

Implementing WGS data collection and analysis for communicable diseases at EU/EEA level

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Campylobacter activities at ECDC

- Annual case-based data collection and \bullet reporting at the EU/EEA-level
- Annual AMR isolate-based data collection and • reporting at the EU/EEA-level
- Food- and waterborne diseases network (FWD-Net) meeting 2020, emphasis on *Campylobacter*
- Joint EURL-AR and FWD-Net workshop on Salmonella and Campylobacter AMR, also • 2020
- WGS support for possible multi-country events through laboratory contractor \bullet
- WGS analysis tools for outbreak investigations







Figure 2. Distribution of confirmed campylobacteriosis cases by month, EU/EEA, 2013–2017



Source: Country reports from Austria, Cyprus, the Czech Republic, Denmark, Estonia, Finland, France, Germany, Hungary, Iceland, Ireland, Italy, Latvia, Lithuania, Luxembourg, Malta, the Netherlands, Norway, Poland, Romania, Slovakia, Slovenia, Spain, Sweden and the United Kinadom.

Indicator-based and event-based surveillance: **TESSy and EPIS**



Notified cases

Whole genome sequencing (WGS) at ECDC



ECDC strategic framework for WGS by 2021:

- Cross-border outbreak investigations
- Control-oriented surveillance (cross-border outbreak detection)
- Strategy-oriented surveillance

Extensive list of diseases and pathogens to be implemented

- Food- and waterborne diseases
- Vaccine-preventable diseases
- Multidrug-resistant tuberculosis
- Antimicrobial resistance
- Influenza
- Any emerging infectious disease threat at the EU level



ECDC WGS system upgrade, design principles



- Re-use existing tools and resources for storage, calculation and visualisation
- Integrate data from several different calculation systems in the same Data Warehouse data model
- Encourage open data sharing while also fulfilling all data protection and access control requirements
- Focus on the user experience and service-oriented architecture (data submission and data exploration)



Data flow schematic





Building blocks



Built by ECDC:

- Data upload interface (API and GUI), TESSy
- Data warehouse for derived data, new
- Web service API for data consumption, new
- Cluster detection scripts, existing
- Web page to hold visualisation components, new

Re-used:

- Storage (currently ENA and network drive, cloud storage to be implemented in 2020)
- Calculation systems and operational databases (currently Bionumerics and BIGSdb, more will be added as needed)
- Interactive visualisation (currently MicroReact, we are also looking at NextStrain)
- Charts and tables libraries (charts.js, datatables.js)





ECDC upgraded system expected to be operational Q1 2020

Currently focus on *Listeria*, in 2020 we expect to have support for continous surveillance for *Listeria*, *Salmonella*, STEC, *Neisseria meningitidis*, Influenza and MDR-TB

Outbreak support for *Campylobacter*

ECDC-EFSA joint database implementation will start Q1 2020



ECDC tool for analysing WGS data for clusters



Visualisation Molecular clusters/datasets My data Settings

These table shows your uploaded data, you can choose to share or not share individual isolates with other users.

Save settings Reload data from source (slow, you must refresh your browser to see the changes)

Column visibility Copy CSV Show 100 T entries

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					LISTISO subr	missions				
Countr y	BatchId 🗘	RecordId	Created date 🔻	Modified date	QC (core loci detect ed)	QC (PASS/FAIL)	DateUsedForStatisti cs	Cluster code	Event 🔶	Share assembly
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			2019-09-19 14:52:02	2019-09-25 08:44:34	97.76	PASS	2019-06-18			
			2019-09-19 14:52:01	2019-09-25 08:44:31	98.56	PASS	2019-05-13			۲
			2019-09-19 14:52:01	2019-09-25 08:44:30	98.45	PASS	2019-05-26	2019-04.LIST.03.CC570		۲
			2019-09-18 18:12:43	2019-09-25 08:44:32	100.0	PASS	2019-08-08			
			2019-09-18 18:12:43	2019-09-25 08:44:28	100.0	PASS	2019-08-28	2017-10.LIST.82.CC1.Asc 1.0053.Aps1.0044		
			2019-09-18 17:02:27	2019-09-20 15:13:34	99.94	PASS			UI-490 Cluster B ST32	
			2019-09-18 17:02:27	2019-09-20 15:13:39	99.88	PASS			UI-490 Cluster B ST32	
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			2019-09-18 12:19:36	2019-09-25 08:44:28	99.82	PASS	2019-06-15	2018-10.LIST.18.ST32	UI-490 Cluster B ST32	
			2019-09-06 10:53:35	2019-09-20 10:16:49	99.88	PASS		2019-09.LIST.01.CC388	<u>UI594</u>	
			2019-08-27 16:23:42	2019-08-28 14:30:00	98.51	PASS		2019-05.LIST.06		
			2019-08-27 16:23:42	2019-09-25 09:23:31	97.99	PASS	2019-04-17	2017-10.LIST.01.CC87		
			2019-08-27 16:23:42	2019-09-25 09:23:41	97.94	PASS	2019-04-19	2017-10.LIST.01.CC87		

Search:



Prototype version 0.17 Powered by <u>MicroReact</u>

LISTISO EBOV IMD ZIKV

Visualisation Molecular clusters/datasets My data Settings

These tables shows molecular clusters detected by ECDC by applying automated algorithms, see respective surveillance protocols for details. Only multi-country clusters and clusters in your country are shown. Included are also generated datasets of particular interest.

Cluster ID	Event ID 🔶	Cluster status 🛛 🔻	Number of isolate s	Countries 🔶	Updated (7d)	First date 🕴	Last date 🔻 🔻	Last updated 🕴	Extended cluster (7AD)	Neighborhood (50 AD)	
Search Cluster ID	Search Event ID	Search Cluster stat	Search Number of	Search Countries	Search Updated (7	Search First date	Search Last date	Search Last update	Search Extended c	Search Neighborhc	
2017-10.LIST.82.CC1.A scI.0053.ApaI.0044		OPEN	5			2012-07-15	2019-08-28	2019-09-25	2017-10.LIST.82.CC1. AscI.0053.ApaI.0044	2017-10.LIST.82.CC1. AscI.0053.ApaI.0044	
2018-10.LIST.18.ST32	UI-490 Cluster B ST32	OPEN	8			2018-01-27	2019-07-18	2019-09-25	2018-10.LIST.18.ST32	2018-10.LIST.18.ST32	
2019-03.LIST.02.CC1		OPEN	8			2018-02-25	2019-06-12	2019-08-06	2019-03.LIST.02.CC1	2019-03.LIST.02.CC1	
2017-10.LIST.01.CC87		OPEN	37			2011-12-13	2019-06-05	2019-09-25	2017-10.LIST.01.CC87	2017-10.LIST.01.CC87	
2019-06.LIST.10.CC8	<u>UI563</u>	OPEN	5			2019-05-15	2019-05-15	2019-07-09	2019-06.LIST.10.CC8	2019-06.LIST.10.CC8	
2017-10.LIST.69.CC1		OPEN	3			2010-08-27	2019-02-19	2019-04-30	2017-10.LIST.69.CC1	2017-10.LIST.69.CC1	
2016-11.LIST.01	<u>UI452</u>	OPEN	25			2014-07-23	2019-02-18	2019-05-29	2016-11.LIST.01	2016-11.LIST.01	
2017-06.LIST.70.CC1.A scI.0053.ApaI.0040		OPEN	48			2015-08-26	2019-02-18	2019-07-03	2017-06.LIST.70.CC1. AscI.0053.ApsI.0040	2017-06.LIST.70.CC1. AscI.0053.ApaI.0040	
2019-07.LIST.14.CC87		OPEN	2			2018-08-15	2018-11-30	2019-08-07	2019-07.LIST.14.CC87	2019-07.LIST.14.CC87	
2019-07.LIST.12.CC2		OPEN	2			2014-09-26	2018-11-26	2019-08-07	2019-07.LIST.12.CC2	2019-07.LIST.12.CC2	
2019-07.LIST.13.CC6		OPEN	2			2013-10-15	2018-11-21	2019-08-07	2019-07.LIST.13.CC6	2019-07.LIST.13.CC6	
2018-05.LIST.07.CC15 5		OPEN	7			2015-08-30	2018-10-18	2019-08-07	2018-05.LIST.07.CC15 5	2018-05.LIST.07.CC15 5	
2015-04.LIST.04.CC6.A scI.0002.ApsI.0072	UI583_test	OPEN	26			2010-01-27	2015-03-26	2019-09-10	2015-04.LIST.04.CC6. AscI.0002.ApsI.0072	2015-04.LIST.04.CC6. AscI.0002.ApsI.0072	
2019-06.LIST.11.CC6	<u>UI552</u>	OPEN	3					2019-07-09	2019-06.LIST.11.CC6	2019-06.LIST.11.CC6	
2019-08.LIST.01.CC9		CLOSED	2			2018-01-17	2019-06-04	2019-08-06	2019-08.LIST.01.CC9	2019-08.LIST.01.CC9	
2019-04.LIST.03.CC57 0		CLOSED	3			2019-05-26	2019-05-26	2019-09-25	2019-04.LIST.03.CC57	2019-04.LIST.03.CC57 0	
2017-11.LIST.11.CC87		CLOSED	7			2015-05-30	2019-05-01	2019-07-10	2017-11.LIST.11.CC87	2017-11.LIST.11.CC87	
2019-03.LIST.07.CC6		CLOSED	3			2017-03-24	2019-02-20	2019-07-08	2019-03.LIST.07.CC6	2019-03.LIST.07.CC6	
2019-07.LIST.01.CC2		CLOSED	2			2016-05-24	2019-02-10	2019-07-10	2019-07.LIST.01.CC2	2019-07.LIST.01.CC2	



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Visualisation Molecular clusters/datasets My data Settings



MicroReact Tables and figures Distance matrix Isolate table and download





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LISTISO EBOV IMD ZIKV

Visualisation Molecular clusters/datasets My data Settings

Pathogen: MicroReact Tables and figures Distance matrix Isolate table and download Listeria T Clustering method: Single linkage Include data from SRA and ENA 5 Include data from all countries Sampled within X days: 4 Uploaded within X days: Search for specific cluster/dataset: 3 2017-10.LIST.01.CC87 Search for specific UI: 2 Search for specific RecordIds: 1 After filtering, also include all matches within X genetic differences: 0 7 2011-12 Update

URL for sharing: http://microreact.ecdcnet.e URL for recreating: http://zbiodev1.idmdevdmz



Age and gender distribution







Software options for WGS data upload



- ECDC WGS upload application
- 🛸 ECDC WGS upload app 17 Data Setup Submission View Isolate table Total entries: 3 Selected: Submit data to configured syst Modified date Assembly file Selected for TE .. TESSy batch TESSy validation ENA run MISEQ_2X300 MISEQ_2X300 2019-01-30 10:2. SUBM-UPDATE-F. SUBM-UPDATE-F. TESSy - Create batch from selected and test 019-01-30 10:2.. TESSy - Upload batch... TESSy - Approve batch. TESSy - Reject batch... SFTP - Upload raw data for isolates in TESSy
- ECDC Bionumerics client plugin

• Direct TESSy submission

All three options include mandatory submission to TESSy isolate-based subjects

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Data	baldbase entries												
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•	Key	Level	Modified date	MLST PubMLST A	ReportingCountry	DateUsedForStatistics	DateOfSampling	DateUploaded	DateReleased	SampleOrigin	InsdcR 👻		
<	SRR5193935		2019-01-30 00:00:09		UK	2015-07-01	2015-07-01		2017-01-24	Homo sapiens			
1	SRR5193925		2019-01-29 20:06:00		UK	2016-11-01	2016-11-01		2017-01-24	Homo sapiens			
1	SRR5193934		2019-01-29 21:25:04		UK	2016-09-01	2016-09-01		2017-01-24	Homo sapiens			
1	SRR5193926		2019-01-30 03:16:06		UK	2016-12-01	2016-12-01		2017-01-24	Homo sapiens			
1	SRR5193931		2019-01-30 00:05:38		UK	2016-10-01	2016-10-01		2017-01-24	Homo sapiens			
1	SRR5193929		2019-01-29 21:41:17		UK	2016-08-01	2016-08-01		2017-01-24	Homo sapiens			
1	SRR5193927		2019-01-29 18:55:37		UK	2016-08-01	2016-08-01		2017-01-24	Homo sapiens			
1	SRR5193922		2019-01-29 17:50:59		UK	2016-04-01	2016-04-01		2017-01-24	Homo sapiens			
1	SRR5193923		2019-01-30 02:31:41		UK	2016-06-01	2016-06-01		2017-01-24	Homo sapiens			
1	SRR5193920		2019-01-30 09:21:15		UK	2016-09-01	2016-09-01		2017-01-24	Homo sapiens			
✓	SRR5193932		2019-01-30 01:30:06		UK	2016-09-01	2016-09-01		2017-01-24	Homo sapiens			
1	SRR5193947		2019-01-30 05:04:05		UK	2016-06-01	2016-06-01		2017-01-24	Homo sapiens			
1	SRR5193946		2019-01-30 09:21:06		UK	2016-12-01	2016-12-01		2017-01-24	Homo sapiens			
1	SRR5193945		2019-01-30 09:21:12		UK	2016-05-01	2016-05-01		2017-01-24	Homo sapiens			
~	SRR5193943		2019-01-30 09:21:15		UK	2016-11-01	2016-11-01		2017-01-24	Homo sapiens			



File Edit Database Analysis Scripts ECDC WGS WGS tools Window Held

ECDC WGS upload application



How to use

- Download and install the application
- Read the manual and configure the data upload for each pathogen, personal support is available through the ECDC FWD mailbox
- Configure data import/mapping from Excel/csv, your LIMS system, or enter data directly
- Start uploading in a few clicks according to the reporting protocol
- The epidemiological data are submitted to TESSy only, the WGS reads/assembly is submitted to the selected system(s)



Features

- Can be configured to import data from databases or local files (MySQL, SQL Server, SQLite, Excel, csv)
- Configure only once, single click upload
- Can upload assemblies to TESSy and SFTP, raw reads to SFTP and ENA (configurable)
- Data sharing through SFTP and ENA

