Eleven things you need to know NOW about COVID-19

(As of November 2022)

Number of cases and virulence of COVID-19

- 1. How many people have been diagnosed with COVID-19 in Japan?
- 2. How many people will become very sick or die after being diagnosed with COVID-19?
- 3. Who is at higher risk for developing severe COVID-19 illness?
- 4. Is the number of individuals diagnosed with COVID-19 higher in Japan than in other countries?

Infectivity of COVID-19

- 5. How long does a person with COVID-19 remain infectious?
- 6. Does everyone with COVID-19 infect others?
- 7. What precautions should we take to prevent the spread of COVID-19?

Testing and Treatment for COVID-19

- 8. What tests are being used to diagnose COVID-19?
- 9. What are the treatment options for COVID-19?
- 10. Which COVID-19 vaccine is used, and how far along is the vaccination process in Japan?

Variants of COVID-19

11. What is known about the COVID-19 variants?

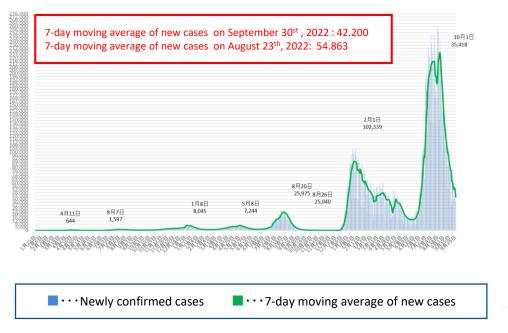
1. How many people have been diagnosed with COVID-19 in Japan?

As of 0:00 November 1st 2022, **22,360,872 people have been diagnosed with COVID-19 in Japan. This accounts for 17.7 %** of the total population.

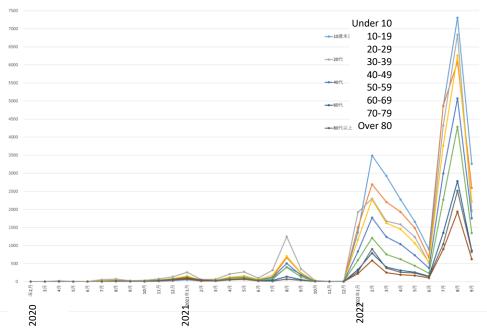
For latest updates, please refer to the following link:

https://www.mhlw.go.jp/stf/covid-19/kokunainohasseijoukyou.html

Number of newly confirmed cases per 100,000 population



Number of newly confirmed cases per 100,000 population by age group



^{*}The number of cases include only those who tested positive. Therefore it may miss those people who were actually infected but without symptoms and therefore did not seek medical care.

2. Among those diagnosed with COVID-19, how many people will get very sick or die?

Proportion of individuals getting severely ill or death differs by age; older tend to higher and younger tend to be lower. Overall severity and case death rates seem to have been in declining trends in the past, and among those diagnosed with COVID-19 from March to April 2022 when the Omicron strain was the mainstay of the epidemic,

- The rate of severity rate is 0.03% in age 50 or younger, 1.50% in those over 60s.
- The rate of case death rate is 0.01% in age 50 or younger, 1.13% in those over 60s.

* "Severe illnesses" include either one or more of followings: admission to the Intensive Care Units, use of respirators, or deaths.

At diagnosis

Fever, cough and other symptoms

Severe illness

0.01% of those in their 50s or younger
1.50% of those over age 60

Death

Severity rates among diagnosed cases (%)

Death rates among diagnosed cases(%)

	0 - 9	10- 19	20- 29	30- 39	40- 49	50- 59	60- 69	70- 79	80- 89	90-		0 - 9	10- 19	20- 29	30- 39	40- 49	50- 59	60- 69	70- 79	80- 89	90-
July-Oct,2021	0.05	0.05	0.01	0.31	1.04	2.25	3.62	4.43	9.17	12.68	July-Oct,2021	0.00	0.00	0.00	0.06	0.08	0.42	0.86	2.35	6.21	11.97
Jan-Feb,2022	0.02	0.00	0.00	0.01	0.05	0.12	0.58	2.03	4.25	6.48	Jan-Feb,2022	0.00	0.00	0.00	0.00	0.02	0.03	0.29	1.23	3.67	6.21
Mar-Apl,2022	0.02	0.00	0.00	0.02	0.03	0.16	0.32	1.54	3.10	4.32	Mar-Apl,2022	0.00	0.00	0.00	0.00	0.01	0.05	0.10	0.94	2.67	4.05

Using data from Ishikawa, Ibaraki, and Hiroshima prefectures that provided cooperation, we tentatively calculated the severe disease and fatality rates by age group and vaccination history infected with COVID-19 during the period. The number of death cases includes those who were COVID-19 positive and died of any cause. Data for July-Oct 2021, Jan-Feb 2022, and Mar-Apr 2022 are based on the status of the patient end of the treatment and hospitalization period or at least 2 month after notification, and as of 3/31 and 5/31. The number of severity ad death cases may increase in the future.

Death rates from 2022/6/1 to 2022/8/30													
	0 - 9	10-19	20-29	30-39	40-49	50-59	60-69	70-79	80-89	90-			
2022/6/1 ~ 8/30	11	4	13	25	80	163	373	1,212	2,828	2,279			

COVID-19 Advisory Board as of 2022/9/7: https://www.mhlw.go.jp/content/10900000/000987078.pdf

3. Among those diagnosed with COVID-19, who is at increased risk for severe illness and death?

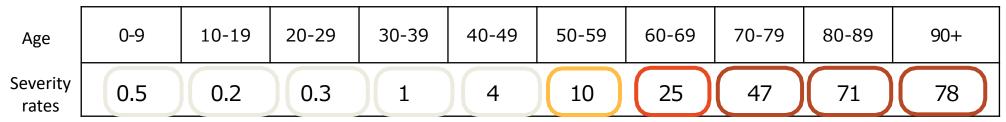
Among those diagnosed with COVID-19, the elderly, those with underlying medical conditions, and some pregnant women in their third trimesters are more likely than others to develop severe illness or die.

Comorbidities known to cause more severe illness include:

Chronic Obstructive Pulmonary Diseases (COPD), Chronic Kidney Diseases, Diabetes, Hypertension, Cardiovascular diseases, Obesity, and Smoking.

Full vaccination (two doses) is effective in preventing severe illness.

Proportion of individuals becoming severely ill by age group (shown as a relative risk when the risk of those in their 30s is considered as a reference (1.00))

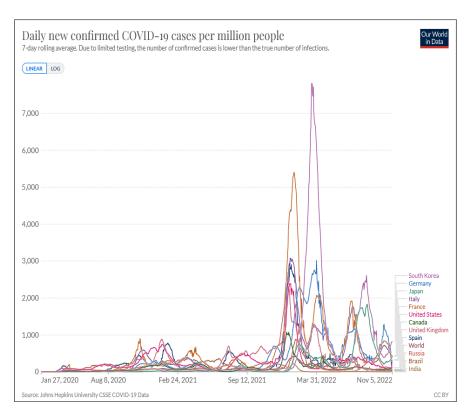


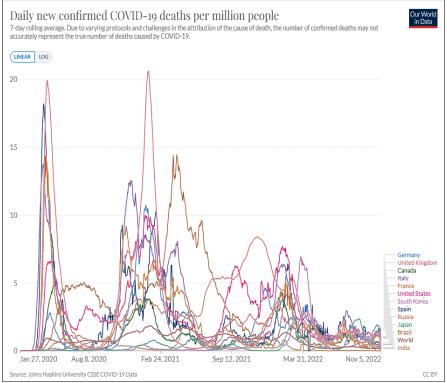
Comorbidities known to cause more severe illness:



4. Is the number of individuals diagnosed with COVID-19 higher in Japan than in other countries?

The number of confirmed cases of infection and deaths per 1 million population in Japan is higher than in the major countries. (This difference should be understood with caution, as each country has the different methods of reporting the number of infections and deaths.)





5. How long does someone with COVID-19 remain infectious?

Individuals with COVID-19 become infectious from **2 days prior to until approximately 7 to 10 days** after the onset of symptoms.*Viral shedding is believed to be particularly high around the time of symptom onset.

Therefore, individuals diagnosed with COVID-19 should **refrain from going out for non-urgent reasons**, **regardless of symptoms**, **to stop the spread of infection**.

* From "Clinical Management of Patients with COVID-19." Version 6.2

6. What proportion of individuals with COVID-19 ends up infecting others?

Less than 20% of individuals diagnosed with COVID-19 infect others, while others do not. Therefore, unless a super-spreading event where an individual with COVID-19 infect many others in an environment like 3Cs without proper precautions happens, spread of SARS-CoV-2 (which causes COVID-19) could be controlled. You are strongly encouraged to take precautions such as not going out for non-urgent purposes when you are sick, and wearing a mask when meeting with others.

Ueki, H., Furusawa, Y., Iwatsuki-Horimoto, K., Imai, M., Kabata, H., Nishimura, H., & Kawaoka, Y. (2020). Effectiveness of Face Masks in Preventing Airborne Transmission of SARS-CoV-2. mSphere, 5(5), e00637-20.

^{*}Wearing masks reduce the amount of virus inhaled by individuals in close proximity to those with COVID-19. (By 60-80% when worn by individuals with COVID-19, and by 20-40% when worn by someone contacting with individuals with COVID-19).

7. What precautions should we take to prevent the spread of COVID-19?

Situations such as social gatherings, long feasts in large groups, having conversation without masks, living together in small limited spaces, and switching locations increase the risk of infection and should be avoided. This is why the risk of infection increases in a 3Cs (Closed spaces, Crowded places, Close-contact settings) environment. Please aim at No 3Cs.

"5 situations" that increase the risk of infection

Social gatherings with drinking alcohol

- Drinking alcohol improves mood and at the same time decreases attention. In addition, hearing is dulled and it leads to speaking in a louder voice.
- The risk of infection increases when large numbers
- of people are in a small space for a long time.

 In addition, sharing glasses and chopsticks increases the risk of infection.



Situation[®]

Long feasts in large groups

- Long-term meals, dinner receptions, drinking alcohol at night increase the risk of infection compared to a short meal.
- The risk of infection is increased by eating and drinking in a large group of people, for example, 5 or more people, because in groups you have to talk louder and droplets of saliva spread more often.



Situation(3)

Conversation without a mask

- Talking at close range without a mask increases the risk of airborne or micro-droplet infection.
- Cases of infection without masks were observed during gatherings in karaoke machines.
- Please be careful when traveling by car or bus.



Situation ®

Living together in a small limited space

- Living together in a small limited space increases the risk of infection because the enclosed space is shared by several people for a long time.
- There have been reports of suspected infections in common areas such as dormitory bedrooms and bathrooms.



Situation ⑤

Switching locations

- When you move to another location, such as when you take a break in a workplace, the risk of infection may increase due to the feeling of relaxation and changes of the environment.
- Suspicious cases of infection were identified in breaking rooms, smoking areas and changing rooms.



8. What tests are being used to diagnose COVID-19?

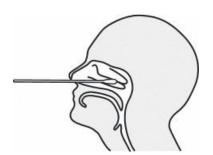
Tests for diagnosing COVID-19 include Nucleic acid detection tests (including PCR test), qualitative antigen test, and quantitative antigen test. These tests are all designed to detect SARS-CoV-2 virus (which causes COVID-19) in one's body and therefore can be used to see if someone is currently infected with the virus. Recent development of new method has enabled use of saliva and nasal cavity swab samples in selected situations and selected patient groups.

Antibody tests are only used to see if a person has previously been infected with SARS-CoV-2 (virus that causes COVID-19). Antibody tests are not for use to diagnose acute infection.

		acid detec uding PCR	tion tests (test)		Antigen te Quantitativ			Antigen tes Qualitative			
Intended to	use for···	Nasopharynx	Nasal cavity	Saliva	Nasopharynx	Nasal cavity	Saliva	Nasopharynx	Nasal cavity	Saliva	
Symptomatic	Within 9 days of symptom onset	0	0	\circ	0	0	0	0	0	○ _{*3}	
Individuals 10 days or more from symptom onset		0	0	×	0	0	×	△ *1	△ *1	×	
Asymptomatic i	ndividuals	0	\circ	\circ	0	× *2	\circ	× *2	× *2	×	

^{*1} Additional Nucleic acid detection tests (e.g. nasopharyngeal PCR tests) is advised when tested negative. *2 Not advised to use for confirmatory tests. Can be used as screening tests in limited settings such as hospitals or nursing homes in disease spreading areas on condition that preventive measures should still be continued for individuals tested negative. *3 Applicable to products with regulatory approval for saliva specimens.

Example of Specimen Collection for Qualitative Antigen Test



Insert a swab through the nose and rub the nasopharynx several times (Performed by a healthcare provider, only)



Insert a swab about 2 cm from the nose, rotate it 5 times, and let it stand for about 5 seconds.
(Self specimen collection is possible)

Nasopharyngeal specimen collection

Nasal cavity Specimen collection

9. What are the treatment options for COVID-19?

Most patients with mild COVID-19 recover without any specific treatment. So only supportive therapy such as antipyretics would be provided when necessary. Patients who are at risk of severe illness are encouraged to receive neutralizing antibody drugs or oral antivirals for prevention. In case of respiratory failure, oxygen therapy along with antivirals, steroids(to control inflammation), immunomodulators, and neutralizing antibodies will be given. And if not responding well enough, intensive care with mechanical ventilation might be an option.

Establishment of treatment options described above seems to have resulted in low case fatality rates for the hospitalized COVID-19 patients. Be sure to consult your GP or nearby clinic in case you have fever, cough or symptoms suggestive of COVID-19.

Drugs approved for the treatment of COVID-19 in Japan

- Remdesivir (Veklury®)
- Molnupiravir (LAGEVRIO®)
- Nirmatrelvir/Ritonavir

(Paxlovid®)

- Sotrovimab (XEVUDY®)
- Casirivimab/Imdevimab

(Ronapreve®)

Tixagevimab/Silgavimab

(Evusheld®)

- Dexamethasone
- Baricitinib (olumiant®)
- Tocilizumab (ACTEMRA®)

Mild Moderate I Moderate II Severe Breathing treatment Oxygen therapy Intubation **HFNC** included Ventilation/ Use of filtered CPAP and **ECMO** NPPV when necessary Aggressive repositioning, including prone therapy **Antivirals** Remdesivir *1 Nirmatrelvir/Ritonavir *2 Molnupiravir *2 *1 Not indicated for patients with mild COVID-19, but Neutralizing antibody drugs should consider for patients at risk for severe illness. *2 Patients at risk for severe illness are eligible. Sotrovimab *3 Patients infected with the Omicron are not eligible. Casirivimab/Imdevimab *3 *4 Use with steroids. Steroids (Dexamethasone, etc.) Immunomodulator/ **Immunosuppressant** Baricitinib Tocilizumab *4 **Anticoagulants** Heparin

(As of October 11^{th} , 2022) * From "Clinical Management of Patients with COVID-19." Version 6.2

10. Which COVID-19 vaccine is used, and how far along is the vaccination process?

- **Vaccines and Vaccination Eligibility:** Pfizer Inc., Takeda/Moderna Inc., AstraZeneca plc., and Takeda/Novavax are approved, as of November 1st,2022. [Messenger RNA Vaccines] · Moderna; Initial vaccination (1st and 2nd doses) is administered to those aged 12 years and older.
- •The Pfizer; The first vaccination (*) is administered to each subject using three types of vaccines: for ages 12 and older, for ages 5-11, and for ages 6 months-4 years old. In addition, one dose of Omicron strain-compatible vaccine is administered to each person aged 12 years and older who has completed the 1st and 2nd doses. For those aged 5-11 years, 3rd dose is administered using the vaccine for 5-11 years (conventional type). * The vaccines for 12 years of age and older and for 5-11 year olds are administered as two doses in the initial vaccination, while the vaccine for 6 months to 4 years of age is administered as three doses in the initial vaccination.
- [Viral Vector Vaccines] •Takeda/Moderna; viral vector vaccine whose production technology was transferred from Novavax, is available for 1st and 2nd doses for persons aged 12 years and older. In addition, one additional dose per person after the third dose is available for those who have completed 1st and 2nd doses and are 18 years of age or older.
- Efficacy: Based on the current knowledge, experts have concluded that the additional vaccination with the Omicron strain-compatible vaccine is expected to be more effective than the conventional vaccine in preventing severe cases and, although possibly for a shorter period of time, in preventing the onset of disease and infection against the Omicron strain. It is also expected to be more likely to be effective against future mutant strains. In addition, the vaccine for children aged 5 to 11 years (conventional type) and the vaccine for children aged 6 months to 4 years (conventional type) have also been reported to be effective in preventing the onset of disease under an outbreak of Omicron strains.
- Safety: It has been reported that the injected area may cause pain, fatigue, headache, muscle and joint pain, chills, diarrhea, fever, and other symptoms after vaccination. No serious concerns have been identified based on the information available at this time.

Vaccination status in Japan (As of November 1st ,2022)

Source: Webpage of Prime Minister's Office of Japan Vaccination Rate 5~11 12~19 20's 30's 40's 50's 60~64 65~69 70's 80's 90's Over 100 20.0% 75.5% 82.2% 81.8% 84.4% 91.1% 92.6% 89.8% 94.4% 96.8% 98.3% 94.5% More Than Once 94.2% 19.1% 74.7% 81.5% 81.3% 84.0% 90.8% 92.4% 89.6% 96.5% 97.8% 93.6% Two Doses 4.0% 42.0% 53.2% 56.8% 64.6% 78.6% 85.9% 85.8% 91.3% 93.2% 93.6% 88.5% Third Doses

Number of Vaccinations	5~11	12~19	20's	30's	40's	50's	60~64	65~69	70's	80's	90's	Over 100
Population	7,317,297	8,887,053	12,680,325	14,065,328	17,965,890	17,245,283	7,378,055	7,770,002	16,284,498	9,382,746	2,404,985	86,607
More Than Once	1,464,623	6,707,774	10,428,540	11,503,154	15,163,391	15,704,650	6,832,311	6,977,751	15,374,043	9,084,299	2,363,120	81,832
Two Doses	1,398,455	6,641,412	10,336,977	11,429,431	15,097,240	15,660,278	6,818,991	6,964,938	15,343,336	9,055,292	2,351,575	81,106

11. What is known about the SARS-CoV-2 (virus that causes COVID-19) variants?

Viruses usually mutate and change gradually over time as they grow or spread, and is believed to show approximately one base mutation in two weeks. Currently variants of SARS-CoV-2 (B.1.1.529 lineage, called Omicron variant) have been reported mainly from around the world and we still have to be vigilant against such new variants.

In response to this, the Ministry of Health, Labour and Welfare (MHLW) has been conducting a nationwide genomic sequencing of the SARS-CoV-2 to monitor for the variants. The MHLW communicates closely with the World Health Organization (WHO) and experts to analyze and characterize variants, and is strengthening the national surveillance system. The MHLW also strengthens its testing capacity and epidemiological investigation (tracing) when cases of such variants are reported to prevent further spread.

Preventive measures such as avoiding the "3Cs" (and "Five situations" that increase the risk of infection), wearing masks, and washing hands remain as effective against the variants. Therefore individuals are strongly encouraged to continue taking these precautions.

■ COVID-19 Variants

https://www.mhlw.go.jp/stf/covid-19/kokunainohasseijoukyou.html#h2_1

■ COVID-19 Control Advisory Board

https://www.mhlw.go.jp/stf/seisakunitsuite/bunya/0000121431 00395.html

■ National Institute of Infectious Diseases

https://www.niid.go.jp/niid/ja/2019-ncov/2551-cepr/10745-cepr-topics.html