

HPAI A(H5) Scenario-Based Human Health Risk Assessment for the United States Center for Outbreak Response Innovation (CORI) Updated as of January 9, 2024

In this update, the Center for Outbreak Response Innovation (CORI) reports the latest developments in the H5N1 outbreak; the risk levels remain unchanged.

Recent developments highlight the potentially severe human and animal health consequences of H5N1 and the continued, increasing importance of enhanced surveillance as well as coordinated containment and prevention activities but do not immediately change the current risk level. For the risk scenario to increase, human-to-human transmission would need to be confirmed, and/or evidence of increasing severity of disease would need to be confirmed. For the risk level to decrease, there would need to be a decline in human cases and a reduction in opportunities for reassortment (eg widespread utilization of PPE by farm workers and others in contact with animals and/or a decline in animal cases).

	Risk to farm workers	Risk to other people in contact with affected workers and animal populations	Risk to healthcare workers	Risk to the US general public
Scenario for Increased Potential for Human	High	Moderate	Low*	Low*
Adaptation and Increased Human				
Reports: Increased potential for reassortment				
and human adaptation, increased reports of				
human infections, potential early				
laboratory/epidemiological/sequencing				
evidence for human-to-human transmission but				
still no human-to-human transmission				
confirmed				

*While the immediate risk to the general public and healthcare workers is still currently low, the long-term consequences of continued, uncontrolled transmission present a high risk to all populations. For this reason, along with the uncertainty and complexity of these events, CORI will continue to monitor the situation and update this risk assessment. For a detailed analysis, including limitations and recommendations see the next page.

Critical Updates:

- The <u>Louisiana Department of Health</u> has reported the first H5N1-related death in the United States. The individual tested positive and was hospitalized in November with known risk factors, including age over 65, and reported preexisting conditions.
- Genetic sequencing of the Louisiana mortality case conducted by the CDC indicates that the individual had several genetic mutations found in other severe H5N1 cases, one of which was the E186E/D mutation identified in the British Columbia severe case in late 2024 that is known to be associated with better binding to the human lung receptors. Genetic analysis of the two specimens collected from the Louisiana human case and the specimens from the wild birds in Louisiana and on the individual's property indicate the mutation most likely arose during the course of human clinical infection. There is no evidence of human-to-human transmission currently.
- Reports of H5N1 in domestic house cats are becoming common. Pet owners can take
 precautions by not feeding their animals raw food, being mindful to check for and remove
 any recalled contaminated food, limiting their pets' exposure to wild birds or other outdoor
 flocks, and contacting their vets if they exhibit signs of illness (lethargy, loss of appetite, etc)



Routine Surveillance Updates

- The Centers for Disease Control and Prevention (CDC) is reporting 66 confirmed human cases
 of H5 in the United States Outbreak as of January 9, 2024. Two of these cases have not had a
 known source of infection and reported no contact with animals or raw milk, while 23 have had
 contact with poultry, 40 have had contact with dairy cows, and one had exposure to a backyard
 flock.
- CDC reports that for the duration of the outbreak, 160 tests have been conducted for individuals with exposure to dairy cows and 400 tests have been conducted for individuals exposed to birds and other animals, resulting in a positivity rate since March 2024 of 25.00% (40/160), 5.75% (23/400) respectively.
- The United States Department of Agriculture Animal and Plant Health Inspection Service (<u>USDA APHIS</u>) has reported 151 new infected cattle herds across three states (CA, TX, MI) in the last 30 days, bringing the total for the outbreak to 923 cattle herds in 16 states.

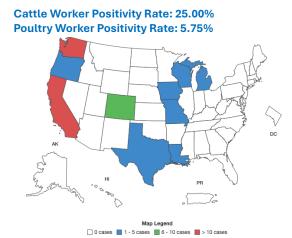


Figure 1: 2024 Map of Human H5 Infections (available from CDC)

Total Cattle Herds Affected: 923 in 16 states Total Swine Affected: 1 in 1 state In the Total Outbreak, in Livestock (all), there were: 925 Confirmed Cases in Livestock (all) by State, Total Outbreak

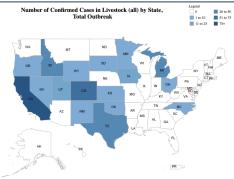


Figure 2: HPAI Confirmed Cases in All Livestock (available from USDA APHIS)

Critical Limitations

- Information about the true prevalence of live H5N1 in the raw US milk supply is greatly needed, along with clear communication efforts to individuals still consuming raw milk about the potential risks
- Information on the true prevalence and incidence of affected animals remains limited because testing is only required by USDA when moving cattle across state lines. This gap in information may be ameliorated in part by new efforts for bulk milk testing
- Testing in humans remains limited due to many factors: 1) commercial testing is unavailable, 2) testing must first be completed for seasonal flu, and then tests that are positive for influenza A but cannot be subtyped must be sent to CDC for H5N1 confirmation, 3) targeted surveillance



efforts are limited to commercial farms that are aware of the infection in their animals, and are open to health department involvement, 4) symptom reporting by affected farm workers is likely vastly underreported due to the complex relationship between farm workers and owners, and may be impacted by stigma, fear of government involvement, and concerns about missing work

CDC <u>human case numbers</u> are updated on Mondays, Wednesdays, and Fridays, while the
 <u>number of tests conducted</u> by each exposure category (poultry, dairy cows) are based on
 weekly counts, this can lead to some fluctuations in positivity rates

Recommendations

In the current scenario, it is vital to:

1. Prevent reassortment opportunities, especially during the current flu season

- Members of the general public are vaccinated against seasonal flu
- Individuals consume only pasteurized dairy products (milk, cheeses, etc.)
- Farm workers diligently use personal protective equipment (PPE; including masks, goggles, gloves, gowns, head covers, and boot covers) when working directly with or closely to cattle and poultry, other infected or potentially infected animals, and potentially infected environments
- Farm workers receive the seasonal flu vaccine as early as possible in the current flu season
- Individuals working with agricultural animals who are sick do not report to work, especially if they exhibit respiratory or flu-like symptoms, and seek medical care for diagnosis
- Individuals working with agricultural animals who are sick and unable to stay home wear a KN95 mask when in contact with animals
- Individuals planning agricultural or other events that bring together birds, cattle, and swine, should consider testing animals before exhibits or events, promoting good hygiene at events, and taking extra steps to ensure that sick animals remain at home, and animals who become sick at or recently following an event are seen by a veterinarian

2. Ensure timely, accurate surveillance and prevention of H5N1 in agricultural animals

- Increase diagnostic testing and genomic surveillance in cattle and poultry.
- Separation of infected cattle during convalescence
- Enforce cattle import restrictions to limit the movement of infected cattle across state borders
- Stringent control of potentially infected food products (removal of milk or other infected products)
- 3. Continue enhanced public health activities to prevent H5N1 transmission to and among humans



- Increase focus on sentinel surveillance, wastewater surveillance, and education of clinicians to consider H5N1 as a possible diagnosis for people who present with new respiratory illness
- Implementation of and support for recommended isolation of human cases and
 quarantine of close contacts of cases through escalated case finding and contact
 tracing, antiviral (eg, Tamiflu) prophylaxis for those exposed, compensation for
 individuals who are isolated/quarantined and cannot report to work, and social support
 to provide for essential needs of those in isolation/quarantine
- Continue development and widespread implementation of antigen and molecular testing in both hospital and outpatient healthcare settings
- Increase public health surveillance for H5N1 cases in local communities
- 4. Continue enhanced, open communication about the current situation and potential risks
 - Information sharing between the agricultural and public health sectors to increase transparency and monitor for increases in animal-to-human or human-to-human transmission
 - Enhanced communication with the public about the situation and the measures being taken to address it, as well as efforts to mitigate the spread of rumors and disinformation
- 5. Continue and consider strengthening political support for public health response
 - Policy preparation for the possibility of a pandemic, including congressional deliberations about emergency funding and emergency planning by healthcare institutions, workplaces, and federal, state, territorial, local, and tribal public health agencies
 - Increase investment and urgent development, testing, and production of vaccines and treatment options

Appendix

Scenarios

*Please note: We are evaluating the risks to human health should each scenario occur, not the relative risk of any one scenario occurring.

Features that would characterize each scenario include:

Scenario for Minimal Spread in Cattle: The virus is predominantly infecting cattle but there is minimal spread within herds and to other animals. The likelihood of widespread human infections is low. Population health consequences are low. The overall risk to human health in this scenario is low.



Scenario for Widespread Transmission in Cattle: Widespread transmission in cattle, few human infections, no human-to-human transmission. The virus is predominantly infecting cattle but spreads widely within herds. There is also occasional cow-to-human transmission. There are few human infections and no human-to-human transmission. The likelihood of widespread human infections is low. Population health consequences are low. The overall risk is low, but population-specific risk is increased for farm workers.

Scenario for Potential Human Adaptation: There is increased potential for reassortment and human adaptation, but still no human-to-human transmission. The virus begins to infect swine or other animal species which could facilitate the mixing and spreading of influenza viruses. This increases the likelihood that the virus reassorts with other influenza viruses and adapts to humans. Although the opportunities for reassortment are present, there are no specific mutations, or laboratory or epidemiological evidence to indicate that the virus has adapted for human-to-human transmission. The likelihood of widespread human infections is low. Population health consequences are low. The overall risk of widespread transmission in humans is low, but the risk is increased for farm workers. The relative risk of a future pandemic has increased, but the absolute risk remains low.

Scenario for Increased Potential for Human Adaptation and Increased Human

Reports: There is increased potential for reassortment and human adaptation, increased reports of human infections, and potential early laboratory/epidemiological/sequencing evidence for human-to-human transmission but still no human-to-human transmission confirmed. The virus has been observed in animal mixing vessels, including pigs, and additional reassortment opportunities are present, such as mixing of the H5N1 virus with the seasonal flu virus, due to the ongoing seasonal respiratory virus season (October to April), which increases the risk of human adaptation. There are more reports of human infections due to contact with infected animals like cattle, swine, and/or poultry. Viral mutations and laboratory or epidemiological evidence may be reported that indicate the potential for human-to-human transmission, but there are no confirmed reports of human-to-human transmission. Population health consequences are low. The overall risk of widespread transmission in humans is low, but risk is increased for farm workers and individuals who work with animals, and close contacts of those workers. The relative risk of a future pandemic has increased, but the absolute risk remains low.

Scenario for Limited Human Transmission: There is continued potential for reassortment, increasing reports of human infections, and limited human-to-human transmission between close contacts. There are more reports of human infections due to contact with infected animals like cattle, swine, and/or poultry. Limited human-to-human transmission is reported among close contacts of infected individuals, including healthcare workers, but there is no efficient human-to-human transmission. The likelihood of widespread human infections is moderate. Population health consequences are low. The overall risk of widespread



transmission is low, but population-specific risk is increased for farm workers, close contacts of farm workers, and healthcare workers. The likelihood of a future pandemic is increased.

Scenario for Sustained Human Transmission: There are reports of efficient human-to-human transmission. The likelihood of human infections is high because the virus now transmits efficiently and will be very difficult to contain. Population health consequences are high. Overall risk is high for all populations. The likelihood of a pandemic is very high.

H5N1 Human Health Risk Assessment Scenario Table

	Risk to farm workers	Risk to other people in contact with affected workers and animal populations	Risk to healthcare workers	Risk to the US general public
Scenario for Minimal Spread in Cattle:	Low	Low	Low	Low
The virus is predominantly infecting				
cattle but there is minimal spread within				
herds and to other animals				
Scenario for Widespread	Moderate	Low	Low	Low
Transmission in Cattle: Widespread				
transmission in cattle, few human				
infections, no human-to-human				
transmission				
Scenario for Potential Human	Moderate-	Low	Low	Low
Adaptation: Increased potential for	High			
reassortment and human adaptation,				
still no human-to-human transmission				
CURRENT- Scenario for Increased	High	Moderate	Low	Low
Potential for Human Adaptation and				
Increased Human Reports: Increased				
potential for reassortment and human				
adaptation, increased reports of human				
infections, potential early				
laboratory/epidemiological/sequencing				
evidence for human-to-human				
transmission but still no human-to-				
human transmission confirmed				
Scenario for Limited Human Transmission: Continued potential for	High	Moderate-High	Moderate	Low- Moderate



reassortment, increasing reports of				
human infections, limited human-to-				
human transmission between close				
contacts				
Scenario for Sustained Human	High	High	High	High
Transmission: Efficient human-to-				
human transmission				

References

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