

Status of COVID-19 in Spain and recommendations

Scientific Report

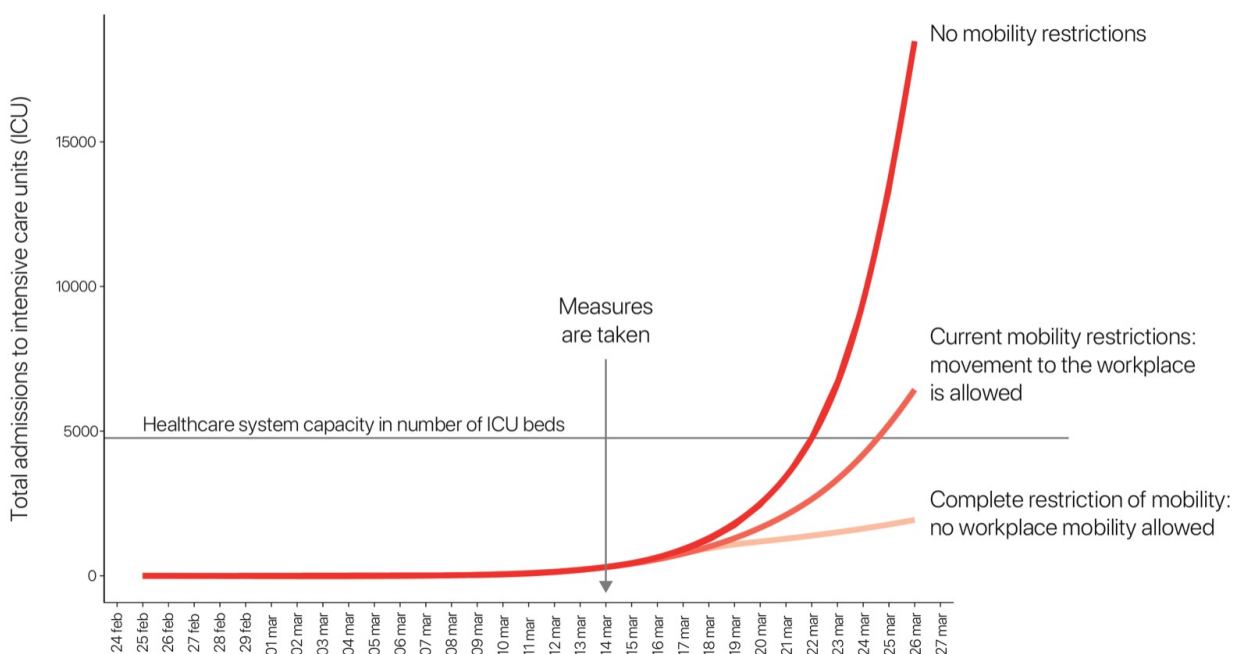
The undersigned are members of the scientific community, including 36 experts working in 25 research groups investigating the dynamic and spread of epidemics in the setting of infectious and respiratory diseases. We would like to express our concern about the limited capacity of actions taken by the Spanish Government to successfully control the SARS-CoV-2 outbreak and leave the exponential growth phase of new cases, once we can see the effects in the coming weeks. The measures taken so far, consisting primarily of partial restriction mobility, are in the right direction, although some researchers have warned about the pressure placed on the building blocks of the health system.¹

Given that the actual extent of an epidemic can only be assessed retrospectively, governments and policymakers are forced to make decisions based on mathematical models of other diseases and previous experiences of other countries taking different actions.² In the case of Spain, various models simulating the spread of infection and using different assumptions converge in a landscape of a high number of new cases within the next few weeks. Simulations have been conducted using the URV and UNIZAR models to predict the progression of the number of patients who will require hospitalization in intensive care units because of CoVID-19 in three scenarios (Supplementary material; mathematical method): scenario 1, no mobility restrictions; scenario 2, partial restriction of mobility (i.e., movement to the workplace allowed for 50% of people); scenario 3, complete restriction of mobility (i.e., no workplace mobility allowed except for essential services).

The model suggests that the actions taken to date will be insufficient to prevent hospitals and intensive care units from being overwhelmed in the next few weeks. The foreseen collapse of the health system, along with the age distribution of the Spanish population (i.e., 18% of people aged ≥ 65 years), warns about the potential high mortality rate associated with CoVID-19 in our

country. According to our models, the current state is associated with a very high risk of saturation of our health system, which is expected to occur by March 25 (Figure 1). Therefore, we urge the Spanish Government to implement, as swiftly as possible, more drastic measures to minimize the impact of the pandemic on the Spanish population.

Incidence curves for patients requiring ICU hospitalization due to CoVID-19 in Spain



As a reference framework (to be adjusted, if applicable), we suggest the following measures:

Establish regional categories according to the number of cases per 100,000 population and implement a package of multiple interventions that fit each category. For example, type A areas (≥ 25 cases/ 10^6 inhabitants) implement a complete shutdown of the region and citizen lockdown, except for essential services (e.g., hospitals, healthcare, and research centers) for a minimum period of 15 to 21 days. This category includes, at this moment, the Autonomous Communities of Madrid (132 cases/ 10^6 inhabitants), Castilla y León (27), Castilla-La Mancha (39), La Rioja (132), Basque Country (44), Navarra (59) and Catalonia (27). Type B areas (< 25 cases/ 10^6 inhabitants) apply partial confinement (50% of work activity and 25% of internal movement allowed) and

close monitoring of the growth rate of new infections; switch to type A in the event of a sharp rise in the number of cases. This category includes all regions not considered type A. Total interruption of non-essential inter-regional land, sea, and air transport for a minimum period of 15 days (e.g., until April 4, approximately) is also warranted.

To implement combined non-pharmacological interventions for several weeks, including complete restriction of movement, work interruption, and social distancing, banning all travel and all non-basic economic activities, together with the intensified use of diagnostic tests in suspected cases has proven to yield good results.³ Also, there is an urgent need to establish a purchasing and supply channel for personal protective equipment, which is currently insufficient for health personnel who are highly exposed to contagion and are prone to contagion. The recent finding on the spreading capacity of SARS-CoV-2 by contamination of eyelashes and hair reinforces this need.⁴

The proposed suppression policies will not mean the end of CoVID-19 in Spain in the initial three to four weeks; therefore, the development of strategies to sustain the gains is critical. A key lesson from South Korea is the need to create a robust surveillance system capable of collecting and reporting epidemiological data down to the individual or household level.⁵ There are two pillars for the development of such a system: first, increased laboratory capacity to perform PCR diagnostic tests on all individuals with symptoms for early isolation, and second, the development and implementation of a universal mobile application for self-reporting of observations suspicious of COVID-19. The identification of an increase in the number of cases in an area would trigger quick remedial measures like the implementation of early and targeted suppression actions.

Importantly, we beg the Spanish Government to facilitate the access of the scientific community to outbreak data, thus providing artificial intelligence support in simulation and modelling, and to create core support groups that coordinate a comprehensive, objective, and transparent scientific response.

References

1. Legido-Quigley H, Mateos-García JT, Campos VR, Gea-Sánchez M, Muntaner C, McKee M. The resilience of the Spanish health system against the COVID-19 pandemic. *Lancet Public Heal*. 2020 Mar;0(0).
2. European Centre for Disease Prevention and Control. Rapid risk assessment: Novel coronavirus disease 2019 (COVID-19) pandemic: increased transmission in the EU/EEA and the UK – sixth update – 12 March 2020. Stockholm [Internet]. 2020. Available from: <https://www.ecdc.europa.eu/sites/default/files/documents/RRA-sixth-update-Outbreak-of-novel-coronavirus-disease-2019-COVID-19.pdf>
3. Li R, Pei S, Chen B, Song Y, Zhang T, Yang W, et al. Substantial undocumented infection facilitates the rapid dissemination of novel coronavirus (COVID-19). *medRxiv*. 2020 Mar 16;2020.02.14.20023127.
4. Yan Y, Chen H, Chen L, Cheng B, Diao P, Dong L, et al. Consensus of Chinese experts on protection of skin and mucous membrane barrier for health professions fighting against coronavirus disease 2019. *Chinese J Dermatology*. 2020 Mar 13;
5. COVID-19 National Emergency Response Center, Epidemiology & Case Management Team KC for DC& P. Contact Transmission of COVID-19 in South Korea: Novel Investigation Techniques for Tracing Contacts. *Osong public Heal Res Perspect*. 2020 Feb;11(1):60–3.

Signatures

Surname, Name	Institution	Positiona
Arenas, Alex	Universitat Rovira i Virgili, Tarragona	<i>Director de Cátedra Ciencia y Humanismo; Editor Asociado, Physical Review</i>
Ares, Saúl	Department of Systems Biology, Centro Nacional de Biotecnología (CSIC), Madrid	<i>Científico Titular</i>
Benlloch, José María	I3M (CSIC-UPV), Valencia	<i>Director</i>
Bruña Romero, Óscar	Departamento de Microbiologia, Inmunologia y Parasitología, Universidad Federal de Santa Catarina, Brasil	<i>Associate Professor</i>
Buceta, Javier	Bioengineering Department, Chemical and Biomolecular Engineering Department, Lehigh University, USA	<i>Associate Professor</i>
Cale, Malu	Research Group in Bioinformatics and Medical Statistics, Facultat de Medicina i Facultat de Ciències de la Salut i Benestar, Universitat de Vic – Universitat Central de Catalunya, Vic.	<i>Associate Professor</i>
Casabona, Jordi	CIBERESP-UAB, Barcelona	<i>Director CEEISCAT</i>
Casals, Martí	Facultat de Medicina i Facultat de Ciències de la Salut i Benestar a la Universitat de Vic – Universitat Central de Catalunya	<i>Profesor</i>
Caylà, Joan	Fundació de la Unitat d'Investigació en Tuberculosi de Barcelona	
Chaccour, Carlos	Institut de Salut Global de Barcelona	<i>Assistant Research Professor</i>
Clotet, Bonaventura	Instituto Investigación Germans Trias i Pujol	<i>Director IrsiCaixa</i>
Cota, Wesley	Universidade Federal de Viçosa, Minas Gerais, Brasil	<i>PhD Candidate</i>
Cuesta, José A.	Universidad Carlos III de Madrid	<i>Catedrático de Universidad</i>
de León, Manuel	Instituto de Ciencias Matemáticas (ICMAT-CSIC) y Real Academia de Ciencias	<i>Fundador y Profesor de Investigación</i>
Domingo Solans, Esteban	Profesor "ad honorem" del CSIC, Centro de Biología Molecular Severo Ochoa.	<i>Profesor de Investigación</i>
Fernández Oliva, Albert	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigador predoctoral</i>

Fernández Soto, Daniel	Centro Nacional de Biotecnología (CSIC), Madrid	<i>PhD Candidate</i>
Figueras, Antonio	Instituto de Investigaciones Marinas (CSIC), Madrid	<i>Profesor de Investigación</i>
Gallego Cámara, Beatriz	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigadora postdoctoral</i>
Garcia-Aymerich, Judith	Institut de Salut Global de Barcelona	<i>Jefa de Programa, Enfermedades no transmisibles y medio ambiente</i>
Gasset, Maria	Instituto Química-Física Rocasolano (CSIC)	<i>Investigadora científica (Associate professor)</i>
Gómez Melis, Guadalupe	GRBIO: Research Group in Biostatistics and Bioinformatics, Departament d'Estadística i Investigació Operativa, Universitat Politècnica de Catalunya-BarcelonaTech	<i>Professor of Statistics</i>
Gómez Rodríguez, Carmen	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Research Scientist</i>
Gómez-Gardeñes, Jesús	Universidad de Zaragoza, Zaragoza	<i>Profesor Titular</i>
Gómez, Sergio	Universitat Rovira i Virgili, Tarragona	<i>Profesor Titular</i>
Granell, Clara	Universidad de Zaragoza, Zaragoza	<i>Investigadora postdoctoral</i>
Guillén, Francisco	de Medicina Preventiva y Salud Pública, Facultad de Ciencias de la Salud, Universidad Pública de Navarra, Pamplona.	<i>Catedrático de Universidad</i>
Gutiérrez Álvarez, Francisco J.	Department of Molecular and Cell Biology (Coronavirus Laboratory), Centro Nacional de Biotecnología, Madrid	<i>Investigador postdoctoral</i>
Hueso-Gil, Ángeles	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigadora predoctoral</i>
Jarrillo-Herrero, Pablo	Massachusetts Institute of Technology, Boston, MA, USA	<i>Cecil and Ida Green Professor of Physics</i>
López, Leonardo Rafael	Institut de Salut Global de Barcelona	<i>Investigador postdoctoral</i>
Lowe, Rachel	London School of Hygiene & Tropical Medicine, Londres, Reino Unido	<i>Associate Professor</i>
Lozano, Andrés	University of Toronto	<i>Jefe de Neurocirugía</i>
Maiques, Ana	Neuroelectronics, USA	<i>CEO</i>
Manrubia, Suana	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigadora científica (Associate professor)</i>
Martín Buldú, Jaime	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigador principal</i>

Martín-Benito, Jaime	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Professor</i>
Martínez González, Miguel Ángel	Universidad de Navarra y Harvard University - Harvard School of Public Health	<i>Catedrático de Medicina Preventiva y Salud Pública y Catedrático adjunto</i>
Martínez Sánchez, José María	Universitat Internacional de Catalunya	<i>Responsable del Grupo de Evaluación de Determinantes de la Salud y Políticas Sanitaria</i>
Matamalas, Joan T.	Harvard Medical School, Boston, MA, USA	<i>Research Associate</i>
Mérida, Isabel	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigadora científica (Associate professor)</i>
Mitjà, Oriol	Instituto Investigación Germans Trias i Pujol	<i>Associate Professor, European Research Council researcher</i>
Morales, Maria	Departamento de Medicina Preventiva y Salud Pública, Universitat de València	<i>Catedrática</i>
Moreno del Alamo, María	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigadora postdoctoral</i>
Pascual-Leone, Álvaro	Center for Memory Health, Hebrew Senior Life, Boston, MA, USA	<i>Senior scientist</i>
Pereda, María	Departamento Ingeniería de Organización, Administración de empresas y Estadística, Universidad Politécnica de Madrid	<i>Assistant Professor</i>
Pérez Losada, Jesús	Instituto de Biología Molecular y Celular del Cáncer de Salamanca	<i>Científico titular del CSIC</i>
Poyatos, Juan F.	Logic of Genomic Systems Laboratory. CNB-CSIC	<i>Group Leader</i>
Puig Reixac, Maria Teresa	Hospital de la Santa Creu i Sant Pau, Universitat Autònoma de Barcelona, Barcelona.	<i>Associate Professor</i>
Real, Jordi	Unidad Bioestadística, USR Barcelona DAP-CAT	<i>Bioestadista</i>
Risco Ortiz, Cristina	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Directora del Laboratorio de Estructura Celular</i>
Rodó, Xavier	Institut de Salut Global de Barcelona	<i>ICREA, Jefe de Programa, Clima y Salud</i>
Ruffini, Giulio	Starlab Barcelona, Neuroelectronics Corp USA	<i>CEO, President, European Research Council researcher</i>
Sardanyés, Josep	Centre de Recerca Matemàtica (CRM)	<i>Investigador Ramón y Cajal</i>
Sánchez Villegas, Almudena	Departamento de Ciencias Clínicas, Universidad de las Palmas de Gran Canaria, Las Palmas de Gran Canarias	<i>Catedrática,</i>
Seoane Blanco, Mateo	Centro Nacional de Biotecnología (CSIC), Madrid	<i>PhD Candidate</i>

Solé, Ricard	Complex Systems Lab- UPF, Barcelona	<i>ICREA Research Professor</i>
Soriano-Paños, David	Universidad de Zaragoza, Zaragoza	<i>Investigador postdoctoral</i>
Steinegger, Benjamin	Universitat Rovira i Virgili, Tarragona	<i>PhD Candidate</i>
Sunyer, Jordi	Institut de Salut Global de Barcelona	<i>Jefe de Programa, Infancia y Medio Ambiente</i>
Tamames, Javier	Centro Nacional de Biotecnología (CSIC), Madrid	<i>Investigador</i>
Tobías, Aurelio	Instituto de Diagnóstico Ambiental y Estudios del Agua (CSIC - IDAEA), Barcelona	<i>Investigador Científico (Associate professor)</i>
Toledo, Estefania	Centro de Investigación Biomédica en Red- Fisiopatología de la Obesidad y Nutrición, Universidad de Navarra.	<i>Associate Professor</i>
Valls, Joan	Institut de Recerca Biomèdica de Lleida	<i>Experto en Estadística y Metodología de la Investigación</i>
Valverde, Sergi	Institut de Biologia Evolutiva (CSIC-UPF), Barcelona	<i>CSIC Tenured Scientist</i>
Vilà-Freixa, Jordi	Departamento de Ciencias Básicas, Universitat Internacional de Catalunya	<i>Vicerrector de investigación</i>
Vioque, Jesús	Universidad Miguel Hernandez	<i>Catedrático de Medicina Preventiva y Salud Pública</i>

Coordinating author: Oriol Mitja, Instituto Investigación Germans Trias i Pujol

Supplementary material – Mathematical model

The simulation was conducted using the last version of a family of discrete-time epidemiological models,¹ specifically tailored to describe the transmission dynamics of SARS-COV-2, the etiological agent of COVID-19. The model aims to estimate the risk rate for each Spanish municipality, considering the following parameters: (1) the transmission dynamics of SARS-COV-2, (2) the usual movement patterns of the Spanish population, and (3) the demographics of the Spanish population.

Transmission dynamics

Virus transmission was described using a compartmental model that groups the population according to the infection status into the following categories:

- **Susceptible:** individuals without COVID-19, but still at risk.
- **Exposed:** infected individuals within the incubation phase and no transmission capacity.
- **Asymptomatic:** infected individuals without clinically relevant symptoms but transmission capacity.
- **Infected:** individuals with symptoms that are likely to be attributed to COVID-19.
- **Hospitalized:** infected individuals who have been identified and admitted to a hospital, thus blocking the transmission capacity.
- **Recovered:** individuals who have been infected but cannot transmit the disease because they have either died or recovered and developed immunity.

Movement patterns of the Spanish population

Transitions between the infection states drive the probabilities of transmission, recovery, etc., obtained from COVID-19 studies published to date. Regarding movements, we have included data from the National Institute of Statistics (NIS) on travel to the workplace between and within towns.² The NIS record includes all movements between Spanish municipalities greater than 100 inhabitants and reporting more than 10 trips. The inclusion of individual movements into the model is essential to understand how the virus spreads across the country and allows simulating the outcome of movement restrictions both globally and locally.

Spanish demographics

The Spanish population has been divided into three age groups:

- Young (from 0 to 25 years old).
- Adult (from 26 to 65 years old).
- Older (more than 65 years old).

The rationale for age grouping has been based on the recent evidence on COVID-19 that shows a different effect of the infection in each age group.⁴ In our model, the following differences between age groups have been considered:

- Young and older are less likely to move across the territory than adults.
- Young are more likely to experience an asymptomatic disease (or with mild symptoms) and are, therefore, more challenging to identify.
- Older people are more likely to require hospitalization than young and adults.

Limitations

- The model does not predict or consider international inputs of infected individuals.
- The model has been based on epidemiologic parameters reported to date; however, these parameters may change in the near future.
- The model assumes the movement data reported by the NIE, which may vary in case of mobility restrictions.

Strengths

- The model allows modifying the epidemiologic parameters as new reports come up.
- The model allows for investigating the influence of the asymptomatic period and associated infectivity.
- Based on the current parameters, we can build risk maps of new cases and foresee infection spread by asymptomatic subjects.
- General mobility restrictions can be quickly introduced into the model to yield new predictions with these assumptions. This feature is of particular interest for policymakers

and health authorities, which will be able to explore the foreseen impact of the previewed restrictions on infection spread.

References

1. D. Soriano-Panos, L. Lotero A. Arenas and J. Gomez-Gardenes, Spreading processes in multiplex metapopulations containing different mobility networks, *Physical Review* 2018, X 8, 031039.
2. D. Soriano-Panos, G. Ghoshal, A. Arenas and J. Gomez-Gardenes Impact of temporal scales and recurrent mobility patterns on the unfolding of epidemics, *Journal of Statistical Mechanics: Theory and Experiment* 2020; 2, 024006.
3. Critical regimes driven by recurrent mobility patterns of reaction-diffusion processes in networks, J. Gomez-Gardeñes, D. Soriano-Paños and A. Arenas, *Nature Physics* 2018 14, 391–395.
4. Dawei Wang, Bo Hu, Chang Hu, et Clinical Characteristics of 138 Hospitalized Patients With 2019 Novel Coronavirus–Infected Pneumonia in Wuhan, China. *JAMA* 2020;323(11):1061-1069.