



Nový korona virus 2019-nCoV a klinická charakteristika onemocnění COVID-19

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Obsah

- Charakteristika a morfologie viru 2019-nCoV (SARS-CoV-2)

BRIEF REPORT

A Novel Coronavirus from Patients with Pneumonia in China, 2019

Na Zhu, Ph.D., Dingyu Zhang, M.D., Wenling Wang, Ph.D., Xingwang Li, M.D., Bo Yang, M.S., Jingdong Song, Ph.D., Xiang Zhao, Ph.D., Baoying Huang, Ph.D., Weifeng Shi, Ph.D., Roujian Lu, M.D., Peihua Niu, Ph.D., Faxian Zhan, Ph.D., Xuejun Ma, Ph.D., Dayan Wang, Ph.D., Wenbo Xu, M.D., Guizhen Wu, M.D., George F. Gao, D.Phil., and Wenjie Tan, M.D., Ph.D., for the China Novel Coronavirus Investigating and Research Team

- Klinická charakteristika onemocnění COVID-19

ORIGINAL ARTICLE

Clinical Characteristics of Coronavirus Disease 2019 in China

W. Guan, Z. Ni, Yu Hu, W. Liang, C. Ou, J. He, L. Liu, H. Shan, C. Lei, D.S.C. Hui, B. Du, L. Li, G. Zeng, K.-Y. Yuen, R. Chen, C. Tang, T. Wang, P. Chen, J. Xiang, S. Li, Jin-lin Wang, Z. Liang, Y. Peng, L. Wei, Y. Liu, Ya-hua Hu, P. Peng, Jian-ming Wang, J. Liu, Z. Chen, G. Li, Z. Zheng, S. Qiu, J. Luo, C. Ye, S. Zhu, and N. Zhong, for the China Medical Treatment Expert Group for Covid-19*

- Aktuální situace z 12.03.2020

1) Charakteristika a morfologie 2019-nCoV

- Betacoronavirus (sarbecovirus)
 - obalené RNA viry
 - SARS-CoV (2002-2003), MERS-CoV (2012)
 - genetická diversita a častá rekombinace

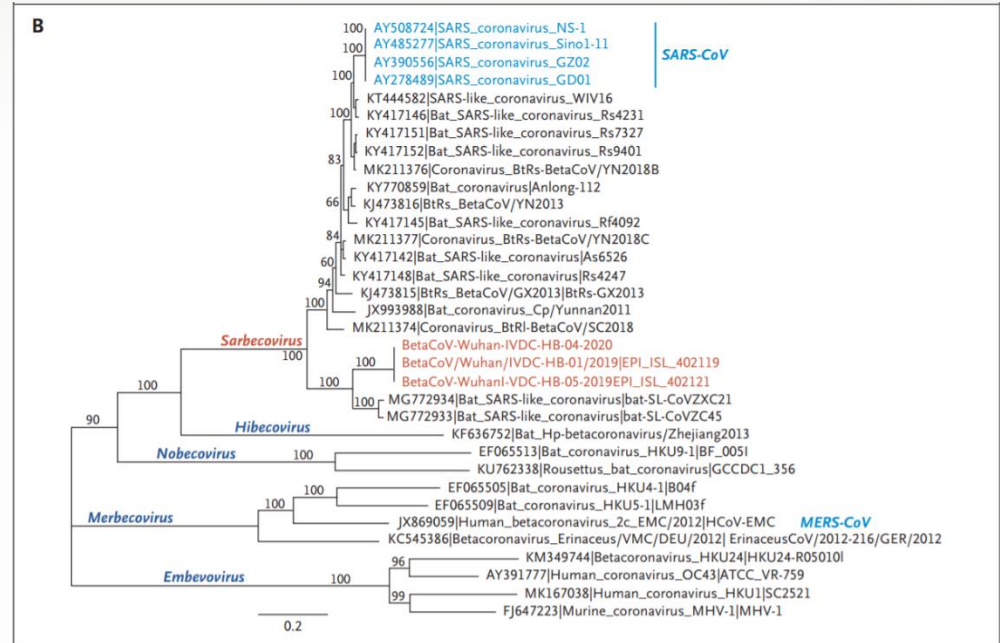


Figure 4. Schematic of 2019-nCoV and Phylogenetic Analysis of 2019-nCoV and Other Betacoronavirus Genomes.

Shown are a schematic of 2019-nCoV (Panel A) and full-length phylogenetic analysis of 2019-nCoV and other betacoronavirus genomes in the Orthocoronavirinae subfamily (Panel B).

Vzorky a izolace viru

- 4x BAL– pneumonie nejasné příčiny – Wuhan 21.12.2019
- 7x kontrolních BAL pacientů s pneumonií známe etiologie – Peking
- Virus transport medium – centrifugace – inokulace supernatantu na epitel.bb DC – denně světél. mikroskopie + RT-PCR
- Epitelie s průkazem cytopatického efektu na povrchu bb s úbykem cílii 72 hod po inokulaci
- Po 6 dnech elektronová mikroskopie

Pacienti s nejasnou pneumonií

- Primární výskyt 12/2019 Wuhan, Hubei Province
 - epidemiologické spojení s trhem mořských plodů a zvířat
- PACIENT č.1 – 49 let, Ž, bez chronické medikace, horečka, kašel, dyskomfort na hrudníku s progresí, CT nález, obchodník Wuhan, vyléčena
 - kašel, horečka, po 7 dnech dušnost s RTG progresí, ventilace, smrt po 20 dnech, pravidelný návštěvník trhu
- PACIENT č.2 – 61 let, M,
 - PACIENT č. 3 – 32let, M, vyléčen

Morfologie a kontagiozita SARS-CoV-2

- Sférické partikule s polymorfizmy, \varnothing 60-140 nm, výběžky (9-12nm) – „solar corona“
 - Extracelulární volný virus
 - Intracelulárně inkluze naplněné virovými partikuly s vazbou na cytoplazmat. membránu
- R_0 (attack rate)
 - počet nakažených osob od 1 nemocné osoby
 - WHO: 1.4 – 2.5
 - Studie 1.5 - 3.5
 - Chřipka 1.3
 - SARS 2.0

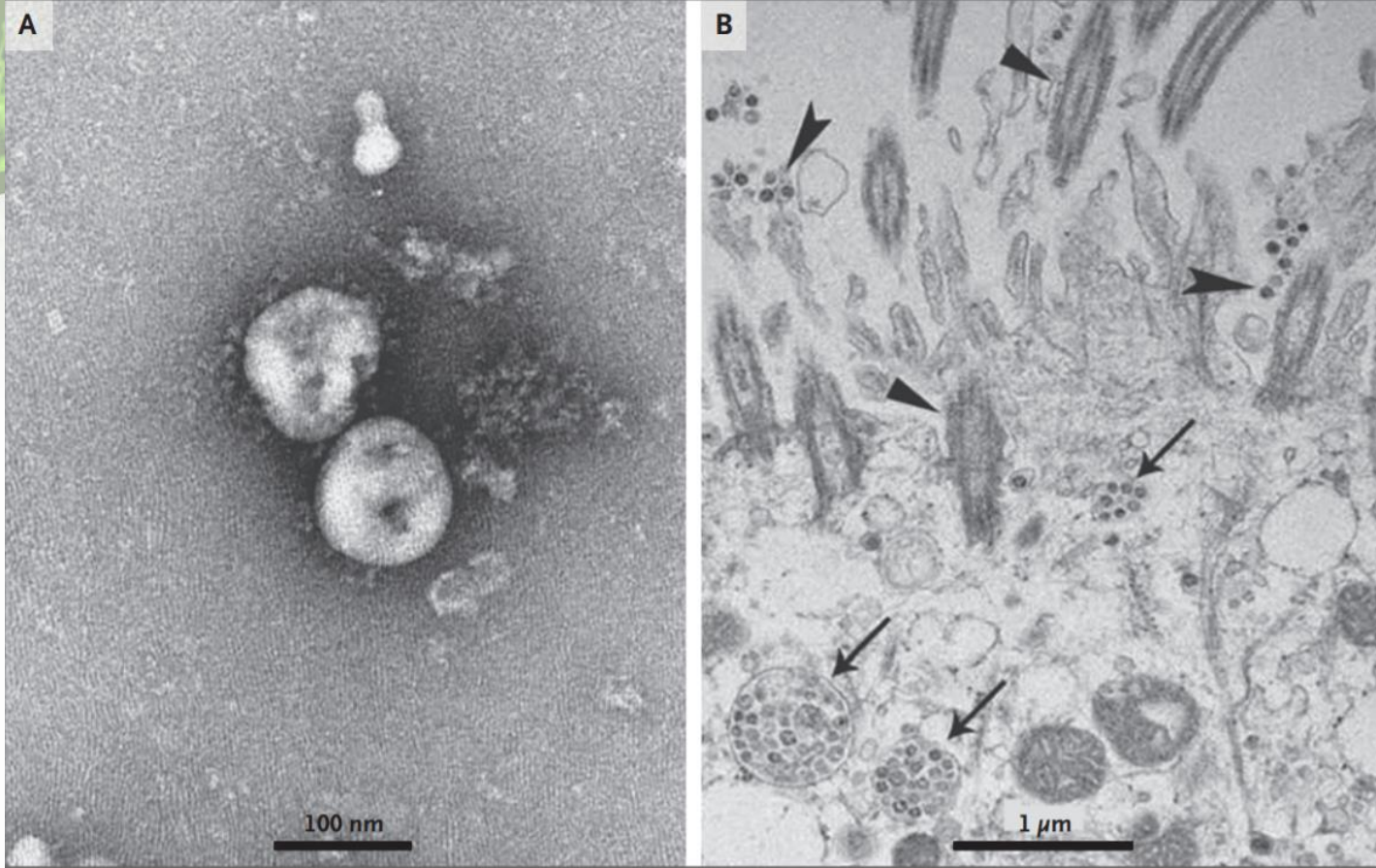


Figure 3. Visualization of 2019-nCoV with Transmission Electron Microscopy.

Negative-stained 2019-nCoV particles are shown in Panel A, and 2019-nCoV particles in the human airway epithelial cell ultrathin sections are shown in Panel B. Arrowheads indicate extracellular virus particles, arrows indicate inclusion bodies formed by virus components, and triangles indicate cilia.

Sekvenace genomu

- RNA extrahována z BAL – templát pro rtPCR
- Illumina sequencing + nanopore sequencing
- 85% identita s bat-SARS-like CoV
- využití sekvenování v diagnostice – detekce virové RNA
– real-time reverse-transcriptase PCR assay



2) Klinická charakteristika COVID-19

Soubor pacientů ve studii

- 1099 pacientů z 552 nemocnic, 30 provincií
- 11.12.2019-31.1.2020
- hospitalizovaní i ambulantní pacienti s lab.dg. – rt RT-PCR assay z nazál. nebo pharyngeál. stěru
- median věku 47 let
- 41,9 % ženy
- při přijetí rozdělení na pacientov v těžkém (173 = 15,7%) a lehkém stavu (926=84,3%) – pacienti s těžkým průběhem v průměru o 7 let starší

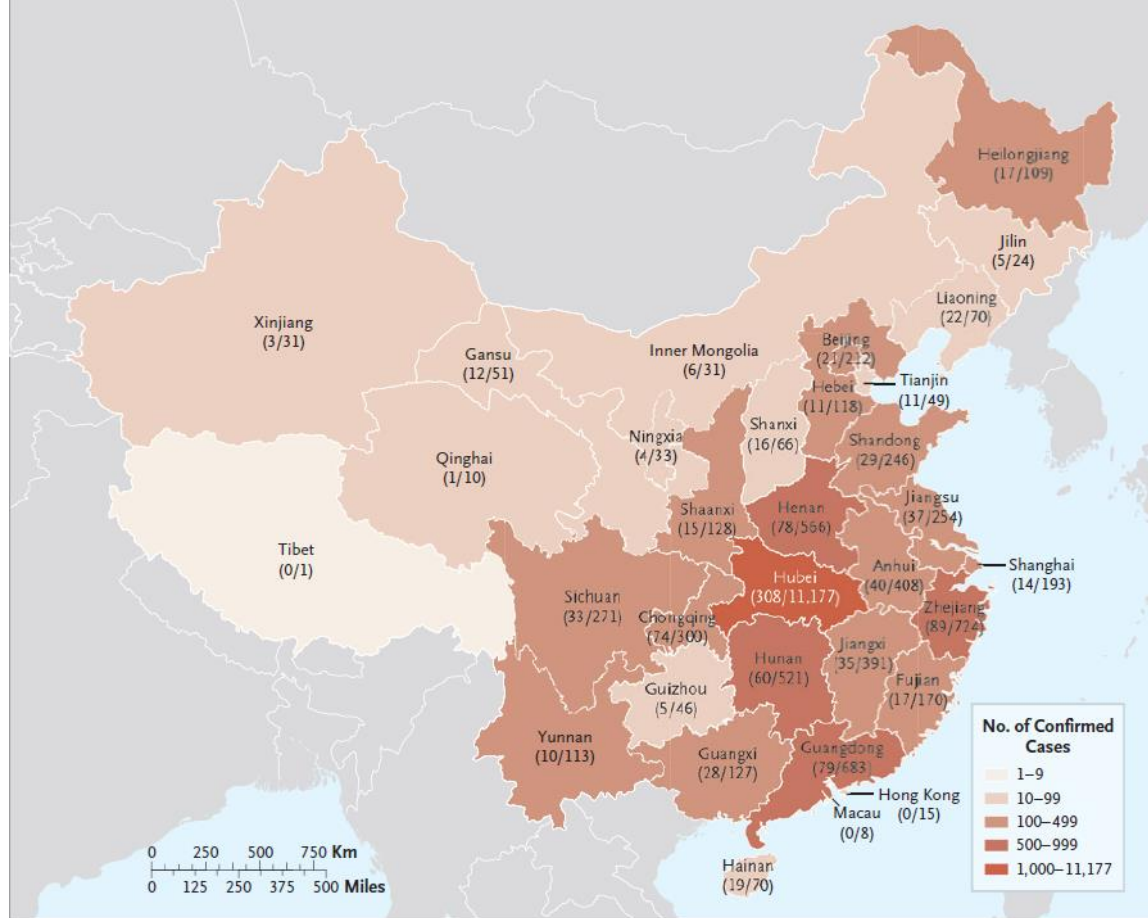


Figure 1. Distribution of Patients with Covid-19 across China.

Shown are the official statistics of all documented, laboratory-confirmed cases of coronavirus disease 2019 (Covid-19) throughout China, according to the National Health Commission as of February 4, 2020. The numerator denotes the number of patients who were included in the study cohort and the denominator denotes the number of laboratory-confirmed cases for each province, autonomous region, or provincial municipality, as reported by the National Health Commission.

Table 1. Clinical Characteristics of the Study Patients, According to Disease Severity and the Presence or Absence of the Primary Composite End Point.*

| Characteristic | All Patients (N = 1099) | Disease Severity | | Presence of Primary Composite End Point† | |
|---|----------------------------|------------------------|---------------------|--|------------------|
| | | Nonsevere (N = 926) | Severe (N = 173) | Yes (N = 67) | No (N = 1032) |
| Age | | | | | |
| Median (IQR) — yr | 47.0 (35.0–58.0) | 45.0 (34.0–57.0) | 52.0 (40.0–65.0) | 63.0 (53.0–71.0) | 46.0 (35.0–57.0) |
| Distribution — no./total no. (%) | | | | | |
| 0–14 yr | 9/1011 (0.9) | 8/848 (0.9) | 1/163 (0.6) | 0 | 9/946 (1.0) |
| 15–49 yr | 557/1011 (55.1) | 490/848 (57.8) | 67/163 (41.1) | 12/65 (18.5) | 545/946 (57.6) |
| 50–64 yr | 292/1011 (28.9) | 241/848 (28.4) | 51/163 (31.3) | 21/65 (32.3) | 271/946 (28.6) |
| ≥65 yr | 153/1011 (15.1) | 109/848 (12.9) | 44/163 (27.0) | 32/65 (49.2) | 121/946 (12.8) |
| Female sex — no./total no. (%) | | | | | |
| 459/1096 (41.9) | 386/923 (41.8) | 73/173 (42.2) | 22/67 (32.8) | 437/1029 (42.5) | |
| Smoking history — no./total no. (%) | | | | | |
| Never smoked | 927/1085 (85.4) | 793/913 (86.9) | 134/172 (77.9) | 44/66 (66.7) | 883/1019 (86.7) |
| Former smoker | 21/1085 (1.9) | 12/913 (1.3) | 9/172 (5.2) | 5/66 (7.6) | 16/1019 (1.6) |
| Current smoker | 137/1085 (12.6) | 108/913 (11.8) | 29/172 (16.9) | 17/66 (25.8) | 120/1019 (11.8) |
| Exposure to source of transmission within past 14 days — no./total no. | | | | | |
| Living in Wuhan | 483/1099 (43.9) | 400/926 (43.2) | 83/173 (48.0) | 39/67 (58.2) | 444/1032 (43.0) |
| Contact with wildlife | 13/687 (1.9) | 10/559 (1.8) | 3/128 (2.3) | 1/41 (2.4) | 12/646 (1.9) |
| Recently visited Wuhan‡ | 193/616 (31.3) | 166/526 (31.6) | 27/90 (30.0) | 10/28 (35.7) | 183/588 (31.1) |
| Had contact with Wuhan residents‡ | 442/611 (72.3) | 376/522 (72.0) | 66/89 (74.2) | 19/28 (67.9) | 423/583 (72.6) |
| Median incubation period (IQR) — days§ | | | | | |
| 4.0 (2.0–7.0) | 4.0 (2.8–7.0) | 4.0 (2.0–7.0) | 4.0 (2.0–7.0) | 4.0 (1.0–7.5) | 4.0 (2.0–7.0) |
| Fever on admission | | | | | |
| Patients — no./total no. (%) | 473/1081 (43.8) | 391/910 (43.0) | 82/171 (48.0) | 24/66 (36.4) | 449/1015 (44.2) |
| Median temperature (IQR) — °C | 37.3 (36.7–38.0) | 37.3 (36.7–38.0) | 37.4 (36.7–38.1) | 36.8 (36.3–37.8) | 37.3 (36.7–38.0) |
| Distribution of temperature — no./total no. (%) | | | | | |
| <37.5°C | 608/1081 (56.2) | 519/910 (57.0) | 89/171 (52.0) | 42/66 (63.6) | 566/1015 (55.8) |
| 37.5–38.0°C | 238/1081 (22.0) | 201/910 (22.1) | 37/171 (21.6) | 10/66 (15.2) | 228/1015 (22.5) |
| 38.1–39.0°C | 197/1081 (18.2) | 160/910 (17.6) | 37/171 (21.6) | 11/66 (16.7) | 186/1015 (18.3) |
| >39.0°C | 38/1081 (3.5) | 30/910 (3.3) | 8/171 (4.7) | 3/66 (4.5) | 35/1015 (3.4) |
| Fever during hospitalization | | | | | |
| Patients — no./total no. (%) | 975/1099 (88.7) | 816/926 (88.1) | 159/173 (91.9) | 59/67 (88.1) | 916/1032 (88.8) |
| Median highest temperature (IQR) — °C | 38.3 (37.8–38.9) | 38.3 (37.8–38.9) | 38.5 (38.0–39.0) | 38.5 (38.0–39.0) | 38.3 (37.8–38.9) |
| <37.5°C | 92/926 (9.9) | 79/774 (10.2) | 13/152 (8.6) | 3/54 (5.6) | 89/872 (10.2) |
| 37.5–38.0°C | 286/926 (30.9) | 251/774 (32.4) | 35/152 (23.0) | 20/54 (37.0) | 266/872 (30.5) |
| 38.1–39.0°C | 434/926 (46.9) | 356/774 (46.0) | 78/152 (51.3) | 21/54 (38.9) | 413/872 (47.4) |
| >39.0°C | 114/926 (12.3) | 88/774 (11.4) | 26/152 (17.1) | 10/54 (18.5) | 104/872 (11.9) |

Table 1. Clinical Characteristics of the Study Patients, According to Disease Severity and the Presence or Absence of the Primary Composite End Point.*

| Characteristic | All Patients (N = 1099) | Disease Severity | | Presence of Primary Composite End Point† | |
|---------------------------------------|----------------------------|------------------------|---------------------|--|------------------|
| | | Nonsevere (N = 926) | Severe (N = 173) | Yes (N = 67) | No (N = 1032) |
| Conjunctival congestion | 9 (0.8) | 5 (0.5) | 4 (2.3) | 0 | 9 (0.9) |
| Nasal congestion | 53 (4.8) | 47 (5.1) | 6 (3.5) | 2 (3.0) | 51 (4.9) |
| Headache | 150 (13.6) | 124 (13.4) | 26 (15.0) | 8 (11.9) | 142 (13.8) |
| Cough | 745 (67.8) | 623 (67.3) | 122 (70.5) | 46 (68.7) | 699 (67.7) |
| Sore throat | 153 (13.9) | 130 (14.0) | 23 (13.3) | 6 (9.0) | 147 (14.2) |
| Sputum production | 370 (33.7) | 309 (33.4) | 61 (35.3) | 20 (29.9) | 350 (33.9) |
| Fatigue | 419 (38.1) | 350 (37.8) | 69 (39.9) | 22 (32.8) | 397 (38.5) |
| Hemoptysis | 10 (0.9) | 6 (0.6) | 4 (2.3) | 2 (3.0) | 8 (0.8) |
| Shortness of breath | 205 (18.7) | 140 (15.1) | 65 (37.6) | 36 (53.7) | 169 (16.4) |
| Nausea or vomiting | 55 (5.0) | 43 (4.6) | 12 (6.9) | 3 (4.5) | 52 (5.0) |
| Diarrhea | 42 (3.8) | 32 (3.5) | 10 (5.8) | 4 (6.0) | 38 (3.7) |
| Myalgia or arthralgia | 164 (14.9) | 134 (14.5) | 30 (17.3) | 6 (9.0) | 158 (15.3) |
| Chills | 126 (11.5) | 100 (10.8) | 26 (15.0) | 8 (11.9) | 118 (11.4) |
| Signs of infection — no. (%) | | | | | |
| Throat congestion | 19 (1.7) | 17 (1.8) | 2 (1.2) | 0 | 19 (1.8) |
| Tonsil swelling | 23 (2.1) | 17 (1.8) | 6 (3.5) | 1 (1.5) | 22 (2.1) |
| Enlargement of lymph nodes | 2 (0.2) | 1 (0.1) | 1 (0.6) | 1 (1.5) | 1 (0.1) |
| Rash | 2 (0.2) | 0 | 2 (1.2) | 0 | 2 (0.2) |
| Coexisting disorder — no. (%) | | | | | |
| Ary | 261 (23.7) | 194 (21.0) | 67 (38.7) | 39 (58.2) | 222 (21.5) |
| Chronic obstructive pulmonary disease | 12 (1.1) | 6 (0.6) | 6 (3.5) | 7 (10.4) | 5 (0.5) |
| Diabetes | 81 (7.4) | 53 (5.7) | 28 (16.2) | 18 (26.9) | 63 (6.1) |
| Hypertension | 165 (15.0) | 124 (13.4) | 41 (23.7) | 24 (35.8) | 141 (13.7) |
| Coronary heart disease | 27 (2.5) | 17 (1.8) | 10 (5.8) | 6 (9.0) | 21 (2.0) |
| Cerebrovascular disease | 15 (1.4) | 11 (1.2) | 4 (2.3) | 4 (6.0) | 11 (1.1) |
| Hepatitis B infection¶ | 23 (2.1) | 22 (2.4) | 1 (0.6) | 1 (1.5) | 22 (2.1) |
| Cancer | 10 (0.9) | 7 (0.8) | 3 (1.7) | 1 (1.5) | 9 (0.9) |
| Chronic renal disease | 8 (0.7) | 5 (0.5) | 3 (1.7) | 2 (3.0) | 6 (0.6) |
| Immunodeficiency | 2 (0.2) | 2 (0.2) | 0 | 0 | 2 (0.2) |

The denominators of patients who were included in the analysis are provided if they differed from the overall numbers in the group. Percentages may not total 100 because of rounding. Covid-19 denotes coronavirus disease 2019, and IQR interquartile range.

The primary composite end point was admission to an intensive care unit, the use of mechanical ventilation, or death.

These patients were not residents of Wuhan.

Data regarding the incubation period were missing for 808 patients (73.5%).

Příznaky a inkubační doba

Nejčastější

- Kašel 67,8 %
- Horečka 43%
- Únava 38,1%
- Zahlenění 33,7%
- Dyspnoe 18,7%

Vzácné

- Exantém 0,2%
- Lymfadenopatie 0,2%
- Konjunktivitis 0,8%
- Hemoptysis 0,9%

| Virus | Incubation Period (typical cases) |
|----------------------------------|---|
| Novel Coronavirus (2019-nCoV) | 2-14 or 0-24 days * |
| SARS | <u>2-7 days</u> , as long as 10 days |
| MERS | <u>5 days</u> (range: 2-14) |
| Swine Flu | <u>1-4 days</u> , as long as 7 days |
| Seasonal Flu | 2 days (1-4 range) |

- ## Inkubační doba
- **median 4 dny (2-7 dnů)**

- **WHO** mezi 2 a 14 dnů
- **National Health Commission (NHC)** of China 10-14 dnů
- **DXY.cn**, a leading Chinese online community for physicians and health care professionals, nejčastěji 3-7 dnů, max 14
 - ??? > 14 dnů

Primární outcome

- hospitalizace na JIP
- nutnost invazivní/neinvazivní ventilace
- smrt pacienta

Table 3. Complications, Treatments, and Clinical Outcomes.

| Variable | All Patients (N= 1099) | Disease Severity | | Presence of Composite Primary End Point | |
|--|---------------------------|------------------------|---------------------|---|------------------|
| | | Nonsevere (N = 926) | Severe (N = 173) | Yes (N = 67) | No (N = 1032) |
| Complications | | | | | |
| Septic shock — no. (%) | 12 (1.1) | 1 (0.1) | 11 (6.4) | 9 (13.4) | 3 (0.3) |
| Acute respiratory distress syndrome — no. (%) | 37 (3.4) | 10 (1.1) | 27 (15.6) | 27 (40.3) | 10 (1.0) |
| Acute kidney injury — no. (%) | 6 (0.5) | 1 (0.1) | 5 (2.9) | 4 (6.0) | 2 (0.2) |
| Disseminated intravascular coagulation — no. (%) | 1 (0.1) | 0 | 1 (0.6) | 1 (1.5) | 0 |
| Rhabdomyolysis — no. (%) | 2 (0.2) | 2 (0.2) | 0 | 0 | 2 (0.2) |
| Physician-diagnosed pneumonia — no./total no. (%) | 972/1067 (91.1) | 800/894 (89.5) | 172/173 (99.4) | 63/66 (95.5) | 909/1001 (90.8) |
| Median time until development of pneumonia (IQR) — days* | | | | | |
| After initial Covid-19 diagnosis | 0.0 (0.0–1.0) | 0.0 (0.0–1.0) | 0.0 (0.0–2.0) | 0.0 (0.0–3.5) | 0.0 (0.0–1.0) |
| After onset of Covid-19 symptoms | 3.0 (1.0–6.0) | 3.0 (1.0–6.0) | 5.0 (2.0–7.0) | 4.0 (0.0–7.0) | 3.0 (1.0–6.0) |
| Treatments | | | | | |
| Intravenous antibiotics — no. (%) | 637 (58.0) | 498 (53.8) | 139 (80.3) | 60 (89.6) | 577 (55.9) |
| Oseltamivir — no. (%) | 393 (35.8) | 313 (33.8) | 80 (46.2) | 36 (53.7) | 357 (34.6) |
| Antifungal medication — no. (%) | 31 (2.8) | 18 (1.9) | 13 (7.5) | 8 (11.9) | 23 (2.2) |
| Systemic glucocorticoids — no. (%) | 204 (18.6) | 127 (13.7) | 77 (44.5) | 35 (52.2) | 169 (16.4) |
| Oxygen therapy — no. (%) | 454 (41.3) | 331 (35.7) | 123 (71.1) | 59 (88.1) | 395 (38.3) |
| Mechanical ventilation — no. (%) | 67 (6.1) | 0 | 67 (38.7) | 40 (59.7) | 27 (2.6) |
| Invasive | 25 (2.3) | 0 | 25 (14.5) | 25 (37.3) | 0 |
| Noninvasive | 56 (5.1) | 0 | 56 (32.4) | 29 (43.3) | 27 (2.6) |
| Use of extracorporeal membrane oxygenation — no. (%) | 5 (0.5) | 0 | 5 (2.9) | 5 (7.5) | 0 |
| Use of continuous renal-replacement therapy — no. (%) | 9 (0.8) | 0 | 9 (5.2) | 8 (11.9) | 1 (0.1) |
| Use of intravenous immune globulin — no. (%) | 144 (13.1) | 86 (9.3) | 58 (33.5) | 27 (40.3) | 117 (11.3) |
| Admission to intensive care unit — no. (%) | 55 (5.0) | 22 (2.4) | 33 (19.1) | 55 (82.1) | 0 |
| Median length of hospital stay (IQR) — days† | 12.0 (10.0–14.0) | 11.0 (10.0–13.0) | 13.0 (11.5–17.0) | 14.5 (11.0–19.0) | 12.0 (10.0–13.0) |

| Characteristic | All Patients (N=1099) | Disease Severity | | Presence of Primary Composite End Point† | |
|--|--------------------------|----------------------|-------------------|--|----------------|
| | | Nonsevere (N=926) | Severe (N=173) | Yes (N=67) | No (N=1032) |
| Clinical outcomes at data cutoff— no. (%) | | | | | |
| Discharge from hospital | 55 (5.0) | 50 (5.4) | 5 (2.9) | 1 (1.5) | 54 (5.2) |
| Death | 15 (1.4) | 1 (0.1) | 14 (8.1) | 15 (22.4) | 0 |
| Recovery | 9 (0.8) | 7 (0.8) | 2 (1.2) | 0 | 9 (0.9) |
| Hospitalization | 1029 (93.6) | 875 (94.5) | 154 (89.0) | 51 (76.1) | 978 (94.8) |

‡ For the development of pneumonia, data were missing for 347 patients (31.6%) regarding the time since the initial diagnosis and for 161 patients (14.6%) regarding the time since symptom onset.

† Data regarding the median length of hospital stay were missing for 136 patients (12.4%).

Primární outcome 6,1% ze všech pacientů, 24,9% u těžkého průběhu

- hospitalizace JIP 5,0 %
- invazivní mechanická podpora dýchání 2,3%
 - letalita 1,4%

**Kumulativní riziko 3,6% celkově,
u pacientů s těžkým průběhem
20,6%**

Laboratorní a RDG nález

- 975 CT – 86,2% abnormality („ground-glass opacity, bilateral patchy shadows“)
 - bez CT nálezu 157/877 (17,9%) s lehkým průběhem, 5/173 (2,9%) s těžkým průběhem
- lymfocytopenie 83,2 %
- trombocytopenie 36,2 %
- leukopenie 33,7%
- elevace CRP ve většině případů
- méně časté elevace jater. enzymů, CK, D-dimerů

Table 2. Radiographic and Laboratory Findings.*

| Variable | All Patients (N = 1099) | Disease Severity | | Presence of Composite Primary End Point | |
|---|----------------------------|-------------------------|---------------------|---|---------------------|
| | | Non severe (N = 926) | Severe (N = 173) | Yes (N = 67) | No (N = 1032) |
| Radiologic findings | | | | | |
| Abnormalities on chest radiograph — no./total no. (%) | 162/274 (59.1) | 116/214 (54.2) | 46/60 (76.7) | 30/39 (76.9) | 132/235 (56.2) |
| Ground-glass opacity | 55/274 (20.1) | 37/214 (17.3) | 18/60 (30.0) | 9/39 (23.1) | 46/235 (19.6) |
| Local patchy shadowing | 77/274 (28.1) | 56/214 (26.2) | 21/60 (35.0) | 13/39 (33.3) | 64/235 (27.2) |
| Bilateral patchy shadowing | 100/274 (36.5) | 65/214 (30.4) | 35/60 (58.3) | 27/39 (69.2) | 73/235 (31.1) |
| Interstitial abnormalities | 12/274 (4.4) | 7/214 (3.3) | 5/60 (8.3) | 6/39 (15.4) | 6/235 (2.6) |
| Abnormalities on chest CT — no./total no. (%) | 840/975 (86.2) | 682/808 (84.4) | 158/167 (94.6) | 50/57 (87.7) | 790/918 (86.1) |
| Ground-glass opacity | 550/975 (56.4) | 449/808 (55.6) | 101/167 (60.5) | 30/57 (52.6) | 520/918 (56.6) |
| Local patchy shadowing | 409/975 (41.9) | 317/808 (39.2) | 92/167 (55.1) | 22/57 (38.6) | 387/918 (42.2) |
| Bilateral patchy shadowing | 505/975 (51.8) | 368/808 (45.5) | 137/167 (82.0) | 40/57 (70.2) | 465/918 (50.7) |
| Interstitial abnormalities | 143/975 (14.7) | 99/808 (12.3) | 44/167 (26.3) | 15/57 (26.3) | 128/918 (13.9) |
| Laboratory findings | | | | | |
| Median PaO ₂ :Fio ₂ ratio (IQR) † | 3.9 (2.9–4.7) | 3.9 (2.9–4.5) | 4.0 (2.8–5.2) | 2.9 (2.2–5.4) | 4.0 (3.1–4.6) |
| White-cell count | | | | | |
| Median (IQR) — per mm ³ | 4700 (3500–6000) | 4900 (3800–6000) | 3700 (3000–6200) | 6100 (4900–11,100) | 4700 (3500–5900) |
| Distribution — no./total no. (%) | | | | | |
| >10,000 per mm ³ | 58/978 (5.9) | 39/811 (4.8) | 19/167 (11.4) | 15/58 (25.9) | 43/920 (4.7) |
| <4000 per mm ³ | 330/978 (33.7) | 228/811 (28.1) | 102/167 (61.1) | 8/58 (13.8) | 322/920 (35.0) |
| Lymphocyte count | | | | | |
| Median (IQR) — per mm ³ | 1000 (700–1300) | 1000 (800–1400) | 800 (600–1000) | 700 (600–900) | 1000 (700–1300) |
| Distribution — no./total no. (%) | | | | | |
| <1500 per mm ³ | 731/879 (83.2) | 584/726 (80.4) | 147/153 (96.1) | 50/54 (92.6) | 681/825 (82.5) |

Table 2. Radiographic and Laboratory Findings.*

| Variable | All Patients (N = 1099) | Disease Severity | | Presence of Composite Primary End Point | |
|--|------------------------------|------------------------------|-----------------------------|---|------------------------------|
| | | Non severe (N = 926) | Severe (N = 173) | Yes (N = 67) | No (N = 1032) |
| Platelet count | | | | | |
| Median (IQR) — per mm ³ | 168,000 (132,000–207,000) | 172,000 (139,000–212,000) | 137,500 (99,000–179,500) | 156,500 (114,200–195,000) | 169,000 (133,000–207,000) |
| Distribution — no./total no. (%) | | | | | |
| <150,000 per mm ³ | 315/869 (36.2) | 225/713 (31.6) | 90/156 (57.7) | 27/58 (46.6) | 288/811 (35.5) |
| Median hemoglobin (IQR) — g/dl‡ | 134.0 (119.0–148.0) | 135.0 (120.0–148.0) | 128.0 (111.8–141.0) | 125.0 (105.0–140.0) | 134.0 (120.0–148.0) |
| Distribution of other findings — no./total no. (%) | | | | | |
| C-reactive protein ≥10 mg/liter | 481/793 (60.7) | 371/658 (56.4) | 110/135 (81.5) | 41/45 (91.1) | 440/748 (58.8) |
| Procalcitonin ≥0.5 ng/ml | 35/633 (5.5) | 19/516 (3.7) | 16/117 (13.7) | 12/50 (24.0) | 23/583 (3.9) |
| Lactate dehydrogenase ≥250 U/liter | 277/675 (41.0) | 205/551 (37.2) | 72/124 (58.1) | 31/44 (70.5) | 246/631 (39.0) |
| Aspartate aminotransferase >40 U/liter | 168/757 (22.2) | 112/615 (18.2) | 56/142 (39.4) | 26/52 (50.0) | 142/705 (20.1) |
| Alanine aminotransferase >40 U/liter | 158/741 (21.3) | 120/606 (19.8) | 38/135 (28.1) | 20/49 (40.8) | 138/692 (19.9) |
| Total bilirubin >17.1 μmol/liter | 76/722 (10.5) | 59/594 (9.9) | 17/128 (13.3) | 10/48 (20.8) | 66/674 (9.8) |
| Creatine kinase ≥200 U/liter | 90/657 (13.7) | 67/536 (12.5) | 23/121 (19.0) | 12/46 (26.1) | 78/611 (12.8) |
| Creatinine ≥133 μmol/liter | 12/752 (1.6) | 6/614 (1.0) | 6/138 (4.3) | 5/52 (9.6) | 7/700 (1.0) |
| D-dimer ≥0.5 mg/liter | 260/560 (46.4) | 195/451 (43.2) | 65/109 (59.6) | 34/49 (69.4) | 226/511 (44.2) |
| Minerals§ | | | | | |
| Median sodium (IQR) — mmol/liter | 138.2 (136.1–140.3) | 138.4 (136.6–140.4) | 138.0 (136.0–140.0) | 138.3 (135.0–141.2) | 138.2 (136.1–140.2) |
| Median potassium (IQR) — mmol/liter | 3.8 (3.5–4.2) | 3.9 (3.6–4.2) | 3.8 (3.5–4.1) | 3.9 (3.6–4.1) | 3.8 (3.5–4.2) |
| Median chloride (IQR) — mmol/liter | 102.9 (99.7–105.6) | 102.7 (99.7–105.3) | 103.1 (99.8–106.0) | 103.8 (100.8–107.0) | 102.8 (99.6–105.3) |

Lymphocytopenia was defined as a lymphocyte count of less than 1500 per cubic millimeter. Thrombocytopenia was defined as a platelet count of less than 150,000 per cubic millimeter. To convert the values for creatinine to milligrams per deciliter, divide by 88.4.

Data regarding the ratio of the partial pressure of arterial oxygen to the fraction of inspired oxygen (PaO₂:FiO₂) were missing for 894 patients (81.3%).

Data regarding hemoglobin were missing for 226 patients (20.6%).

Data were missing for the measurement of sodium in 363 patients (33.0%), for potassium in 349 patients (31.8%), and for chloride in 392 patients (35.7%).

Přenos COVID-19

- Kapénková nákaza
 - Cca 1,8 m
- Vzdušný přenos
 - Typické pro respirační viry
 - Kontroverzní
 - FFP3
 - Povrch – sliznice
 - Hodiny až dny (týdny?)

Riziko úmrtí při COVID-19

Na základě věku

| Baseline characteristics | Confirmed cases, N (%) | Deaths, N (%) | Case fatality rate, % |
|--------------------------|---------------------------|------------------|--------------------------|
| Overall | 44,672 | 1,023 | 2.3 |
| Age, years | | | |
| 0–9 | 416 (0.9) | – | – |
| 10–19 | 549 (1.2) | 1 (0.1) | 0.2 |
| 20–29 | 3,619 (8.1) | 7 (0.7) | 0.2 |
| 30–39 | 7,600 (17.0) | 18 (1.8) | 0.2 |
| 40–49 | 8,571 (19.2) | 38 (3.7) | 0.4 |
| 50–59 | 10,008 (22.4) | 130 (12.7) | 1.3 |
| 60–69 | 8,583 (19.2) | 309 (30.2) | 3.6 |
| 70–79 | 3,918 (8.8) | 312 (30.5) | 8.0 |
| → ≥80 | 1,408 (3.2) | 208 (20.3) | 14.8 |

Na základě pohlaví

| Baseline characteristics | Confirmed cases, N (%) | Deaths, N (%) | Case fatality rate, % |
|--------------------------|---------------------------|------------------|--------------------------|
| Sex | | | |
| Male | 22,981 (51.4) | 653 (63.8) | 2.8 |
| Female | 21,691 (48.6) | 370 (36.2) | 1.7 |

Riziko úmrtí při COVID-19

Na základě komorbidit

| Baseline characteristics | Confirmed cases, N (%) | Deaths, N (%) | Case fatality rate, % |
|---------------------------------|---------------------------|------------------|--------------------------|
| Comorbid condition [†] | | | |
| Hypertension | 2,683 (12.8) | 161 (39.7) | 6.0 |
| Diabetes | 1,102 (5.3) | 80 (19.7) | 7.3 |
| Cardiovascular disease | → 873 (4.2) | 92 (22.7) | 10.5 |
| Chronic respiratory disease | 511 (2.4) | 32 (7.9) | 6.3 |
| Cancer (any) | 107 (0.5) | 6 (1.5) | 5.6 |
| None | 15,536 (74.0) | 133 (32.8) | 0.9 |
| Missing | 23,690 (53.0) | 617 (60.3) | 2.6 |

Porovnání s jinými koronaviry

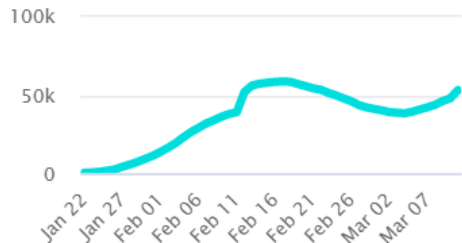
| Virus | Death Rate |
|-------------------------------------|------------|
| Wuhan Novel Coronavirus (2019-nCoV) | 2%* |
| SARS | 9.6% |
| MERS | 34% |
| Swine Flu | 0.02% |

3) Aktuální informace

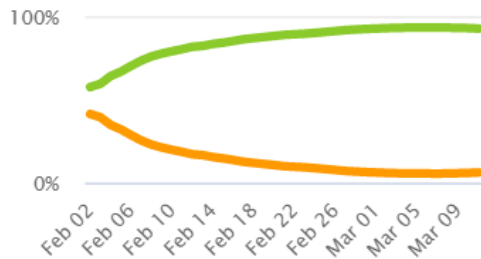
11.03.2020 WHO

vyhlásilo pandemii

ACTIVE CASES



CLOSED CASES



ACTIVE CASES

59,158

Currently Infected Patients

53,319 (90%)
in Mild Condition

5,839 (10%)
Serious or Critical

[Show Graph](#)

CLOSED CASES

73,837

Cases which had an outcome:

68,891 (93%)
Recovered / Discharged

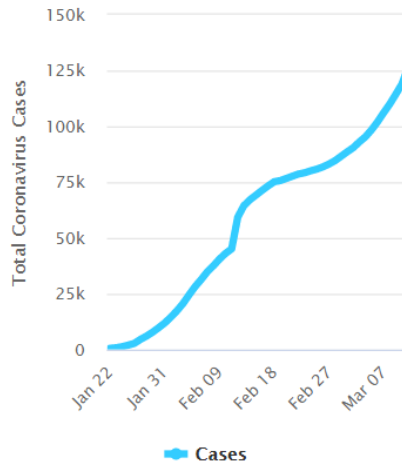
4,946 (7%)
Deaths

[Show Graph](#)

linear logarithmic

Total Cases

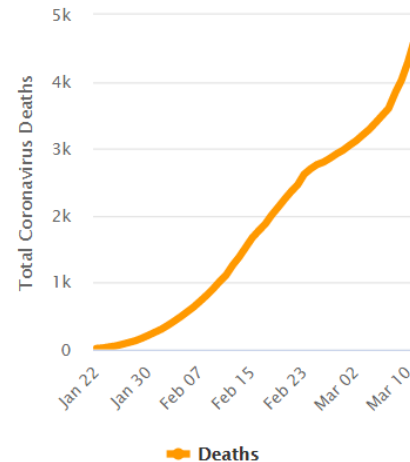
(Linear Scale)



linear logarithmic

Total Deaths

(Linear Scale)



| Country, Other | Total Cases | New Cases | Total Deaths | New Deaths | Total Recovered | Active Cases | Serious, Critical |
|----------------------------------|-------------|-----------|--------------|------------|-----------------|--------------|-------------------|
| China | 80,796 | +18 | 3,169 | +11 | 62,826 | 14,801 | 4,257 |
| Italy | 15,113 | +2,651 | 1,016 | +189 | 1,258 | 12,839 | 1,153 |
| Iran | 10,075 | +1,075 | 429 | +75 | 3,276 | 6,370 | |
| S. Korea | 7,869 | +114 | 66 | +6 | 333 | 7,470 | 54 |
| Spain | 3,059 | +782 | 86 | +31 | 189 | 2,784 | 126 |
| Germany | 2,502 | +536 | 5 | +2 | 25 | 2,472 | 9 |
| France | 2,281 | | 48 | | 12 | 2,221 | 105 |
| USA | 1,396 | +95 | 39 | +1 | 15 | 1,342 | 10 |
| Switzerland | 867 | +215 | 6 | +2 | 4 | 857 | |
| Norway | 727 | +98 | | | 1 | 726 | |
| Diamond Princess | 696 | | 7 | | 325 | 364 | 32 |
| Japan | 691 | +52 | 17 | +2 | 118 | 556 | 29 |
| Sweden | 683 | +183 | 1 | | 1 | 681 | 2 |
| Denmark | 674 | +160 | | | 1 | 673 | 2 |
| Netherlands | 614 | +111 | 5 | | 2 | 607 | 1 |
| UK | 590 | +130 | 10 | +2 | 18 | 562 | |
| Belgium | 399 | +85 | 3 | | 1 | 395 | 2 |

| Country, Other | Total Cases | New Cases | Total Deaths | New Deaths | Total Recovered | Active Cases | Serious, Critical |
|----------------|-------------|-----------|--------------|------------|-----------------|--------------|-------------------|
| Austria | 361 | +115 | 1 | +1 | 4 | 356 | 1 |
| Qatar | 262 | | | | | 262 | |
| Bahrain | 195 | | | | 35 | 160 | 1 |
| Singapore | 187 | +9 | | | 96 | 91 | 12 |
| Malaysia | 158 | +9 | | | 26 | 132 | 2 |
| Australia | 156 | +28 | 3 | | 26 | 127 | 1 |
| Hong Kong | 130 | | 3 | | 77 | 50 | 6 |
| Canada | 122 | +12 | 1 | | 9 | 112 | 1 |
| Finland | 109 | +44 | | | 1 | 108 | |
| Iceland | 109 | +24 | | | | 109 | |
| Israel | 100 | +3 | | | 4 | 96 | 2 |
| Greece | 99 | | 1 | +1 | | 98 | 2 |
| Czechia | 96 | +2 | | | | 96 | |
| Slovenia | 89 | +32 | | | | 89 | |
| UAE | 85 | +11 | | | 17 | 68 | 2 |
| Kuwait | 80 | +8 | | | 5 | 75 | 4 |
| Iraq | 79 | +8 | 8 | +1 | 20 | 51 | |
| Portugal | 78 | +17 | | | 1 | 77 | 1 |

Aktuální data k 12.03.2020

Zdroj: <https://www.worldometers.info/coronavirus/#repro>

124 847

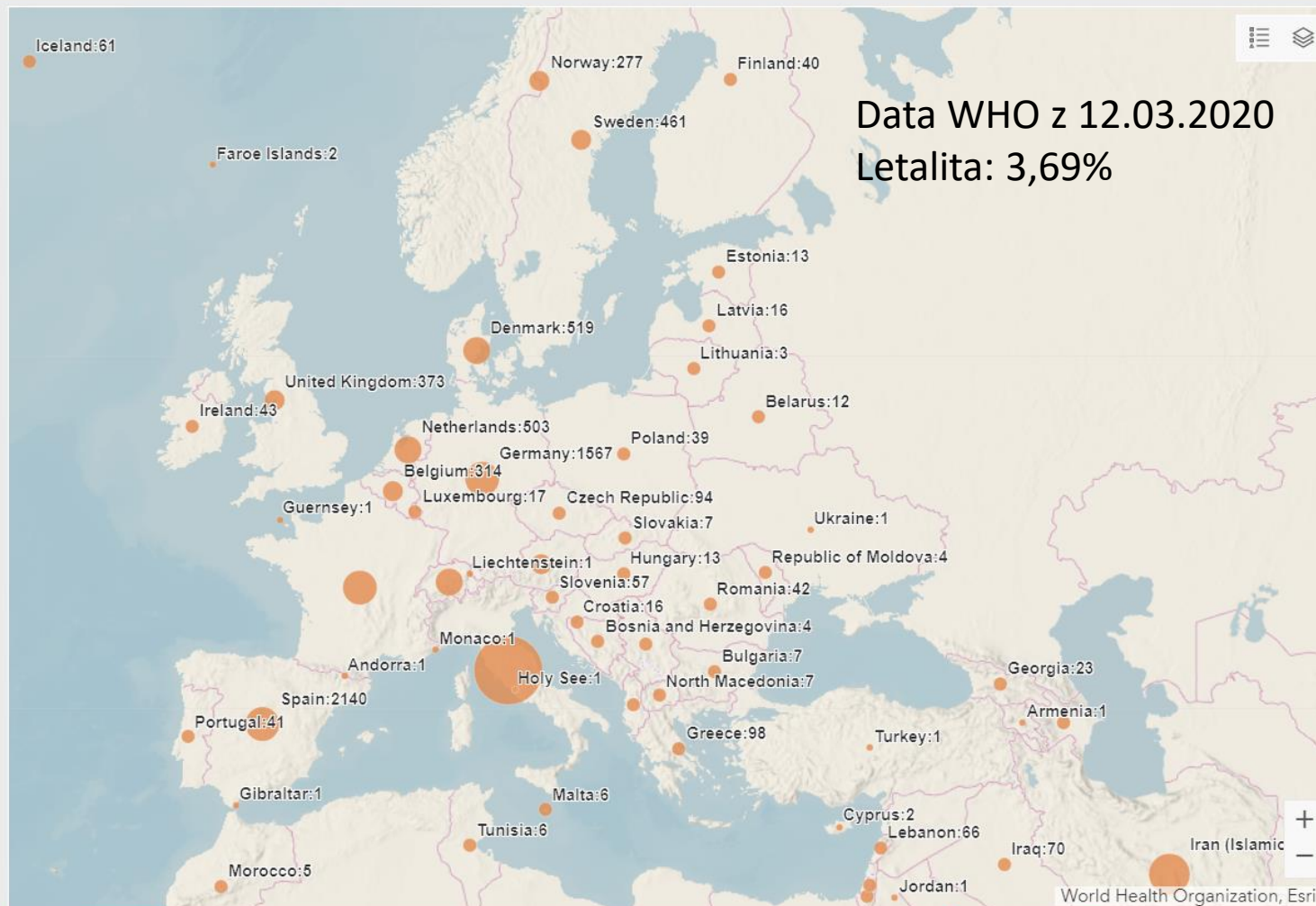
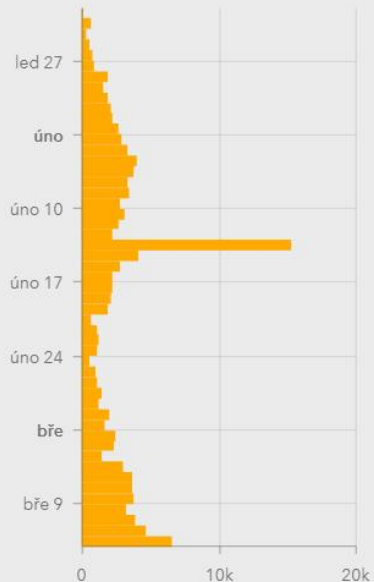
confirmed cases

4 613

deaths

118

countries, areas or territories with cases



Diskuse a závěr



- dominantní symptomy kašel, febrilie, únava, dušnost
- rychlé šíření, ale nižší úmrtnost v porovnání s MERS a SARS
 - Letalita
 - 1,4% - 2,3% - dle aktuálních dat z WHO 3,69 %
 - v skutečnosti se dá předpokládat nižší, do studie nejsou zavzatí pacienti s CoV-SARS-2 bez klinické manifestace
- cesta přenosu kapénkovou nákazou a přenosem tělními tekutinami
- potřeba lépe porozumět nemoci a projevům ještě před rozvojem pneumonie (v této studii u 8,9% pacientů detekován virus před rozvojem pneumonie nebo bez rozvoje)
 - potřeba včasné izolace, diagnostiky a managementu

Zdroje

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- [Report of the WHO-China Joint Mission on Coronavirus Disease 2019 \(COVID-19\)](#) [Pdf] - World Health Organization, March 11, 2020
- <https://www.worldometers.info/coronavirus/> - March 12, 2020
- <https://www.who.int/emergencies/diseases/novel-coronavirus-2019/situation-reports/> - March 12, 2020
- http://www.mzcr.cz/dokumenty/koronavirus-2019-ncov-aktualni-data_18455_4122_1.html – March 12, 2020

A microscopic view of a hand with green virus particles. The hand is shown in a dark green, almost black, background. The fingers are spread out, and several green, spiky virus particles are visible on the surface of the hand. The virus particles have a central core and many thin, hair-like projections extending from the surface. The overall image has a high-contrast, scientific feel.

Děkuju za pozornost!