \rho) \gamma I + hO$$
(1)

N = S + I + O + R is the total number of individuals in a population, the

 $^{^2}$ Notice that we are implicitly absorbing the number of deaths in the R(emoved) compartment of the model, that therefore comprises both the Recovered people (who hopefully have developed antibodies and are not anymore susceptible) and the small fraction of people who are not infective anymore since they did not overcome the epidemics.

transmission coefficient β is the rate at which a susceptible becomes infected upon meeting an infected individual, γ is the rate at which an infected either becomes observable or is removed from the infection cycle. Like the SIR model, the basic reproduction number is $R_0 = \beta/\gamma$; the extra parameters of the SIOR model are ρ , the fraction of infected that become observed from the national health-care system, and h, the rate at which observed individuals are removed from the infection cycle. Notice that we consider that O(bserved) individuals not infecting others, being in a strict quarantine.

3 The Italian Lockdown

The Italian lockdown measures of the 8th and 9th of March [18, 19] were intended to change mobility patterns and to reduce de-visu social contacts, through quarantine measures and to an increased awareness of the importance of social distancing. We analyze an extensive data set on Facebook mobility data³ [20]; our analysis confirms that the lockdown has reduced both the travelled distance and the flow of travelling people.

We consider the effects of such lockdown measures on the parameters of our model. Lockdowns are non-pharmaceutical measures; hence the rate γ is the most unaffected, since it is related to the "medical" evolution of the disease. Analogous arguments apply to the rate h of exiting a condition serious enough to be observed and to the probability ρ of being observed by the national healthcare system (although ρ could be influenced variations in testing schemes and alert thresholds). On the other hand, the transmission coefficient β can be thought as the product $C\lambda$ of a contact rate C times a disease-dependent transmission probability λ . Hence, if we assume that the speed of Covid-19 mutation is irrelevant on our timescales, lockdown strategies mostly influence β by reducing the contact rate C among individuals.

To adapt the SIOR's parameters to the Italian data [21], we compare the reported cumulative number of Covid-19 cases Y^{Obs} with the analogous quantity $Y^{\mathrm{model}} = \int \rho \gamma I dt$ in our model. We want to stress that our model fitting is not aimed to produce an accurate model for detailed predictions, but to work in a realistic region of the parameter space.

We first estimate model's parameters by least square fitting on the prelockdown period. Since in such range the data Y^{Obs} show an exponential growth trend, we are possibly observing a very early phase of the epidemic, where $\beta - \gamma$ equals the growth rate of Y^{Obs} (see SI for observations on the choices of initial parameters). For fixed $\beta - \gamma$, the time of the epidemic start (that we conventionally assume as the time t_0 where the number of infected is 1) and the fraction ρ of serious cases observed by the national healthcare service, allows to vary the values of β and γ as long as their difference is fixed. Hence, estimating

 $^{^3{\}rm Those}$ data are part of the Facebook project "Data for Good", and illustrate mobility patterns of fb users, who allowed the social network to track their location. See https://dataforgood.fb.com/docs/Covid-19/

$\beta = 0.35 day^{-1}$	$\gamma = 10^{-1} day^{-1}$	$h = 1/9 day^{-1}$
$t_0 = -30 days$	$\rho = 40\%$	$\alpha = 0.49$

Tab. 1. Standard parameters used for the SIOR model in the paper.

medical parameters as the rate γ of escaping the infected state is paramount for calibrating mathematical models.

In response to the outbreak of Covid-19, several estimates of model parameters have been proposed in the literature, revealing a certain amount of uncertainty about some fundamental variables of the epidemic contagion. The European Centre for Disease controls reports an infection time duration τ_I between 5 and 14 days [22]; in our model, we will use $\tau_I = 10$ (i.e. $\gamma = \tau_I^{-1} = 1/10 \, days^{-1}$). According to a report of ISS, the Italian National Health Institute, the time from the start of serious symptoms (i.e. when one gets "observed" from ISS) to the resolution of the symptoms can be estimated as $\tau_H \sim 9 \, days$ [23], corresponding in our model to a value $h = 1/9 \, days^{-1}$. Notice the analysis of 12 different models [13] reports varying estimates for the basic reproduction number R_0 , ranging from 1.5 to 6.47, with mean 3.28 and a median of 2.79.

From fitting the 15 days of Y^{obs} (pre-lockdown phase) and by performing a bootstrap sensitivity analysis of the parameters, we obtain $\beta - \gamma \sim 0.25 \pm 0.01$ and $t_0 = -30 \pm 5 \, days$ by assuming that $\rho = 40\%$. Varying ρ in [10%, 100%] varies $\beta - \gamma$ in [0.22, 0.27]. On the other hand, for fixed $\beta - \gamma$, R_0 would vary linearly with τ_I ; as an example, R_0 varies in [2.5, 4.5] for the literature parameters $\tau_I \in [5, 14]$; accordingly, to adjust the difference in growth rate, t_0 varies in [26, 32]. However, despite the variability of the parameter range, the qualitative behavior of the model – and hence our analysis of the key factors of the epidemic evolution – is unchanged.

We then assume that, after the lockdown day $t_{\rm Lock}=15$ (corresponding to the 9^{th} of march), contact rate drops down by a factor α and hence $\beta \to \alpha\beta$. By fitting the observed data Y^{obs} for a symmetric period of 15 days after $t_{\rm Lock}$, and by performing a bootstrap sensitivity analysis, we find $\alpha=0.49\pm0.01$, i.e. a $\sim 50\%$ reduction in infectivity and hence in R_0 . Our figure is in line with the observed reduction in R_0 in response to the combined non-pharmaceutical interventions, that across several countries has and average reduction of 64% compared to the pre-intervention values [24]. Notice that Facebook mobility data show a post-lockdown reduction in mobility of 15% at regional level and of 73% at inter-regional level; however, as we will point out later, mobility is much more related to the beginning of epidemics in new regions/countries than to the evolution of the epidemics in a region/country.

In the following of the paper, we will use the exemplified parameters of Tab. 1 corresponding to a basic reproduction number $R_0 = 3.5$. Moreover, since patients in intensive care represent the highest burden for the national health systems, in the graphs of the paper we will indicate the number of patients in intensive care, estimated as 3.5% of the total patients by using the figures reported by ISS [21].

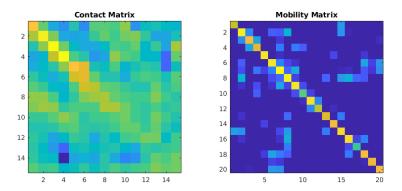


Fig. 1. Left Panel: social contact matrix, from [25]. Right panel: inter-regional mobility matrix, from the Facebook project "Data for Good". The intensity of a color maps the strength of a matrix element (light colors: high values; dark colors: low values). The inter-age social mixing matrix is dense; hence age classes dynamics are strongly coupled. The inter-regional mobility flows is very sparse (i.e. off diagonal elements are order of magnitudes lower than diagonal elements): this mean that most of the people travel within the same region of origin; hence, the regional dynamics can be considered "almost" decoupled.

4 National scenarios and exit mechanisms

Since we are interested on the factors driving the exit dynamics from lockdown, and not on the detailed analysis or production of realistic scenarios, we will consider several lockdown scenarios where the lockdown is abruptly lifted and the system let return to the pre-lockdown parameters. Such an approach amplifies the possible perils and represent a worst-case estimate of the exit strategies. Hence, we consider several simplified scenarios where we use the SIOR model of Eq. 1 with the parameters of Tab. 1: First, in the simple case of a SIOR model fitted on Italian data, we analyze how the post-lockdown dynamics changes according to different starting dates of the epidemics and to different levels of the restrictions implemented by the national authorities. Then, we study the effect of explicitly considering Italy as a collection of separate administrative entities (Regions) independently evolving; finally, we consider the effects of social interactions across age cohorts.

Interestingly, mobility flows [20] and inter-age social mixing [25] lie at the two opposite range of modelling. In fact, the regional social contact matrix is dense (Fig. 1, left panel), indicating that age classes dynamics are strongly coupled. On the other hand, the inter-regional mobility matrix is very sparse (Fig. 1, right panel), indicating that regions have their own independent dynamics.

We first consider a simple exit strategy consisting in lifting the lockdown at a time $t_{\rm Unlock}$ after the peak of O has occurred. For instance, we hypothesize that infection proceeds uncontrolled up to time $t_{\rm Lock}$; in the following lockdown period $[t_{\rm Lock}, t_{\rm Unlock}]$, the transmission coefficient β is reduced by a factor α ;

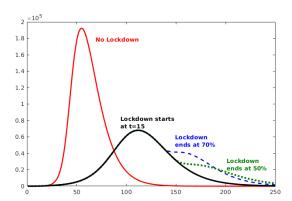


Fig. 2. Comparison of the scenarios where the lockdown is relaxed after the percentage of people with visible symptoms (O) is reached the 70% and the 50% of the reported cases peak. Lifting the lockdown earlier has the epidemics disappear faster, but has higher impact on the number of hospitalized and intensive care patients; moreover, lifting the lockdown too early can resoult in a rebound of the number of cases.

finally, β returns to its initial value and herd immunity is responsible for the dampening of the epidemics.

Our results show that the lockdown lowers the peak of O - i.e. the individuals with noticeable symptoms - to $\sim 70\%$ of the free epidemic one, but also doubles the time of its occurrence from ~ 1.9 months to ~ 3.8 months: an extremely obnoxious effect for the sustainability conditions of the economy of a country. However, since the number of hospitalized patients and - most importantly - the number of patients in intensive care is only a fraction of O, lowering the peak puts less stress on the healthcare system. The ideal situation would be to have accurate data, an accurate model and accurate estimates of the parameters; as an example, in our model lifting the lockdown when the number of infected people per unit time $\beta S(t)I(t)/N$ is lower that the average number of recovering people $\gamma I(t)$ would ensure that the number of infections would continue to decrease. In real life, situations are more fuzzy: having not enough information, we could decide to resort on some heuristics, like lifting the lockdown after the observed people O have dropped to a suitable percentage of the maximum peak. As an example, after ~ 4.7 months the peak has reduced to 70% of its initial value, while after ~ 5.2 months to 50%, i.e. ~ 0.5 months later. Notice that, the earlier the lockdown is lifted, the faster O decays to zero even if it starts from higher figures and could even experience a rebound. All such effects are shown in Fig. 2.

Our framework sustains the identification of several mechanisms. The first is related to the timeliness of the lockdown, i.e. to the choice of anticipating t_{Lock} . As expected, early lockdown (i.e. well before the "free" infection peak) reduces

the height of the peak without much moving it forward in time. Conversely, lifting the lockdown too soon can make epidemic start again and reach values higher than the ones before the release. A peculiar and counter-intuitive effect can be generated if the lockdown is anticipated: in fact, a too early lockdown delays the start of the epidemic without attenuating its severity (see SI for the description of the effects of varying lockdown time). In other words, an early lockdown "buys" time, but it postpones the problem without mitigating its severity.

Another effect is the impact of extreme quarantine measures on the post-lockdown dynamics. Increasing the strength α of the lockdown (i.e., reducing the social contacts), not only delays the time at which the lockdown can be lifted, but it also induces a stronger re-start of the epidemic in the post-lockdown (see SI for the description of the effects of varying lockdown strength), a restart that would trigger the necessity of a new lock-down and so forth. Such scenario could result to be not sustainable for the economy.

An additional counter-intuitive mechanism must be considered. Since to an attenuation α corresponds an effective reproduction number $R_0^{\rm eff} = \alpha R_0$, at the critical value $\alpha_{\rm crit} = 1/R_0$ the epidemic neither grows nor decreases⁴. Thus, after $t_{\rm Lock}$ the system stays stationary until the quarantine is released at $t_{\rm Unlock}$; at this point the epidemic starts growing again as it was before the lockdown. In general, if $\alpha < \alpha_{\rm crit}$, the system looks to ameliorate (infected, hospitalized, all the infective compartments go down) but as soon as the lockdown is lifted, the epidemic starts again to reach its full extent (see SI). Nevertheless, our estimate $\alpha \sim 0.5 > \alpha_{\rm crit} \sim 0.3$ for the Italian lockdown gives us hope that, perhaps, it will not be necessary to follow a repeated seek-and-release strategy in the post-lockdown phase. On the other hand, if it can be attained a lockdown strength $\alpha \sim \alpha_{\rm crit}$ without disrupting the economy, the epidemic could be contained until the creation, production and distribution of a vaccine.

5 Regional Scenarios

Starting with the first confirmed cases in Lombardy on 21 February, by the beginning of March the Covid-19 outbreak had spread to all regions of Italy. While the delay in the beginning of the infection is due to the different interaction among regions, once an epidemic has started it grows exponentially and the intake of external infected people becomes quickly irrelevant (see SI for the description of a metaregional model and its behavior). As a consequence, the growth curves of the epidemic variables should tend to the same shape (see SI about using normalised data). Coherently, looking at the regional info graphics released by the Italian National Healthcare Institute (ISS) [21], one may notice that they have a similar shape but different starting date (see Fig. 3). Such observation can be justified as follows: Italian regions are independent administrative units, where most of the population tend to work inside the resident

⁴To be precise, the decrease becomes sub-exponential, thus taking a practically infinite time in populations as big as a country or even a megacity

Tab. 2. Regional delays (in days)

Lombardia	0.0	Molise	10.6
Emilia Romagna	3.1	Umbria	11.8
Marche	4.3	Abruzzo	13.1
Veneto	5.7	Lazio	14.5
Valle d'Aosta	6.4	Campania	15.0
P.A. Trento	6.6	Puglia	15.7
P.A. Bolzano	8.0	Sardegna	16.2
Liguria	8.1	Sicilia	16.6
Friuli Venezia Giulia	8.9	Calabria	17.2
Piemonte	9.0	Basilicata	19.2
Toscana	10.4		

region [26]. Hence, epidemics propagate from region to region via the fewer inter-regional exchanges (notice that Lombardy is among the Italian regions most involved in international trade connections [27], therefore it appears as one of the most probable candidate for the outbreak of the Italian epidemic). More practically, we estimate these delays by minimizing the distance among the observed curves (see SI for the details of the algorithm); results are reported in Tab. 2. Notice that, assuming Lombardy has been the first region (i.e. delay=0), the resulting regional delays are mostly correlated to geographical distances.

We assume that the Covid-19 outbreak spreads independently in each region of Italy; as argued before, such an approximation is reasonable after the epidemic has started and is even more accurate under lockdown conditions. Hence, we apply the parameters for the whole Italy to regional cases⁵, where now the maximum number of individuals N_i is the population of the i^{th} region⁶. Then, by summing up all the S_i, \ldots, R_i , respectively, we obtain a proxy for the global evolution of Covid-19 epidemic throughout Italy. To evaluate the effect of heterogeneity in time delays, we compare the number of daily cases $O^{\text{Delay}} = \sum O_i^{\text{Delay}}$ (obtained by taking into account the regional delays t_i as reported in Tab. 2) with the number of daily cases $O^0 = \sum O_i^0$ we would observe by considering the epidemics starts at the same time t_0 in all regions. As expected, heterogeneity flattens the curve and shifts its maximum later in time. This is a first source of errors when fitting an heterogeneous dynamics with a global model.

Assuming that the right approach is the one with regional delays, we consider two possible exit strategies: in the first, that we call the Asynchronous scenario, each region i lifts the lockdown at the time $t_i^{\rm Unlock}$ when the peak of $O_i^{\rm Delay}$ decreases by 30%; in the second, that we call the Synchronous scenario, each

⁵Again, we are exploring qualitative scenarios and not trying to predict the real evolution of the epidemics: in fact, Italian regions are different for social contact habits, mobility, healthcare quality and even for factors that could possibly affect the medical parameters like comorbidities, climate or pollution levels.

⁶https://www.istat.it/it/popolazione-e-famiglie?dati

region i lifts the lockdown at the same time t^{Unlock} , i.e., when the global peak of O^{Delay} decreases by 30%. The choice of 30% is arbitrary, similar results would hold for choices of values near the peak; it tries to be a sketch of a situation where, due to economic pressure, lockdown is lifted as soon as possible.

Notice that, once an outbreak begins, the epidemic dynamics in a region i is essentially uncorrelated with the epidemic spreading of any other region $j \neq i$. Therefore, it could be safe and appropriate to decide the lockdown lifting time on a regional basis, instead of lifting restrictions throughout Italy at the same time. Indeed, it could be unreasonabile to keep locked the regions where the epidemic started earlier; on the contrary, regions where the epidemic began with some delay could experience a strong rebound when subjected to a premature lockdown lifting. In Fig. 3 we show the effects of lifting the lockdown at both regional (Async) and national (Sync) level in Lazio and Lombardy. Since not only epidemics, but also the ruin of an economy is a non-linear process, the Sync scenario can turn out to be even more disruptive than the epidemic itself (see also Fig. 2). Notice that analogous arguments hold - mutatis mutandis also for the world/countries scenario.

6 The role of Age

As we have already observed in the previous Section, heterogeneity strongly impacts on the results of a model [28]. Since the transmission coefficient is proportional to the contact rate of individuals, the rates of social mixing among different age classes could represent another important source of heterogeneity. This information has been estimated either through large-scales surveys [25] or through virtual populations modeling [29]. While the POLYMOD [25] matrices have been extensively used to estimate the cost-effectiveness of vaccination for different age-classes during the 2009 H1N1 pandemic [30, 31], here they are used to support the design of lockdown measures and exit strategies. Hence, to account for age classes, we extend our model by rewriting the transmission coefficient as βC (see SI for a full description of the extended model), where β is the transmission probability of the infection, and C is the sociological matrix describing the contact patterns typical of a given country. For lack of further information, we assume β constant among age classes and C as in [25]. To simplify the analysis, we gather POLYMOD age groups into three classes: Young (00-19), Middle (20-69) and Elderly (70+) (see Tab 3). Such aggregation puts together the most "contactful" classes (00-19), the classes with the highest mortality risk (70+) [21], and a god approximation of the active population (20-69).

Fig. 4 shows how the percentage of people with visible symptoms (O) varies once the age class heterogeneity has been added into the model. Notice that, at difference of fig. 2, fully lifting the lockdown results in a conspicuous rebound of the epidemics, that reaches values even more severe than the pre-exit peak. Thus, models not considering heterogeneities could severely mis-forecast the post-exit dynamics. on the other hand, introducing the age structure in the

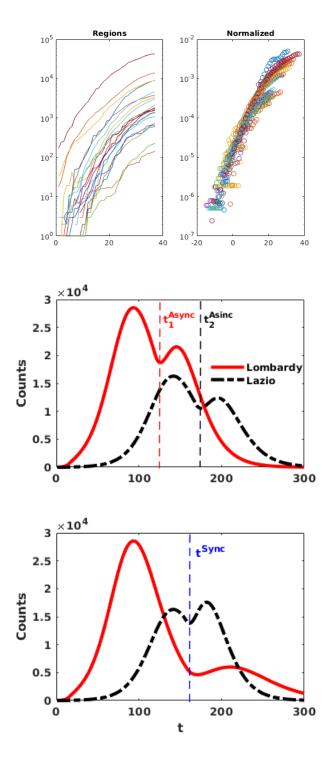


Fig. 3. Upper panels: analysis of time delays among the start of epidemics in different regions (see Tab. 2). Lower panels: sketch of an Async(hronous) exit strategy (i.e. each region lifts the lockdown following its own policy) respect to a Sync(hronous) exit strategy (i.e. the lockdown lift follows the same policy, but applied to a nation wide scale). In particular, t^{Sync} corresponds to lifting the lockdown in all the region after the peak has fallen by 30%, while t_i^{Async} corresponds to lifting the lockdown in the i^{th} region after the peak of such region has fallen by 30%.

	Y	M	Е
Y	2.35	0.44	0.67
M	0.47	0.59	0.50
E	0.50	0.55	0.80

Tab. 3. POLYMOD matrix aggregated for three age classes: Young (00-19), Middle (20-69) and Elderly (70+).

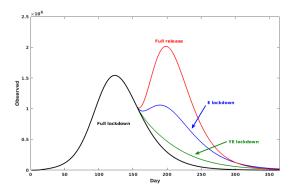


Fig. 4. Comparison of the scenarios where the lockdown is relaxed only for particular age class with respect to a full release policy. Strategies: YE = quarantine young and elderly, E = quarantine elderly. Notice that we have purposefully left the M class fully unrestrained, in order to show how maintaining a partial, age-based lockdown could deeply change the effectiveness of the exit strategy.

model allows to set up different exit strategies based on age-targeted policies for dampening a possible upturn of contagion. Specifically, quarantine measures applied to only elderly people may limit the impact of a renewed upward phase, while maintaining lockdown restrictions to all classes except the middle age class (20-69) could be enough to smooth and lessen the propagation of contagion in the post-lockdown phase. Remember that we are using mock-up exit strategies that lead to worst-case scenarios: in real life, community measures and physical distancing, infection prevention and control in the community, personal hygiene habits and face mask usage could strongly contribute to the dampening of the epidemics [22].

7 Conclusions

In this work we propose and test a general framework to study an epidemic contagion through a compartmental model based on geographical groups and age classes. We reveal how the promptness of lockdown measures has a main effect on the timing of the contagion. Strict social distancing policies reduce the extent of contagion during the lockdown period, but full recover of the contagion occurs once such measures are relaxed. Moreover, we show how local dynamics at regional level can be inappropriately masked when observing the aggregate national system. Regional heterogeneity lowers and widens the curve of the contagion thus determining a shift forward in time for its peak at the aggregate national level. Moreover, by analyzing mobility data, we find that, due to the sparsity of interconnections across regions, contagion develops independently within each region once the epidemic has started. This, in turn, contributes to account for the delays observed in the alignment of the contagion curves across different geographical areas. The independence of regional dynamics allows both to design regional exit strategies almost independently and to maintain – with the appropriate security measures – the movements of goods among the regions. The same reasoning valid for the Italy/regions system can be applied to the world/country system. Finally, we investigate the structure of social contacts to detect the role of different age classes in the spreading of contagion. Both young people (0-19) and elderly people (70+) are the most interconnected classes, and therefore their behavior significantly affects the post-lockdown phase. We show that while confirming quarantine policies only for elderly people can limit the possibility of a renewed upward phase in the contagion dynamics, relaxing the lockdown measures for the middle age class (20-69) can be enough to smooth and lessen the propagation of contagion in the post-lockdown phase.

Although our study is tuned on the Italian Covid-19 contagion, our modeling approach is general enough to help us understand the role of relevant dimensions, beside medical and pharmaceutical ones, in leveraging strategies to contain the epidemics and mitigate its effects. Our framework sustains an assessment of the trade-off between health outcomes and effects on wider economy. In particular, we show how the timeline of post-lockdown measures can benefit from the inclusion of compartmental aspects, such as geographical and age classes. This

feature is general, and it can sustain simulations on specific restrictions, such as those targeting specific age cohorts, especially the most fragile (70+), enforcing social distancing while containing the overall burden on the economy.

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References

- [1] Matt J Keeling. Models of foot-and-mouth disease. *Proceedings of the Royal Society B: Biological Sciences*, 272(1569):1195–1202, 2005.
- [2] Francesco Di Lauro, István Z Kiss, and Joel Miller. The timing of one-shot interventions for epidemic control. *medRxiv*, 2020.
- [3] Vital Surveillances. The epidemiological characteristics of an outbreak of 2019 novel coronavirus diseases (covid-19)—china, 2020. *China CDC Weekly*, 2(8):113–122, 2020.
- [4] Roy M Anderson, Hans Heesterbeek, Don Klinkenberg, and T Déirdre Hollingsworth. How will country-based mitigation measures influence the course of the covid-19 epidemic? *The Lancet*, 395(10228):931–934, 2020.
- [5] Herbert A Simon and Albert Ando. Aggregation of variables in dynamic systems. *Econometrica: journal of the Econometric Society*, pages 111–138, 1961.
- [6] Albert Ando and Franklin M Fisher. Near-decomposability, partition and aggregation, and the relevance of stability discussions. *International Eco*nomic Review, 4(1):53–67, 1963.
- [7] Herbert A Simon. The architecture of complexity. Cambridge, MA: MIT Press, 1996.
- [8] Pierre Jacques Courtois. Decomposability: queueing and computer system applications. Academic Press, 2014.
- [9] Neil Ferguson, Daniel Laydon, Gemma Nedjati Gilani, Natsuko Imai, Kylie Ainslie, Marc Baguelin, Sangeeta Bhatia, Adhiratha Boonyasiri, ZULMA Cucunuba Perez, Gina Cuomo-Dannenburg, et al. Report 9: Impact of non-pharmaceutical interventions (npis) to reduce covid19 mortality and healthcare demand. MRC Centre for Global Infectious Disease Analysis, 2020.

- [10] Kiesha Prem, Yang Liu, Timothy W Russell, Adam J Kucharski, Rosalind M Eggo, Nicholas Davies, Stefan Flasche, Samuel Clifford, Carl AB Pearson, James D Munday, et al. The effect of control strategies to reduce social mixing on outcomes of the covid-19 epidemic in wuhan, china: a modelling study. The Lancet Public Health, 2020.
- [11] Matteo Chinazzi, Jessica T Davis, Marco Ajelli, Corrado Gioannini, Maria Litvinova, Stefano Merler, Ana Pastore y Piontti, Kunpeng Mu, Luca Rossi, Kaiyuan Sun, et al. The effect of travel restrictions on the spread of the 2019 novel coronavirus (covid-19) outbreak. *Science*, 2020.
- [12] Norman TJ Bailey et al. The mathematical theory of infectious diseases and its applications. Charles Griffin & Company Ltd, 5a Crendon Street, High Wycombe, Bucks HP13 6LE., 1975.
- [13] Ying Liu, Albert A Gayle, Annelies Wilder-Smith, and Joacim Rocklöv. The reproductive number of covid-19 is higher compared to sars coronavirus. *Journal of travel medicine*, 2020.
- [14] Francesco Casella. Can the covid-19 epidemic be managed on the basis of daily data? arXiv preprint arXiv:2003.06967, 2020.
- [15] Sebastian Funk, Marcel Salathé, and Vincent AA Jansen. Modelling the influence of human behaviour on the spread of infectious diseases: a review. Journal of the Royal Society Interface, 7(50):1247–1256, 2010.
- [16] Yan Bai, Lingsheng Yao, Tao Wei, Fei Tian, Dong-Yan Jin, Lijuan Chen, and Meiyun Wang. Presumed asymptomatic carrier transmission of covid-19. *Jama*, 2020.
- [17] Hiroshi Nishiura, Tetsuro Kobayashi, Takeshi Miyama, Ayako Suzuki, Sungmok Jung, Katsuma Hayashi, Ryo Kinoshita, Yichi Yang, Baoyin Yuan, Andrei R Akhmetzhanov, et al. Estimation of the asymptomatic ratio of novel coronavirus infections (covid-19). medRxiv, 2020.
- [18] Decreto del Presidente del Consiglio dei Ministri. Ulteriori disposizioni attuative del decreto-legge 23 febbraio 2020, n. 6, recante misure urgenti in materia di contenimento e gestione dell'emergenza epidemiologica da covid-19. Gazzetta Ufficiale, 59(08-03-2020), 2020.
- [19] Decreto del Presidente del Consiglio dei Ministri. Ulteriori disposizioni attuative del decreto-legge 23 febbraio 2020, n. 6, recante misure urgenti in materia di contenimento e gestione dell'emergenza epidemiologica da covid-19. Gazzetta Ufficiale, 62(09-03-2020), 2020.
- [20] Caroline O Buckee, Satchit Balsari, Jennifer Chan, Mercè Crosas, Francesca Dominici, Urs Gasser, Yonatan H Grad, Bryan Grenfell, M Elizabeth Halloran, Moritz UG Kraemer, et al. Aggregated mobility data could help fight covid-19. Science (New York, NY), 2020.

- [21] Istituto Superiore di Sanità. Iss: Sars-cov-2 dati epidemiologici (https://www.epicentro.iss.it/). Technical report, ISS, 2020.
- [22] European Centre for Disease Prevention and Control (ECDC). Coronavirus disease 2019 (covid-19) pandemic: increased transmission in the eu/eea and the uk eigth update, 8 april 2020. 2020.
- [23] Istituto Superiore di Sanitá (ISS) COVID-19 Surveillance Group. Characteristics of covid-19 patients dying in italy report based on available data on march 30th, 2020. 2020.
- [24] Seth Flaxman, Swapnil Mishra, Axel Gandy, H Unwin, H Coupland, T Mellan, H Zhu, T Berah, J Eaton, P Perez Guzman, et al. Report 13: Estimating the number of infections and the impact of non-pharmaceutical interventions on covid-19 in 11 european countries. 2020.
- [25] Joël Mossong, Niel Hens, Mark Jit, Philippe Beutels, Kari Auranen, Rafael Mikolajczyk, Marco Massari, Stefania Salmaso, Gianpaolo Scalia Tomba, Jacco Wallinga, et al. Social contacts and mixing patterns relevant to the spread of infectious diseases. *PLoS medicine*, 5(3), 2008.
- [26] Istituto Nazionale di Statistica. Popolazione insistente per studio e lavoro (https://www.istat.it/it/files//2020/03/Popolazione-insistente.pdf). Technical report, ISTAT, 2020.
- [27] Istituto Nazionale di Statistica. Commercio estero (https://www.istat.it/it/commercio-estero). Technical report, ISTAT, 2020.
- [28] Odo Diekmann, Johan Andre Peter Heesterbeek, and Johan AJ Metz. On the definition and the computation of the basic reproduction ratio r 0 in models for infectious diseases in heterogeneous populations. *Journal of mathematical biology*, 28(4):365–382, 1990.
- [29] Laura Fumanelli, Marco Ajelli, Piero Manfredi, Alessandro Vespignani, and Stefano Merler. Inferring the structure of social contacts from demographic data in the analysis of infectious diseases spread. *PLoS computational biology*, 8(9), 2012.
- [30] Jan Medlock and Alison P Galvani. Optimizing influenza vaccine distribution. Science, 325(5948):1705–1708, 2009.
- [31] Marc Baguelin, Albert Jan Van Hoek, Mark Jit, Stefan Flasche, Peter J White, and W John Edmunds. Vaccination against pandemic influenza a/h1n1v in england: a real-time economic evaluation. *Vaccine*, 28(12):2370–2384, 2010.