

Respiratory Viruses in Luxembourg (ReViLux)

Weekly report - 16 May 2023

Executive summary

The sentinel network detected a low epidemic activity over the last week (2023/19), based on 2.8% of consultations being associated with influenza-like illness. Among the specimen collected by the sentinel network over the last two weeks, the percentage of positive tests for Influenza type A was 3.4% and 1.7% for SARS-CoV-2 and Influenza type B. Overall, during this season, Influenza A (H3) with 88.3% is the most frequent detected strain in submitted specimens. However, between weeks 2023/05 and 2023/16 Influenza type B was predominant.

Concerning SARS-CoV-2 genomic surveillance, the estimated distribution of variants was 35.5 % (95%CI: 26.5- 44.4%) for XBB.1.9 and XBB.1.5. The estimates of other recombinant XBB (excluding XBB.1.5 and XBB.1.9) increased from 19.7 to 27.3% (95%CI: 19.0-35.6%) compared to previous week. Since week 2023/06, XBB (including XBB.1.9 and XBB.1.5) has been dominant.

IN THE SPOTLIGHT

XBB.1.5 sub-lineage

XBB.1.5 is a subtype with additional spike RBD mutation F486P. The lineage was first detected in October 2022 in the USA and is estimated to have a large growth advantage to other previously circulating lineages. Currently, there are no signals that the infection severity of XBB.1.5 is different from that of other circulating Omicron sub-lineages. ECDC categorizes lineage that have similar Spike proteins as **VOI XBB.1.5 lineages** (XBB.1.5 like), which include XBB.1.5, XBB.1.9.1, XBB.1.9.2 and XBB.1.16.

XBB.1.16 has also a similar genetic profile as XBB.1.5, but two additional mutations E180V and K478R in the Spike protein. The sub-variant has been classified as variant of interest and is closely monitored (WHO, 2023).

Introduction

The Laboratoire national de santé, as **National Reference Laboratory for Acute Respiratory Infections** in Luxembourg, performs close surveillance on respiratory viruses, with a special focus on SARS-CoV-2. The main monitoring activities are:

- The Sentinel Surveillance. It provides a broad picture of respiratory diseases affecting the Luxembourgish population, based on clinical and laboratory data.
- The SARS-CoV-2 Genomic Surveillance. It enables detailed observation of SARS-CoV-2 variants and mutations through time and space, as well as studying specific groups of interest.

The objective of the ReViLux report is to inform public health actions in Luxembourg.

Sentinel Surveillance Network

The Sentinel Surveillance aims at monitoring the circulating respiratory viruses, from traditional ones like Influenza to more recent ones like SARS-CoV-2, and hence underpin public health actions. The Sentinel Network is a group of general practitioners and paediatricians spread over the country. They report the weekly number of patients showing symptoms suggestive of acute respiratory infection (ARI) and influenza-like illness (ILI), and those patients are then sampled and tested for a panel of respiratory viruses. The circulation of respiratory viruses in the north hemisphere is generally monitored by seasons that go from week 40 to week 20. The period between weeks 20 and 40 is usually called interseason.

Clinical results

Last week, 2.8% of consultations were reported as ILI, which represents a low epidemic activity for Luxembourg, according to ECDC and the Moving Epidemic Method. The history of ILI consultations is displayed in Figure 1, and a detailed summary of the number of ARI and ILI cases during the last four weeks is included in Table 1.

Laboratory results

Over the last two weeks, the most frequently detected viruses (according to positivity rates) were Parainfluenzavirus (44.4%), Human rhinovirus (33.3%) and Influenzavirus A (3.4%). An overview of the viral circulation during the current season is displayed in Figure 2 and Table 2.

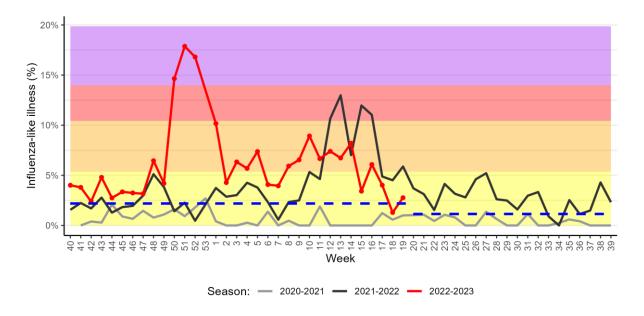


Figure 1. Percentage of patients with influenza-like illness over the last three seasons Background colours according to intensity of circulation: low, medium, high, very high.

Table 1. Syndromic surveillance over the last 4 weeks

Week -	ARI		ILI		Total	
	N	%	N	%	consultations	
2023/16	41	11.33	22	6.08	362	
2023/17	25	12.56	8	4.02	199	
2023/18	24	15.58	2	1.30	154	
2023/19	40	13.84	8	2.77	289	

ARI: Acute Respiratory Infections; ILI: Influenza-Like Illness.

Figure 2. Distribution of respiratory viruses detected within the Sentinel Network, by two-week periods. Results from last weeks are not yet consolidated.

Week (season 2022/2023)

FLU-A: influenzavirus A; FLU-B: influenzavirus B; PIV: parapneumovirus; RSV: respiratory syncytial virus; ADV: adenovirus; MPV: metapneumovirus; HRV: human rhinovirus; SC2: SARS-CoV-2.

Table 2. Distribution of respiratory viruses detected within the Sentinel Network over the last 2 weeks compared to the whole current season.

Vince		Last 2 weeks	Season 2022/2023		
Virus	N*	Positivity rate (%)	N*	Positivity rate (%)	
Parainfluenzavirus	4	44.4	56	2.3	
Human rhinovirus	3	33.3	462	19.1	
Influenzavirus A	2	3.4	409	15.6	
SARS-CoV-2	1	1.7	324	21.9	
Influenzavirus B	1	1.7	311	11.8	
Adenovirus	0	0.0	220	9.1	
Respiratory syncytial virus	0	0.0	209	8.6	
Metapneumovirus	0	0.0	145	6.0	
Total	11		2136		

^{*}Co-infection cases counted once for each virus detected.

Genotyping analyses

In order to assess the circulation of viral strains, genotyping analyses were also performed for Influenzavirus A and Respiratory syncytial virus (RSV) (Figure 3). During the current season, the most frequently detected strain for Influenzavirus A was H3 (88.3%), and for RSV it was type B (74.3%).

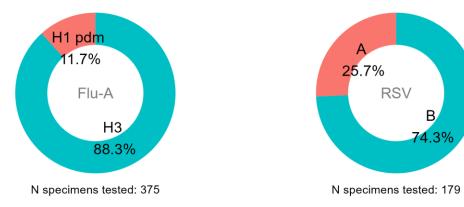


Figure 3. Detection of influenzavirus-A (Flu-A) and Respiratory syncytial virus (RSV) strains in sentinel samples during the current season.

SARS-CoV-2 Genomic Surveillance

On 05.05.2023, the World Health Organisation has lifted the Public Health Emergency of International Concern (PHEIC) for COVID-19. It is now an established and ongoing health issue, therefore testing and sequencing strategy have been reviewed in Luxembourg. The National Reference Laboratory for Acute Respiratory Infections at LNS continues to monitor the genomic characteristics of the SARS-CoV-2 virus detected in Luxembourg.

LNS receives positive specimens (nasopharyngeal or oropharyngeal swabs analysed by RT-PCR) from the national network of laboratories. All hospital specimens are sequenced, as well as a representative selection of community specimens. Illumina and Nanopore devices are used respectively. When needed, an additional screening by targeted PCR is also performed to enable earlier evaluation of the spread of newly emerging variants.

Bioinformatic analyses are based on a standardised pipeline, and lineage assignment is performed through the PANGOLin software (4.2, pango-data 1.19, mode UShER). In order to allow easier visualisation, WHO and Nextstrain categorisations are also used.

Sequencing activity

In week 2023/18, the microbial genomics unit at LNS sequenced 122 specimens from the week of study, including 111 national samples. We extrapolated COVID cases and estimated sequencing coverage of 48.7% (see Figure 4). This probably does not reach the recommended sample size to detect emerging variants at a 5% incidence.

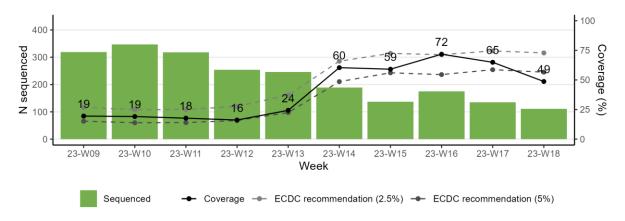


Figure 4. Number of specimens sequenced and national coverage based on weekly number of positive cases in Luxembourg. The coverage from the last two weeks is not consolidated yet.

Variant circulation

The 'XBB.1.5 like' variants remain the main variants of interest within the selected sample, the most frequent sub-variants being XBB.1.9 as well as XBB.1.5. The proportion of XBB.1.5 dropped from 36.1% in week 2023/17 to 35.5% in week 2023/18, and XBB.1.9 decreased from 37.7 to 35.5%, respectively. The estimates of other recombinant XBB (excluding XBB.1.5 and XBB.1.9) increased from 19.7 to 27.3% (95%CI: 19.0-35.6%) compared to previous week. Since week 2023/06, XBB (including XBB.1.9 and XBB.1.5) has been dominant.

An overview of the variants and lineages circulating over the last weeks is displayed in Figure 5, and further details are shown in Table 3. The history of the circulation of each variant since January 2021 is displayed in Figure 6.

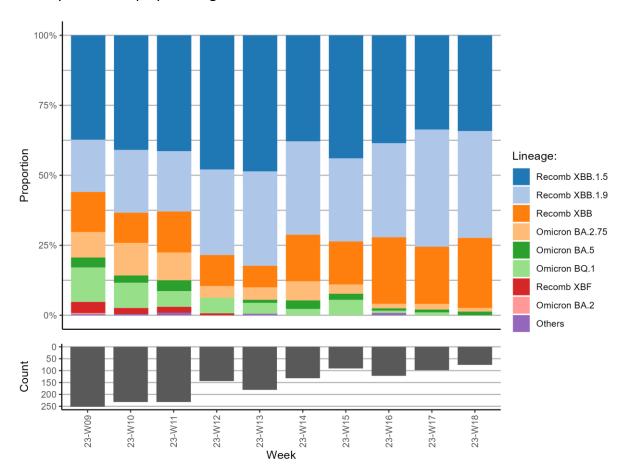


Figure 5. Distribution of lineages within the representative sample during the last 10 weeks.

Table 3. Distribution of SARS-CoV-2 lineages detected within the representative sample during the last two weeks. Previously reported cases might be updated by retrospective analysis.

Lincomo*	V	Veek 17	Week 18		
Lineage*	%	CI %	%	CI %	
Recomb XBB.1.9	37.7	29.1 - 46.3	35.5	26.5 - 44.4	
Recomb XBB.1.5	36.1	27.5 - 44.6	35.5	26.5 - 44.4	
Recomb XBB	19.7	12.6 - 26.7	27.3	19.0 - 35.6	
Omicron BA.2.75	2.5	0.0 - 5.2	0.9	0.0 - 2.7	
Omicron BA.5	2.5	0.0 - 5.2	0.9	0.0 - 2.7	
Omicron BQ.1	1.6	0.0 - 3.9	0.0	-	

^{*}Sub-lineages are not included in the count of their parental lineages (e.g. BQ.1 not included in BA.5). CI: confidence interval at 95%.

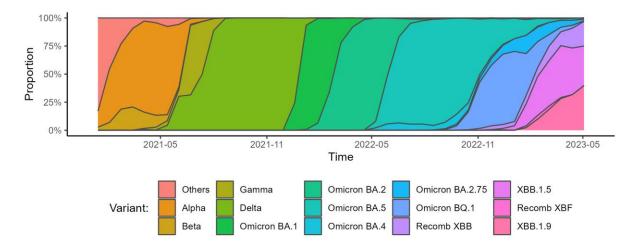


Figure 6. Proportion of each variant circulating in Luxembourg since January 2021.

Clinical and epidemiological factors

In this section, the lineage distribution of all specimens sequenced over the last month is assessed by demographics (sex and age group, Figure 7) and sampling setting (community vs. hospital, Table 4).

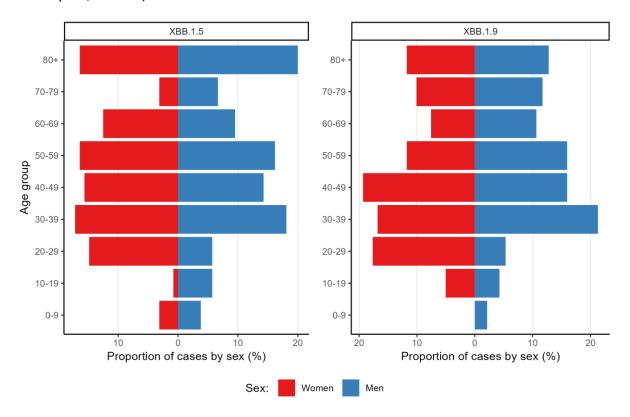


Figure 7. Age group and sex distribution of specimens sequenced over the last 4 weeks, by lineage.

Table 4. Comparison of lineage distribution by sampling setting.

Lineage	Community					Hospital
	Women	Men	Total	Women	Men	Total
Recomb XBB.1.5	55.2%	50.7%	53.3%	42.2%	56.9%	49.6%
Recomb XBB.1.9	44.8%	49.3%	46.7%	57.8%	43.1%	50.4%
Total	100.0%	100.0%	100.0%	100.0%	100.0%	100.0%

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