HPAI A(H5) Scenario-Based Human Health Risk Assessment



HPAI A(H5) Scenario-Based Human Health Risk Assessment for the United States Center for Outbreak Response Innovation (CORI) Updated as of February 11, 2025

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 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 disease severity 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 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	High	Moderate	Low*	Low*

*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. See the next page for a detailed analysis, including limitations and recommendations.

Critical Updates:

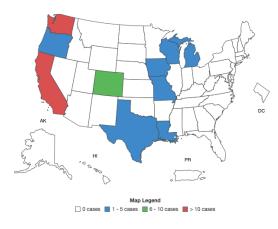
- A <u>Nevada</u> dairy worker has tested positive for H5N1. According to confidential <u>sources</u>, the individual's main symptom has been conjunctivitis. The case was first reported through the media and later confirmed by the CDC. Sequencing and symptom information has not yet been released by CDC.
- The USDA APHIS confirmed last week that the genotype of bird flu that had been primarily circulating in wild birds and poultry, D1.1, has crossed over into cattle. This genotype was first identified through bulk milk testing and is the first detection of D1.1 in cattle, indicating that H5N1 has now spilled over from birds into cattle on at least two occasions. Infected Nevada cattle were sequenced, and some samples showed the critical PB2 mutation, which improves viral replication in mammalian hosts. This mutation had not previously been observed in the B3.13 genotype in cattle or wild birds for the current outbreak but has been observed in H5 human cases in previous years with no onward transmission.
- At least three critical studies related to bird flu remain unpublished due to the pause on federal government communications. One <u>table</u> on H5N1 household transmission was inadvertently posted last week and removed. In the absence of the full report, no conclusions can be drawn.



Routine Surveillance Updates

- The Centers for Disease Control and Prevention (CDC) is reporting <u>68 confirmed human cases</u> of H5 in the United States Outbreak as of February 11, 2025 (Figure 1). One new case, in a dairy worker from Nevada, has been reported since the last publication. For the duration of the outbreak, there have been three cases that have not had a known source of infection and reported no contact with animals or raw milk, while 23 have had contact with poultry, 41 have had contact with dairy cows, and one had exposure to a backyard flock.
- CDC reports that for the duration of the outbreak, <u>171 tests have been conducted for</u> individuals with exposure to dairy cows, and 530 tests have been conducted for individuals exposed to birds and other animals, resulting in a positivity rate of 23.98% (41/171), 4.34% (23/530) respectively, since March 2024.
- Commercial and wild <u>birds</u> continue to be gravely impacted by H5N1, with 103 commercial flocks and 46 backyard flocks affected in the last 30 days.
- The United States Department of Agriculture Animal and Plant Health Inspection Service (<u>USDA</u> <u>APHIS</u>) has reported 41 new infected cattle herds from two states (CA, NV) in the last 30 days, bringing the total for the outbreak to 967 cattle herds in 16 states.

Cattle Worker Positivity Rate: 23.98%Poultry Worker Positivity Rate: 4.34%



Total Cattle Herds Affected: 967 in 16 states Total Swine Affected: 1 in 1 state Total Alpaca Affected: 1 in 1 state

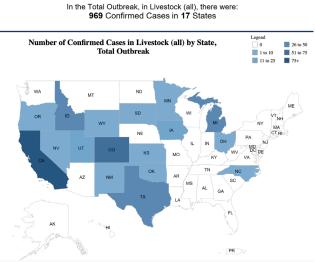


Figure 1: 2024 Map of Human H5 Infections (<u>available</u> <u>from CDC</u>).NOTE: the CDC map has not been updated to reflect the latest case in Nevada as of 2.11.25.

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

Risk Assessment HPAI A(H5) Scenario-Based Human Health Risk Assessment



Critical Limitations

- Information on the true prevalence and incidence of affected animals remains limited because <u>USDA only requires testing when moving cattle across state lines</u>. New information from bulk milk testing has proved influential, with the latest USDA APHIS <u>publication</u> on bulk milk testing providing the first detection of H5N1 genotype D1.1 in cattle and leading to the first detection of the PB2 gene, known for enhanced mammalian adaptation, in circulating H5N1.
- 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

Risk Assessment

HPAI A(H5) Scenario-Based Human Health Risk Assessment



- 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 (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 and emergency response preparation for the possibility of a pandemic, including 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

Risk Assessment HPAI A(H5) Scenario-Based Human Health Risk Assessment



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-tohuman 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 humanto-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.

	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				

H5N1 Human Health Risk Assessment Scenario Table

Risk Assessment

HPAI A(H5) Scenario-Based Human Health Risk Assessment



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	High	Moderate-High	Moderate	Low-
Transmission: Continued potential for				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

- "APHIS Confirms D1.1 Genotype in Dairy Cattle in Nevada | Animal and Plant Health Inspection Service." Accessed February 11, 2025. <u>https://www.aphis.usda.gov/news/program-update/aphis-confirms-d11-genotype-dairycattle-nevada-0.</u>
- "Bird Flu Variant Found in Nevada Cows Shows Signs of Adaptation to Mammals | CNN." Accessed February 11, 2025. <u>https://www.cnn.com/2025/02/08/health/bird-flu-variant-nevada-human-case/index.html</u>.
- Sun, Lena H. "CDC Scientific Report Resumes Publication after Unprecedented Pause." Washington Post, February 6, 2025. https://www.washingtonpost.com/health/2025/02/06/cdc-mmwr-bird-flu-public-healthcommunication/.
- US Centers for Disease Control and Prevention. How CDC is monitoring influenza data among people to better understand the current avian influenza A (H5N1) situation. Updated February 10, 2025. Accessed February 11, 2025. <u>https://www.cdc.gov/bird-flu/h5-monitoring/index.html</u>
- US Centers for Disease Control and Prevention. H5 Bird Flu: Current Situation. Updated February 10, 2025. Accessed February 11, 2025. <u>https://www.cdc.gov/bird-flu/situation-summary/index.html</u>
- US Department of Agriculture Animal and Plant Health Inspection Service. Highly Pathogenic Avian Influenza (HPAI) Detections in Livestock. Updated February 10, 2025. Accessed February 11, 2025. <u>https://www.aphis.usda.gov/livestock-poultry-</u> <u>disease/avian/avian-influenza/hpai-detections/hpai-confirmed-cases-livestock</u>
- US Department of Agriculture Animal and Plant Health Inspection Service. The Occurrence of Another Highly Pathogenic Avian Influenza (HPAI) Spillover from Wild Birds into Dairy Cattle. Published February 7, 2025. Accessed February 11, 2025.

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