

The COVID-19 pandemic and the response by the scientific community in Luxembourg





Chambre de Commerce, Luxembourg 17 November 2021

Welcome



Muriel Bouchet, Director, Fondation IDEA





Rolf Tarrach, President, Les Amis de l'Université



The COVID-19 pandemic and the response by the scientific community in Luxembourg



Prof. Dr Wilmes, Professor of Systems Ecology, Luxembourg Centre for Systems Biomedicine



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COVID-19 pandemic The response by the scientific community in Luxembourg

Paul Wilmes

Luxembourg Centre for Systems Biomedicine **Department of Life Sciences and Medicine** University of Luxembourg



17th November 2021

E GOUVERNEMENT DU GRAND-DUCHÉ DE LUXEMBOURG

INSTITUTE













Direction de la santé

12th March 2020

can we deale peoples' exposures? How can we guarantee enough test capacity? How infectious is this Virus?

How do we organize respirators

Do we have enough intensive care units?

Will we be able to sustain food supp

HOW CAN WE PROTECT OUR FAMILIES AND OURSELV

What is the mortality rate?

What is the impact on logistics? Where do we get our medical personnel?

How can we organise medical and diagnostic supply?

Who is most vulnerable?

How many people are already infected?

to no treat the interiod

Are there drugs out there that help?

How many cases can How can the virus be detected?



CE COVID-19 Task Force A first in Luxembourg's history



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Direction de la santé





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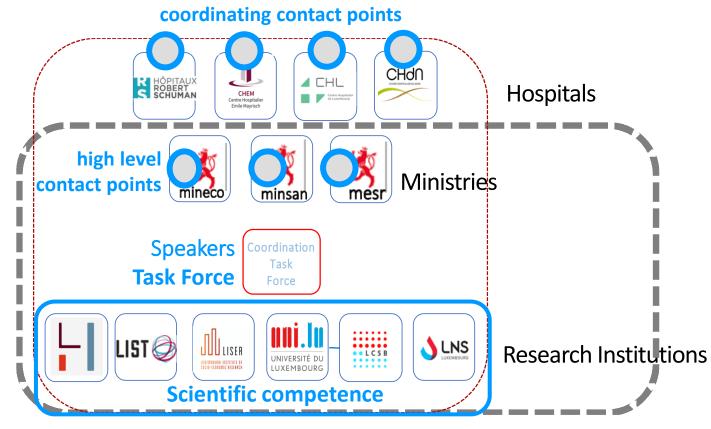


The COVID-19 Task Force

was set up in order to offer the health system and the country the **combined expertise** available within the Luxembourg public research sector supported by the Ministry of Higher Education and Research







SUPPORT TASK COORDINATION BETWEEN MINISTRIES & RESEARCH COMMUNITY PLUS HOSPITALS

- Defined contact points in partner organisations allowed for fast decision making
- Fast initiation of projects from all access points possible
- Straight forward escalation in complex decision processes





Coordination & direction





Spokesperson: Prof. Ulf Nehrbass, CEO Luxemborg Institute of Health



Co-Spokesperson: Prof. Paul Wilmes Professor of Systems Ecology, LCSB

Operational lead: Frank Glod (LIH)

Members: Henry-Michel Cauchie (LIST)

Frank Glod (LIH; coordination: WPs 1, 2, 3 & 4)

Lars Geffers (LIH; coordination: WPs 8 & 9)

Jasmin Schulz (LIH; coordination: WPs 5, 10 & 11)

Paul Wilmes (UL; coordination: WPs 6, 7, 12 & 13)



WORK PACKAGES

Based on a list of priority requests from the ministries

Deputy: Francesco Ferrero

	Workpackage leader		Workpackage leader
Cross-sectional study infection prevalence in Luxembourg	Rejko Krüger	Gauging economic impact of the COVID-19 outbreak	Aline Muller
Predictive markers for COVID-19 severity	Laetitia Huiart Markus Ollert & Guy Fagherazzi	WP8 Mobilising volunteers for support of hospital emergency services	Gilbert Massard
WP3 Interventional clinical trial with existing drugs	Laetitia Huiart & Guy Berchem	WP9 Mobilising and coordinating private partner initiatives	Sasha Baillie
Diagnostic capacity and large-scale testing strategies for Luxembourg	Markus Ollert	COVID-19 centred communication	Didier Goossens/ Deputy: Arnaud D'Agostini
eHealth solutions for hospitalised and ambulatory patients	Damien Dietrich	WP11 Evidence-based review team in the outbreak context	Dirk Brenner
WP6 Statistical pandemic projections	Rudi Balling/ Deputy: Alex Skupin	WP12 Ideas for new initiatives in the pandemic context	Marc Schiltz
		WP13 Logistics and supply chains	Benny Mantin Deputy: Francesco Ferrero



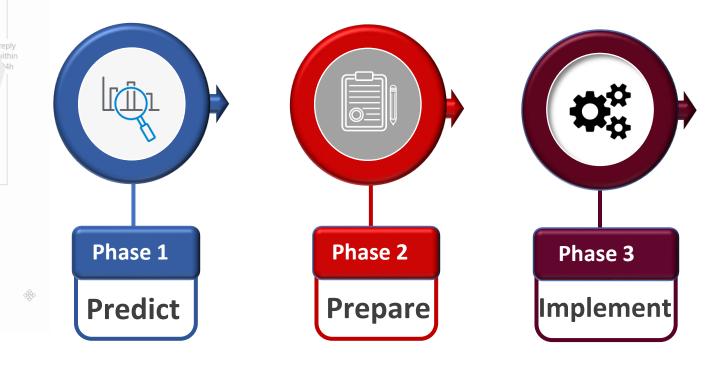


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WP0: Proactive mitigation

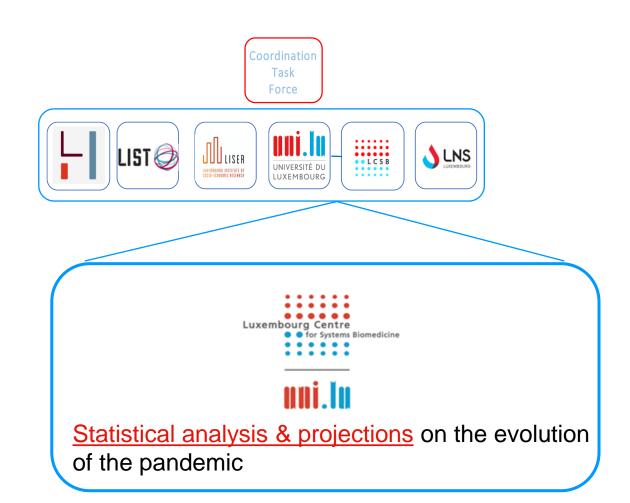
Contribution of various WPs to the same question When and how can we transition from lockdown to revitalise economic activity?



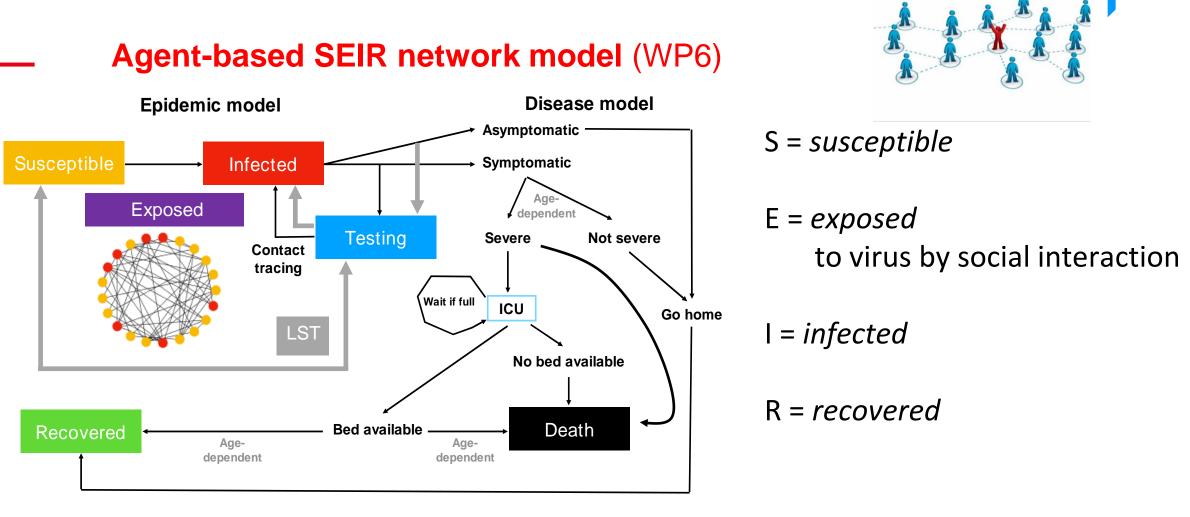










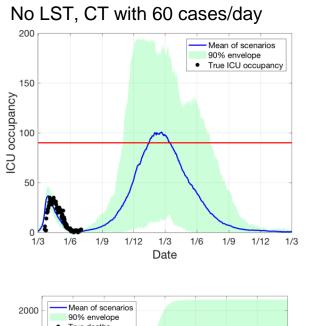


Population level: Epidemic predictions

 \rightarrow Epidemic driven by interaction networks

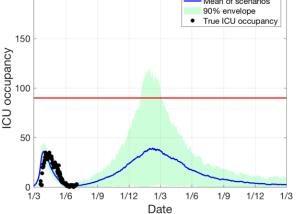
Effect of Large-Scale Testing 53k tests/week

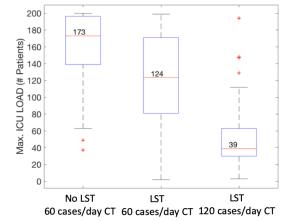
Mass screening and contact tracing to minimize second wave

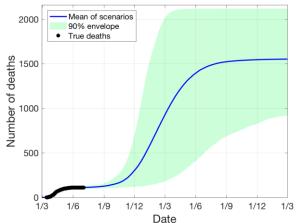


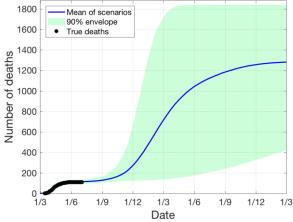
LST, CT with 60 cases/day 200 Mean of scenarios 90% envelope True ICU occupancy 150 occupancy ICU 50 1/3 1/6 1/9 1/12 1/3 1/6 1/9 1/12 1/3 Date

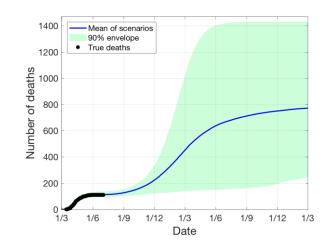
LST, CT with 120 cases/day

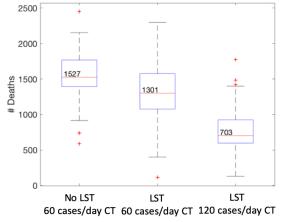








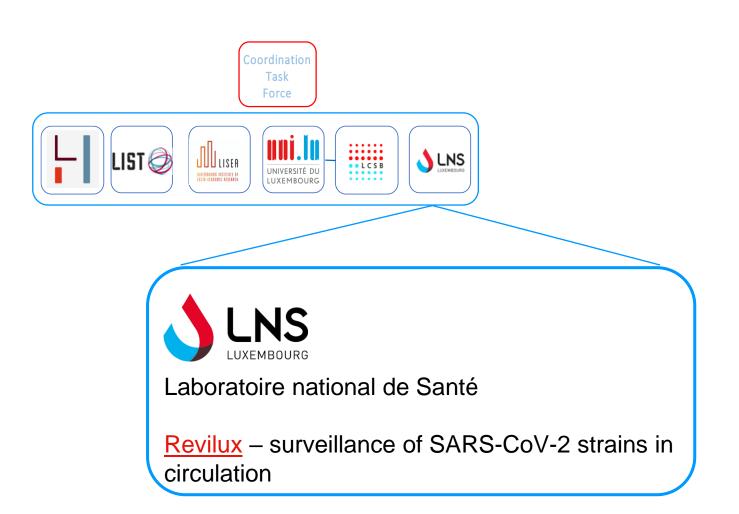










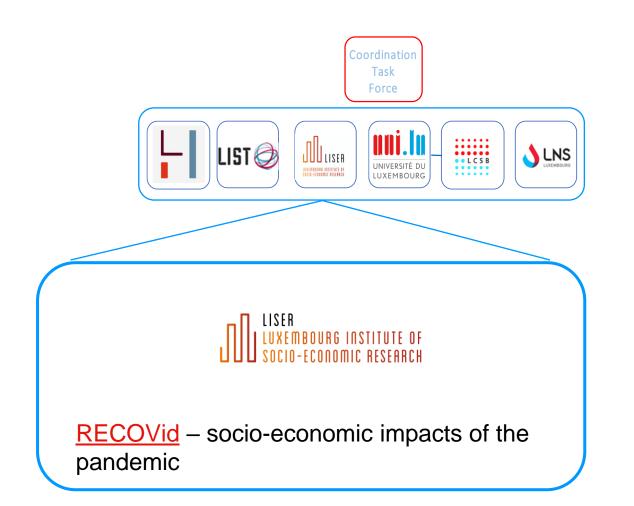










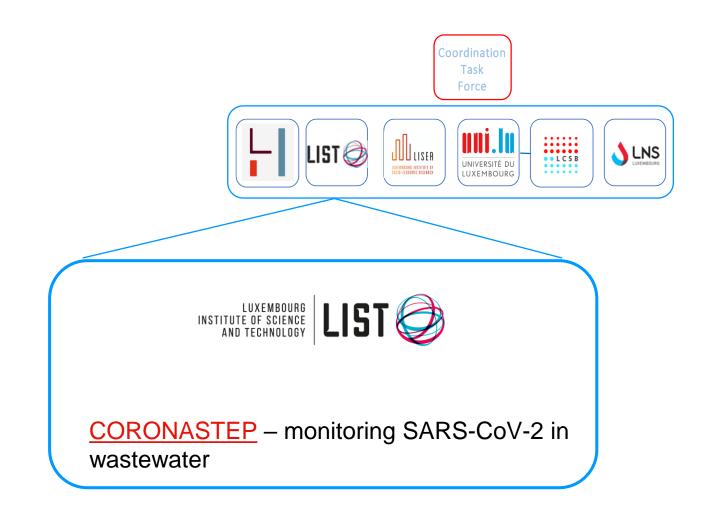


RESEARCH













<mark>.≬_</mark> water



Article Genome Sequencing of SARS-CoV-2 Allows Monitoring of Variants of Concern through Wastewater

Malte Herold ¹⁽¹⁾, Aymeric Fouquier d'Hérouël ², Patrick May ²⁽³⁾, Francesco Delogu ², Anke Wienecke-Baldacchino ³, Jessica Tapp ³, Cécile Walczak ¹, Paul Wilmes ^{2,4}⁽³⁾, Henry-Michel Cauchie ¹⁽³⁾, Guillaume Fournier ³⁽¹⁾ and Leslie Ogorzaly ^{1,*}⁽⁵⁾

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- ² Luxembourg Centre for Systems Biomedicine, University of Luxembourg, 7 Avenue des Hauts-Fourneaux, L-4362 Esch-sur-Alzette, Luxembourg; aymeric.dherouel@uni.lu (A.F.d.); patrick.may@uni.lu (P.M.); francesco.delogu@uni.lu (F.D.); paul.wilmes@uni.lu (P.W.)
- ³ Laboratoire National de Santé, Department of Microbiology, 1 Rue Louis Rech, L-3555 Dudelange, Luxembourg; Anke.Wienecke-Baldacchino@lns.etat.lu (A.W.-B.); Jessica.TAPP@lns.etat.lu (J.T.); Guillaume.Fournier@lns.etat.lu (G.F.)
- ⁴ Department of Life Sciences and Medicine, Faculty of Science, Technology and Medicine, University of Luxembourg, 2 Avenue de l'Université, L-4362 Esch-sur-Alzette, Luxembourg
- * Correspondence: leslie.ogorzalv@list.lu

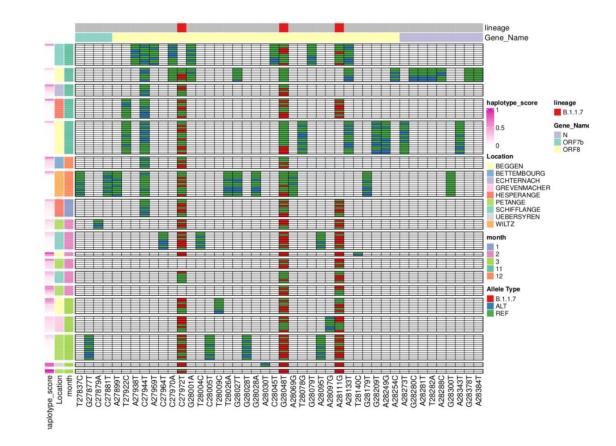
Water 2021, 13(21),3018. 27 October 2021 https://doi.org/10.3390/w13213018

Research supported by the Fondation André Losch and FNR

FONDATION Fonds National de la Recherche Luxembourg

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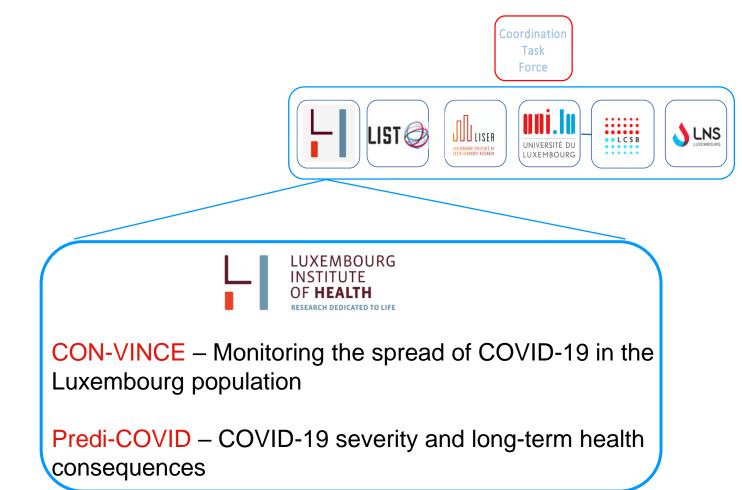


- **Signature mutations** from individual SARS-CoV-2 lineages in wastewater samples in Luxembourg
- Wastewater-based epidemiology









RESEARCH

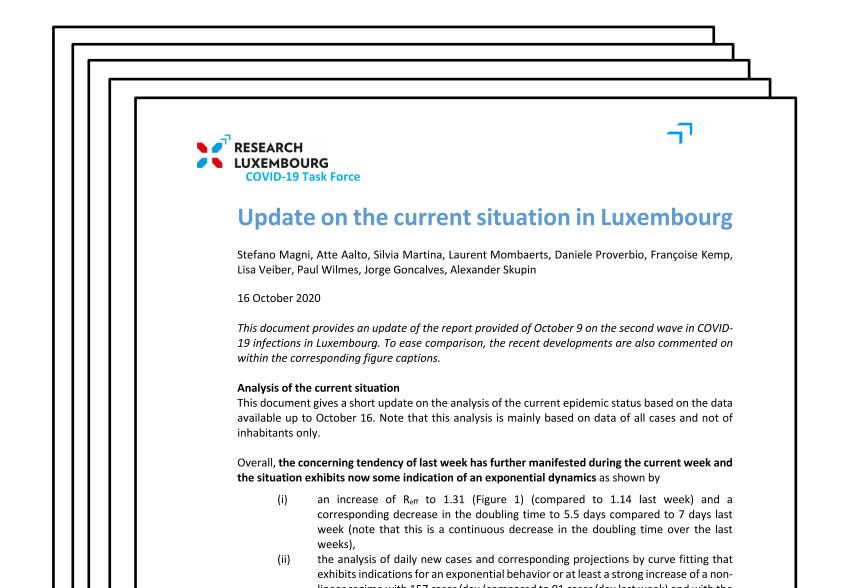


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Weekly briefings for the Government



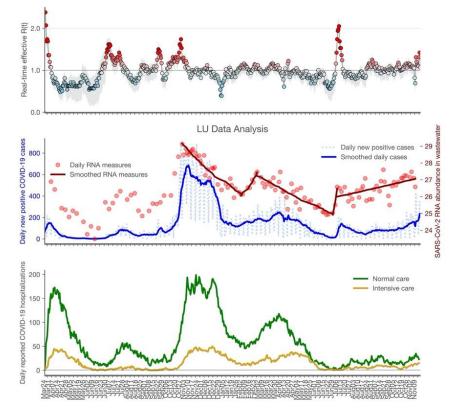




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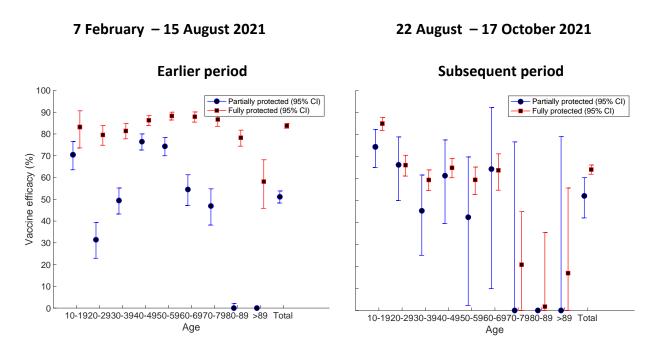
Current situation in Luxembourg

1 Significant increase in daily cases



Research Luxembourg COVID-19 Task Force report - 11 November 2021

2 Waning following vaccination



- Vaccine effectiveness is dropping
- Booster shots (especially for the elderly) is essential to keep population immunity high









Proactive mitigation including large-scale testing



LE GOUVERNEMENT DU GRAND-DUCHÉ DE LUXEMBOURG Ministère de la Santé

LE GOUVERNEMENT DU GRAND-DUCHÉ DE LUXEMBOURG Ministère de l'Enseignement supérieur et de la Recherche









Luxembourg National Research Fund

Direction de la santé

ALL WE NEED



Context planning phase in April 2020

At the onset of the large-scale testing planning phase, the following points of the COVID-19 pandemic were decisive for our strategy:

- 1. The **SARS-CoV-2 virus exhibited a very high infectivity**, with R₀ values well above influenza
- 2. The **case mortality rate was unclear**, and depending on the early overall test rate fluctuated between 2% and 10%
- 3. The **disease aetiology was unclear**, as were clinical treatment strategies and options
- 4. It became obvious that infected individuals were infectious even if they did not develop symptoms of the disease at all (presymptomatic and asymptomatic transmission)

The last point drew into question classical pandemic containment strategies and **required a consequential, new approach**





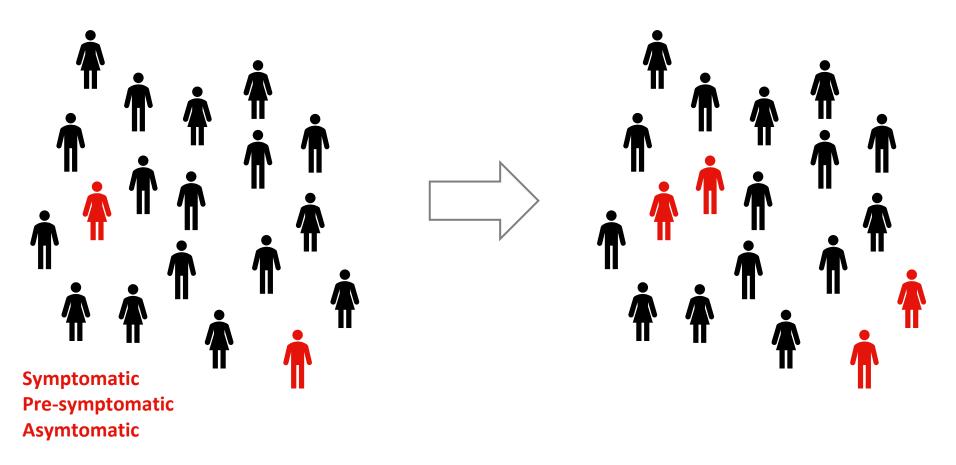
The following conclusions were reached:

- 1. An **rRT-PCR testing strategy** to track down the virus and break infection chains had to include not only symptomatic but also presymptomatic and asymptomatic individuals
- 2. Given the high infectivity and potential fatality rates, the test capacity had to be sufficient to potentially reach a **capacity of 10% of the population per week**
- **3.** Antibody testing was not considered a priority, as the initial goal was to stop pandemic spread into the population, herd immunity was not an option
- 4. This level of PCR testing capacity required a **high-performance PCR pooling process**
- 5. High **operational competence** was needed to guarantee a smooth pre-analytic delivery



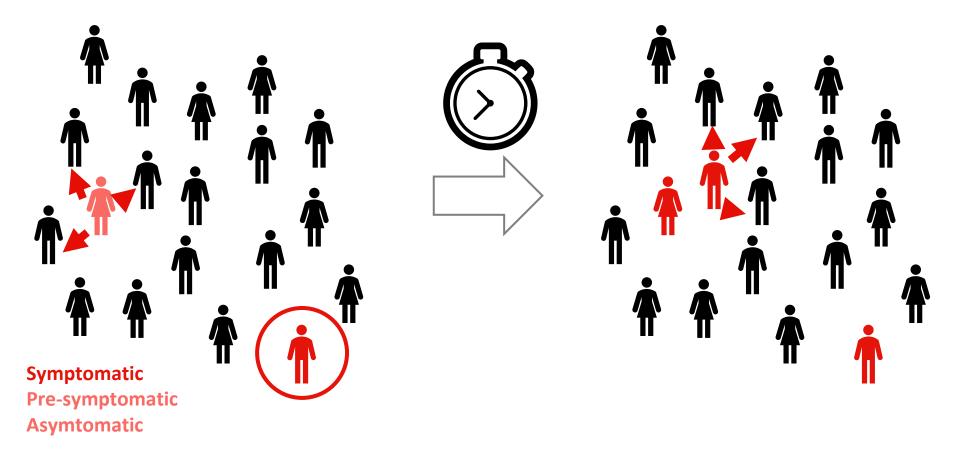


Infection chains





Wilmes*, ..., Nehrbass* (2021) *The Lancet Regional Health – Europe* **4**:100056. doi:10.1016/j.lanepe.2021.100056 Muller (2021) *The Lancet Regional Health – Europe* **4**:100082. doi:10.1016/j.lanepe.2021.100082 • Continuous triggering and sustaining of infection chains



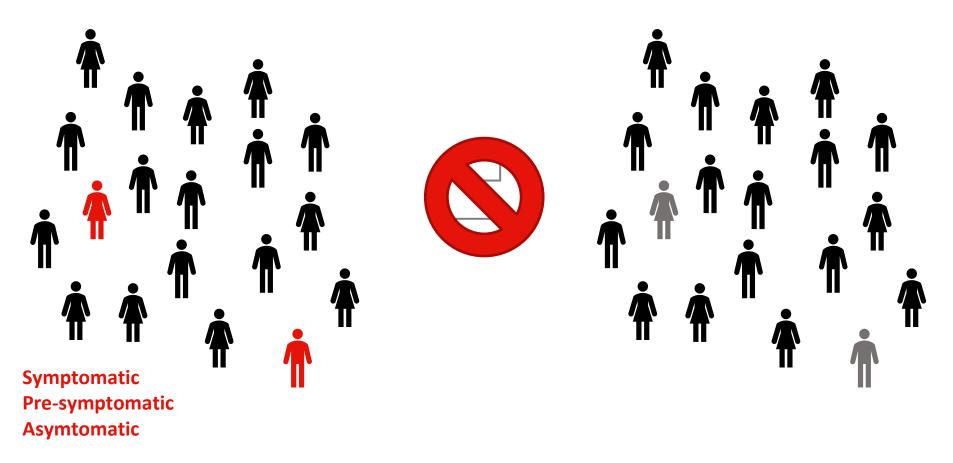


Wilmes*, ..., Nehrbass* (2021) *The Lancet Regional Health – Europe* **4**:100056. doi:10.1016/j.lanepe.2021.100056 Muller (2021) *The Lancet Regional Health – Europe* **4**:100082. doi:10.1016/j.lanepe.2021.100082





Rationale for broad testing



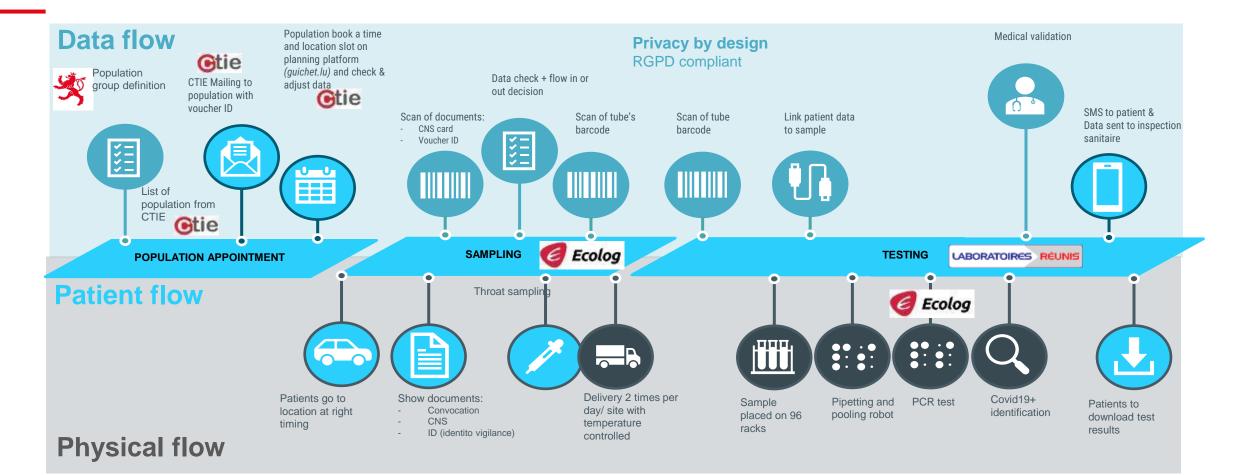


Wilmes*, ..., Nehrbass* (2021) *The Lancet Regional Health – Europe* **4**:100056. doi:10.1016/j.lanepe.2021.100056 Muller (2021) *The Lancet Regional Health – Europe* **4**:100082. doi:10.1016/j.lanepe.2021.100082



Testing - process





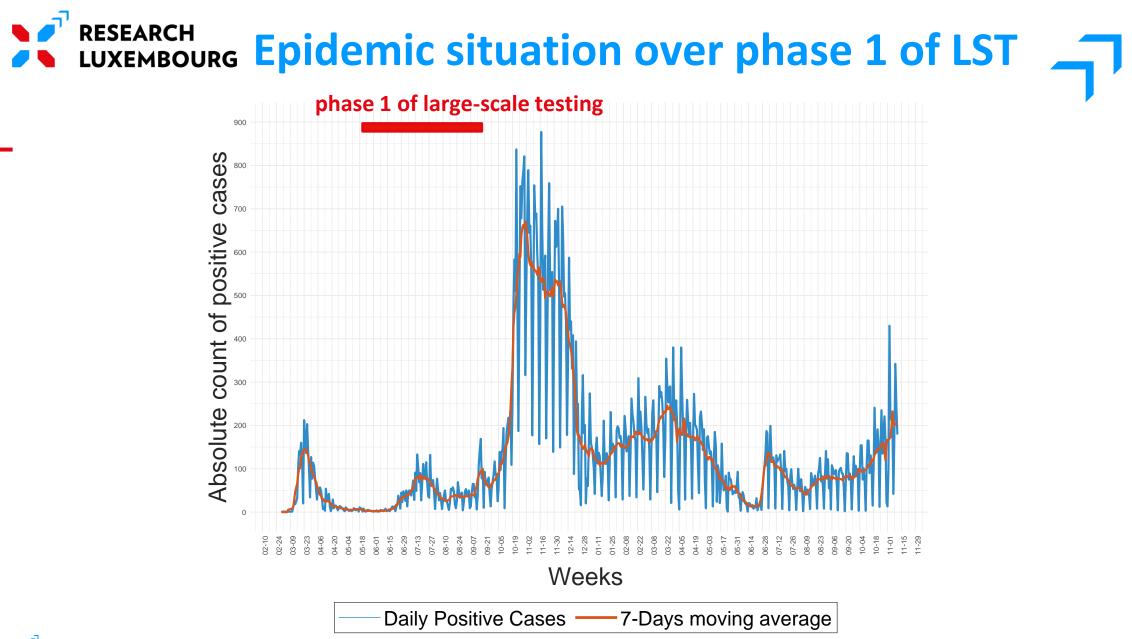




Testing station Grevenmacher, 14th September 2020



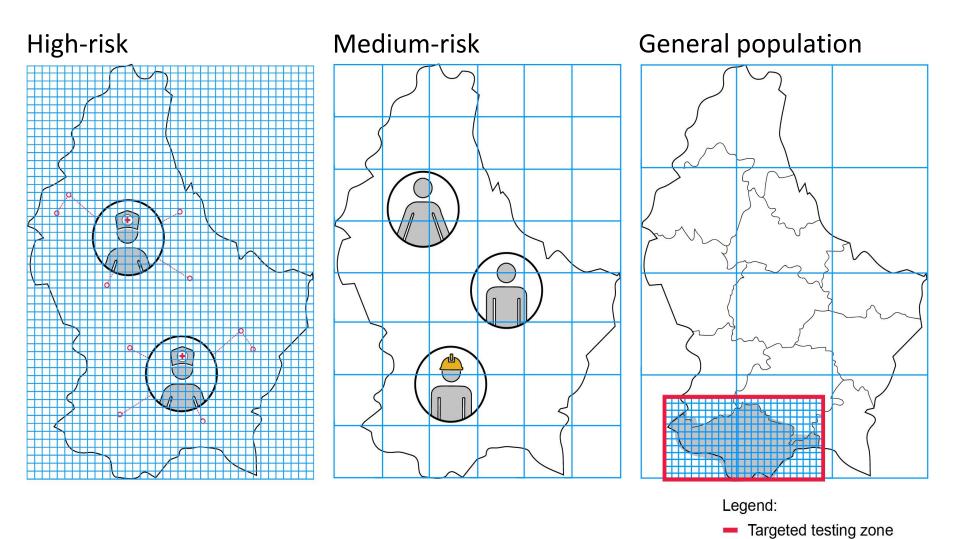
LST hand-over, 15th September 2020



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Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056







Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056

O Focus on population group



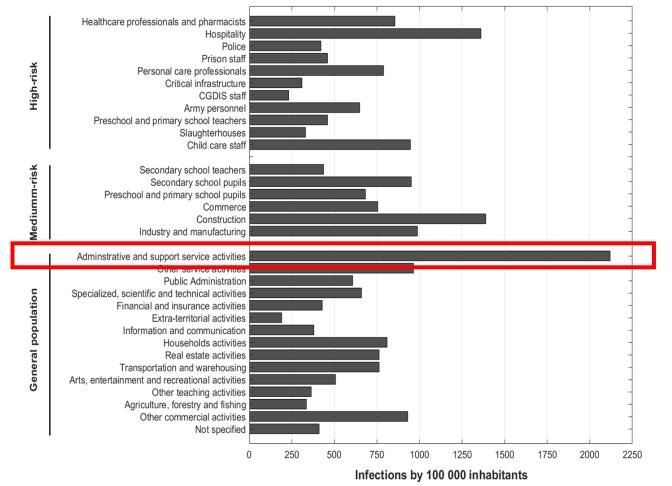
Category	Number	
Overall number of LST invitations	1,436,00	
Number of residents that participated	307,751	
Number of cross-borders that participated	87,198	
Total LST tests performed	566,320 (70% of all tests performed over the period)	
Positive LST cases	850	
Positive cases from LST-based contact tracing	249	
Positive cases directly and indirectly due to LST	1,099	

Effective capacity allowed testing 10% of population/week

- Three-pronged strategy addressing risk groups, general population and regional hot spots
- The test campaign turned out to have worked almost flawlessly, producing high-quality, reliable test results







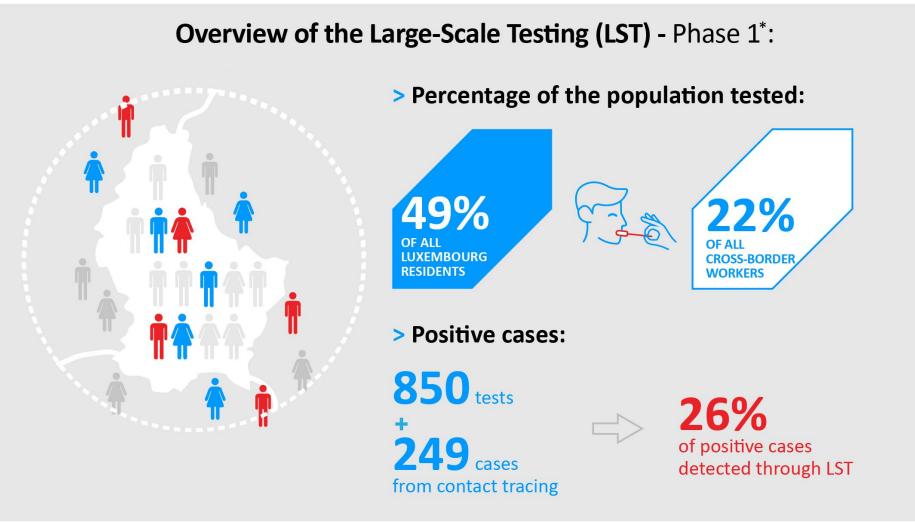
Sectors with high prevalence were Luxembourg-specific

- Administrative and support service activities as well as the construction sector had highest incidences
- Other sectors included in LST high risk group did not stand out (police, pharmacists, ...)

RESEARCH

Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056





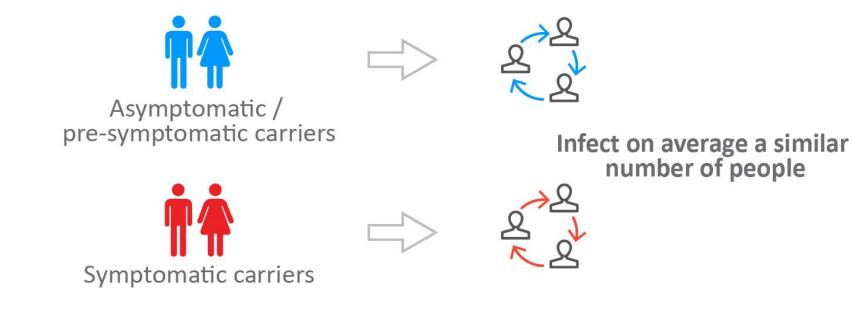


*Phase 1: 27th May – 15th September, 2020

Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056



Asymptomatic SARS-CoV-2 carriers result in a similar number of positive contacts





Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056



Greater odds of being tested positive for:

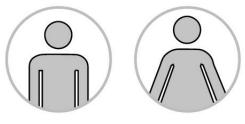


Activity sectors with high social connectivity





Individuals with a disposable income

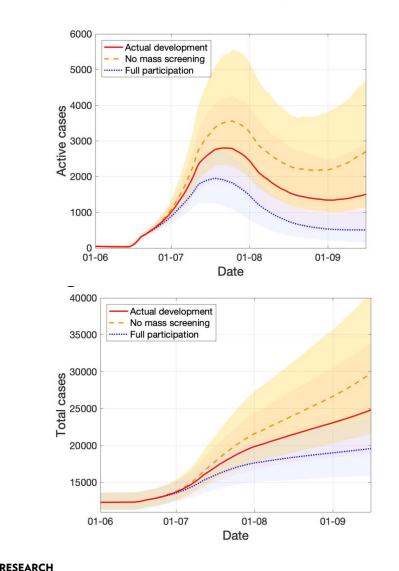


< 30k€/year



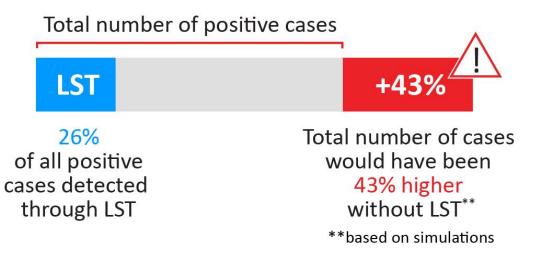
Wilmes*, ..., Nehrbass* (2021) The Lancet Regional Health – Europe 4:100056. doi:10.1016/j.lanepe.2021.100056





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> Efficacy of LST



Opportunity cost is compelling:

- Lockdown: €3,200 per Luxembourg resident
- Test: **€30**

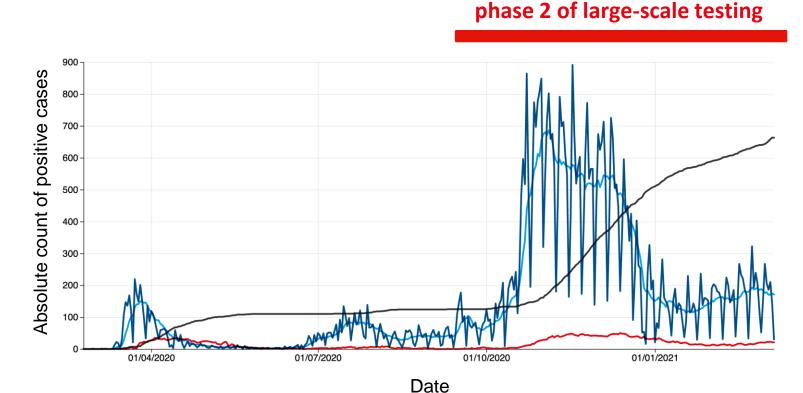




DU GRAND-DUCHÉ DE LUXEMBOURG

Direction de la santé

- 2.1 million invitations
- Participation rate: around 30 %
- 9,200 positive cases: around 20 % of cases





Comparison – Liverpool (Slovakia)

- 6 months: 6 November 2020 30 April 2021
- 283,338 (57%) residents took rapid lateral flow test (LFT); 47% had more than one test (27% of residents)
- 152,609 residents took a PCR test either because they had symptoms or to confirm a positive asymptomatic LFT result
 - 6,300 individuals (0.9 %) declaring no symptoms were tested positive by LFT (case positivity 2.1%)
 - 22,567 individuals declaring symptoms tested positive by PCR (case positivity 14.8%)

Liverpool Covid-SMART Community Testing Pilot – Evaluation Report; https://www.liverpool.ac.uk/media/livacuk/research/Mass,testing,evaluation.pdf

• Impact:

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> 21% reduction in cases up to mid-Dec. vs control areas

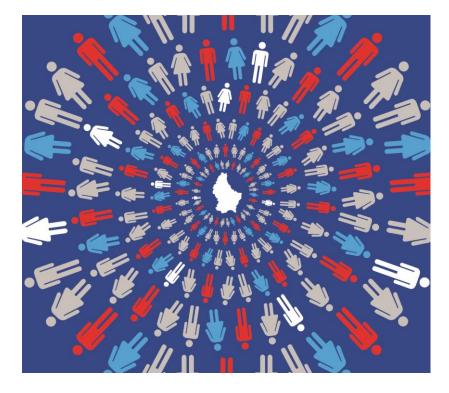
- Test uptake was lower and infection rates were higher in deprived areas
- Slovakia: 65 % of population, 1 week, decline of prevalence by 80 %

Second round

García-Fiñana et al. (2021) BMJ 374:1637. doi: 10.1136/bmj.n1637.

Pavelka et al. (2021) Science 372:635. doi: 10.1126/science.abf9648.





- Not much difference in viable virus between symptomatic and asymptomatic individuals
- 3/100,000 asymptomatics become longterm carriers
- May become **virus reservoirs**, with the potential to cause **recurrent outbreaks**
- During acute phase of a COVID-19 wave, asymptomatic individuals should definitively be included in the testing strategy and their contacts traced
- Need to learn more about the role of presymptomatic carriers







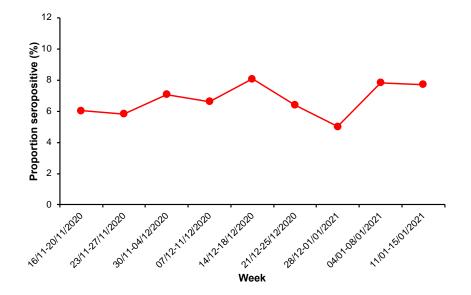
- Incentivisation: challenge
- Age, altruism, conformism, the tendency to abide by government-imposed rules, concern about contracting COVID-19, patience and risk-seeking increase willingness to get tested
- Risk aversion, unemployment, and conservative political orientation correlate negatively with willingness to get tested
- Similar with respect **vaccination**?



Fallucchi, ..., Suhrcke (2021) Health Policy in press. doi: 10.1016/j.healthpol.2021.05.003.



- Representative seroprevalence: 7.7 % (15 January 2021; before vaccination)
- Case ascertainment: <u>almost complete</u>
 - Belgium: 37.7 %
 - No other data available



• Case fatality rate: **1.2 %**, 21st March 2021 (1.2 %, 18th April 2021)

Belgium: 2.8 % (2.5 %), France: 2.1 % (1.9 %), Germany: 2.8 % (2.5 %)



Wilmes*, Mossong & Dentzer (2021) The Lancet Regional Health – Europe 5:100116. doi: 10.1016/j.lanepe.2021.100116.









- LST turned out to be the **only feasible strategy** as non-symptomatic carriers are infectious
- Over the last 18 months Luxembourg kept schools and shops open, while in France, Germany and Belgium they were in lock-down



May 2021

What is the prevalence of the different categories of Long COVJD symptoms?

DOES VACCINATION PREVENT LONG COVID?

Which demographic 4socio-economic factors influence the Vaccination willingness and beliefs towards Vaccination?

What are the characteristics and different categories of Long COVID symptoms and their trajectories over time?

What is the long-term effects of COVID on children?

What are the clinical symptoms in case of re-infection/infection post-vaccination of asymptomatic, mild and moderate COVID-19 patients?

What is the impact of re-infection on Long COVID patients?

Which other respiratory or other pathogens are presently or were prevalent in COVID and long-COVID patients?

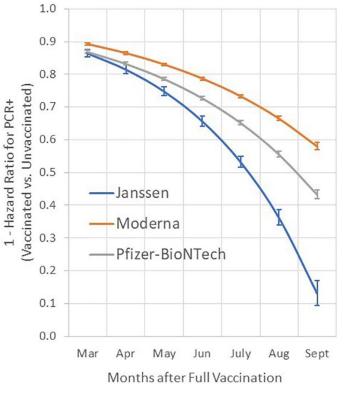
What is the impact of the different variants on Long COVID?

What are the environmental factors contributing to Long COVID?



The COVID-19 Pandemic

- COVID-19 affected **249 millions of individuals worldwide** with 5 million deaths
- 82,842 cases and 849 deaths reported in Luxembourg
- Vaccination campaigns are advancing but herd immunity is difficult to reach
- Variants of concerns represent a threat in particular with the dominance of the Delta variant
- Breakthrough infections are shown to correlate with a decrease in antibody titres, especially at 6 months or more after the second dose
- Waning immunity is a specific concern as recent studies conducted in US and UK have shown that COVID-19 vaccines become less effective after few months although they offer good protection against severe illness and death
- Debate on the use of **booster shots** ongoing: studies have shown that booster shots could flatten the curves of breakthrough cases



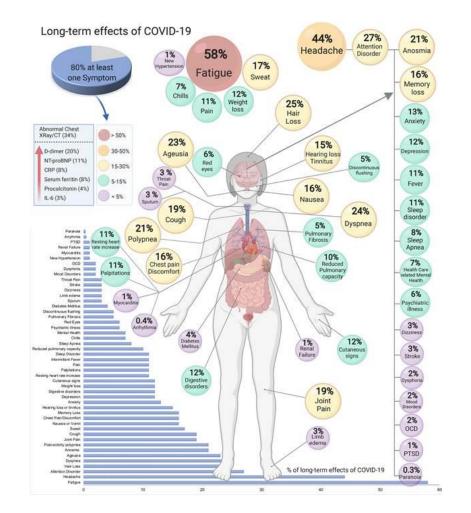
Cohn *et al.* SARS-CoV-2 vaccine protection and deaths among US veterans during 2021. *Science* (2021) <u>DOI:</u> <u>10.1126/science.abm0620</u>





Long COVID

- Many patients who survive the initial infection do not fully recover and experience **a spectrum of symptoms** that persist for several months
- Long COVID symptoms include fatigue, headaches, anxiety, shortness of breath, loss of smell or taste and cognitive impairment and can affect hospitalized and non hospitalized people.
- Long COVID has become an **important public health issue**
- Long COVID could affect 25 to 40% of the COVID-19 patients and may represent around 25,000 persons in Luxembourg
- Luxembourg launched in August 2021 a pilot project for the multidisciplinary management of Long COVID patients; 300 persons are already followed in Long COVID consultations (involvement of the CHL, REHAZENTER and Domaine Thermal Mondorf)







The CoVaLux study

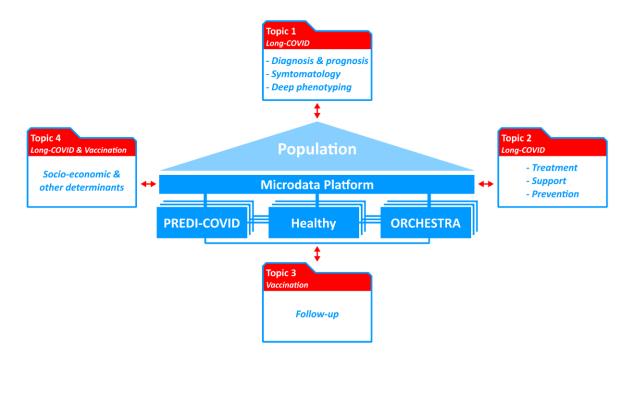
Two main research questions Four complementary topics

Q1

What is the short- to mid-term impact of vaccination in relation to the evolution of immunity and breakthrough infections by variants, and how could this knowledge be used to inform the future vaccination strategy in Luxembourg?

Q2

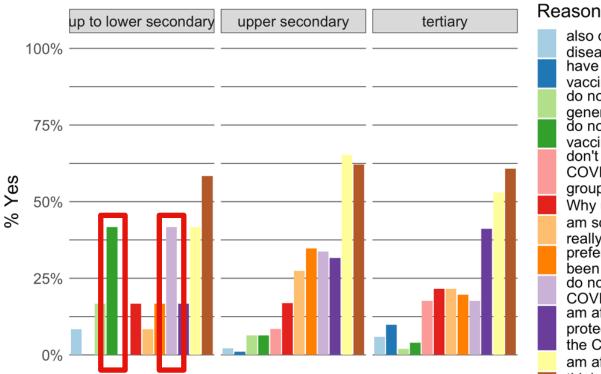
How can we use a better understanding of the symptomatology, prevalence and socioeconomic factors of long COVID for enhanced prevention, diagnosis, prognosis, support and treatment of this public health emergency in Luxembourg?



CHNF







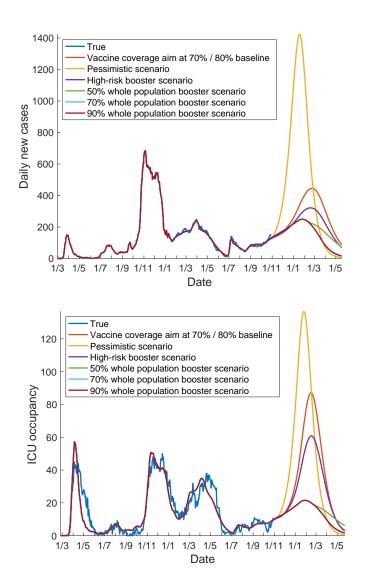
Education

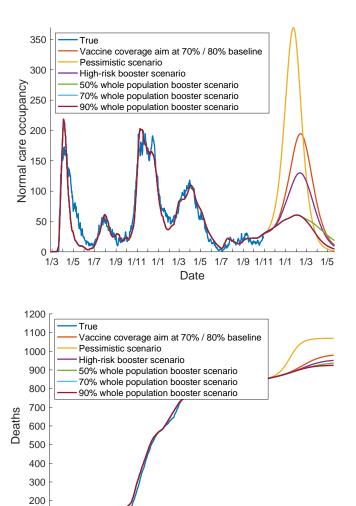
Reasons to be undecided also do not get vaccinated against other diseases have had bad experiences with other vaccinations do not believe in vaccinations in general do not feel well enough informed about vaccinations in general don't think I need a vaccination against COVID-19 because I am not in the risk group Why undecided: Other reasons am sceptical that the COVID-19 vaccine really protects prefer to wait until more people have been vaccinated do not feel well enough informed about COVID-19 vaccinations am afraid that COVID-19 vaccine does not protect against future mutated forms of the Coronavirus am afraid of possible side effects think that the vaccine has not been tested sufficiently



Effects of booster shots

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1/5 1/7 1/9 1/11 1/1 1/3 1/5 1/7 1/9 1/11 1/1 1/3 1/5

Date



- In the COVID-19 pandemic, Luxembourg's academic institutions, Ministries and government managed to plan, implement and execute an entirely novel mitigation strategy with a minimal time delay
- All participating organisations as well as the Luxembourg population established an effective and smooth **routine to deal with the pandemic**
- After initally low attendance, the test campaign turned out to have worked almost flawlessly, producing **high-quality, reliable test results**
- As a consequence, Luxembourg has succeded in **suppressing its second wave**





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• Broad testing and contact tracing: very high case ascertainment

- Strong evidence why Luxembourg's management of the crisis has worked
 - Broad testing and systematic contact tracing: very high case ascertainment
 - Avoidance of multiple lockdowns and severe measures to contain the virus
 - Close to normal life, with almost all services open. Even schools have been functioning in-person
 - In contrast to situation in neighbouring countries





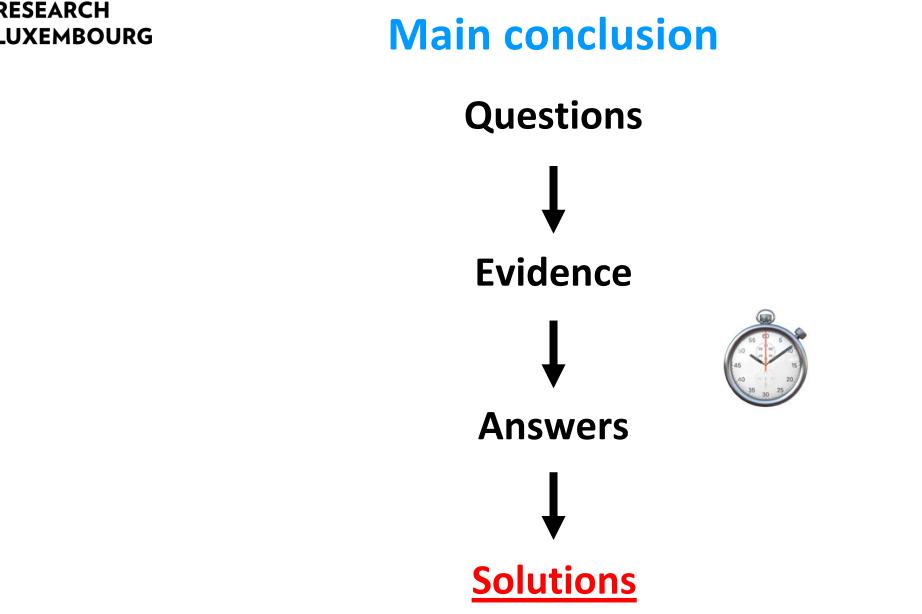
 Luxembourg, by having broadly screened and by continuing to screen its population for SARS-CoV-2, represents an ideal study population for studying longer-term effects of COVID-19 (asymptomatics)

 Comparable results obtained in other countries and regions (LFT), although our positive predictive values were much higher (rRT-PCR)

 Impact of vaccination and longer-term consequences of COVID: essential research questions











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Thank you very much for your attention!



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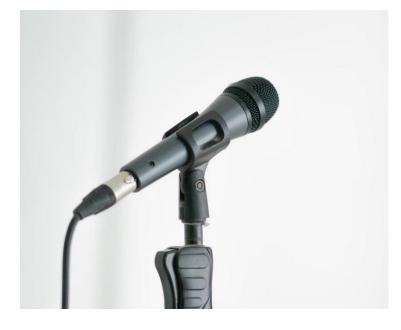






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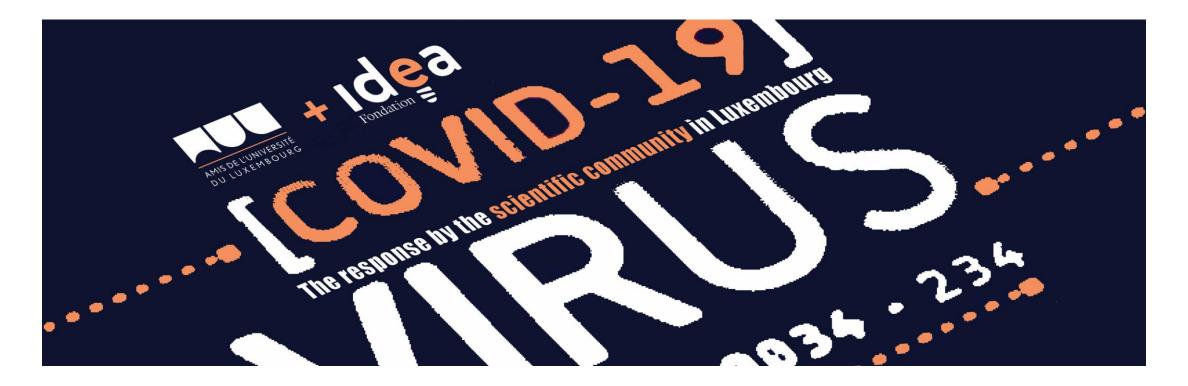
The COVID-19 pandemic and the response by the scientific community in Luxembourg



Q&A session







The COVID-19 pandemic and the response by the scientific community in Luxembourg





Chambre de Commerce, Luxembourg 17 November 2021













UXEMBOURG

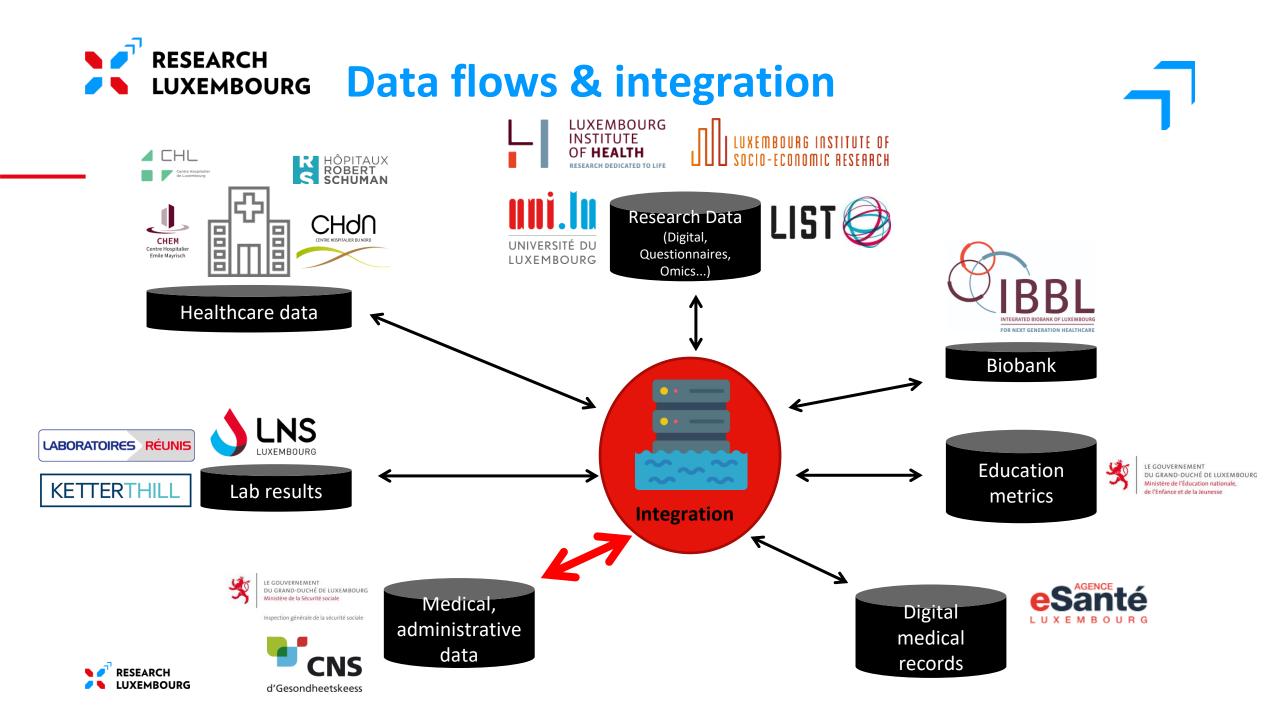
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VS

VS











Luxembourg: 58,955 cases based on 2,280,826 tests (21 March 2021)
64,746 cases based on 2,570,608 tests (18 April 2021)

• Overall positivity rate: **2.6 %** (2.5 %)

Belgium: 8.1 % (7.9 %), France: 7.4 % (7.4 %), Germany: 5.6 % (5.8 %)
Norway: 2.0 % (2.2 %), The Netherlands: 17.0 % (16.6 %)



Wilmes*, Mossong & Dentzer (2021) The Lancet Regional Health – Europe 5:100116. doi: 10.1016/j.lanepe.2021.100116.



Diagnostic Methods for COVID-19 in Luxembourg RT-PCR for SARS-CoV-2 – Quality Aspects LST

	FastTrack, FTD SARS-CoV-2 (RUO)							
Std dilution	-	1:10	1:20	1:100	1:200	1:1000		
copies/well (10ul)	2000	200	100	20	10	2		
repli 1	27.19	30.51	31.36	33.87	34.85	36.4		
repli 2	27.09	30.59	31.37	33.45	34.6	38.17		
repli 3	27.22	30.36	31.31	33.52	34.82	36.65		
repli 4	27.12	30.44	31.28	33.46	35.16	37.45		
repli 5	27.16	30.36	31.35	33.63	34.42	37.87		
repli 6	27.13	30.39	31.39	33.56	35.03	36.35		
repli 7	27.16	30.34	31.48	33.45	34.47	37.46		
repli 8	27.26	30.34	31.24	33.54	35.03	37.35		
repli 9	27.19	30.54	31.45	33.84	34.63	36.18		
repli 10	27.13	30.41	31.3	34.24	34.65	38.05		
mean	27.17	30.43	31.35	33.66	34.77	37.19		
stdev	0.048	0.085	0.071	0.242	0.239	0.705		

	Generic E gene (Charité, Berlin)							
Std dilution	-	1:10	1:20	1:100	1:200	1:1000		
copies/well (5ul)	1000	100	50	10	5	1		
repli 1	28.43	31.51	32.46	35.97	36.04	45		
repli 2	28.33	32.21	32.73	35.68	37.22	45		
repli 3	28.35	31.42	32.81	35.05	37.03	37.76		
repli 4	28.44	31.48	32.64	35.16	35.1	45		
repli 5	28.39	31.59	32.21	36.69	35.62	36.64		
repli 6	28.16	31.43	32.62	33.95	36.4	45		
repli 7	28.26	31.66	32.83	35.02	37.63	37.47		
repli 8	28.16	31.26	32.48	36.01	35.12	45		
repli 9	28.2	32.02	32.61	35.06	36.93	45		
repli 10	28.31	31.37	32.54	34.97	35.17	45		
mean	28.30	31.60	32.59	35.36	36.23	42.69		
stdev	0.099	0.284	0.175	0.717	0.902	3.543		

	Generic N gene (CDC)						
Std dilution	-	1:10	1:20	1:100	1:200	1:1000	
copies/well (5ul)	1000	100	50	10	5	1	
repli 1	27.04	30.13	31.37	34.35	34.43	35.71	
repli 2	27.04	30.34	31.24	33.97	34	45	
repli 3	26.96	30.07	31.25	33.54	35.24	45	
repli 4	26.82	30.29	30.8	33.36	34.14	36.8	
repli 5	27.05	30.15	31.06	34.19	35.52	45	
repli 6	26.91	30.1	31.17	33.14	35.82	36.08	
repli 7	26.78	29.83	30.72	33.48	33.67	37.83	
repli 8	27.03	30.14	31.3	33.06	36.34	37.65	
repli 9	26.89	30.15	30.92	33.38	33.99	36.64	
repli 10	27.01	29.89	31.2	33.15	34.16	36.7	
mean	26.95	30.11	31.10	33.56	34.73	39.24	
stdev	0.093	0.148	0.209	0.432	0.874	3.817	

imit of detection (LoD) of the generic N gene assay is 5 RNA copies/reaction.

Limit of detection (LoD) of the FTD assay is below 2 RNA copies/reaction

Through its set-up and design as a single well and dual-target assay, the FTD SARS-CoV-2 (CE-IVD; FDA-EUA) assay was found to be ideally suited for population-based and large-scale screening due to its high sensitivity (<u>detection of as low as 2 RNA copies per reaction</u>) paired with no compromise in specificity and technical reliability.





Diagnostic Methods for COVID-19 in Luxembourg Validation of 1+3 Pooling Strategy

Sensitivity

- 30 pools tested with FTD assay:
 - Ct value <30 (n=6)
 - Ct value between 30 and 35 (n=11)
 - Ct value >35 (n=13)
- All pools correctly identified as positive, only slight shift in of Cq values between pool and single sample ∠ 100% sensitivity

Specificity

• 20 pools of negative samples compared with deconvoluted samples ∠ specificity reached 100%

