Avian Influenza Virus in Bangladesh

Subjects: Zoology

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Highly pathogenic H5N1 and low pathogenic H9N2 avian influenza viruses are circulating in Bangladesh since 2007 causing serious economic losses to the country. Multiple virus introductions of different clades of HPAIV H5N1, reassorted genotypes, and on-going diversification of LPAIV H9N2 create a highly volatile virological environment which potentially implicates increased virulence, adaptation to new host species, and subsequent zoonotic transmission.

Keywords: Bangladesh, Poultry, Avian Influenza Virus

1. Definition

Influenza A viruses (IAVs), belonging to the family *Orthomyxoviridae* [1], are an important cause of respiratory infections of humans and many other species of mammals and birds. Avian influenza A viruses (AIV) are potentially zoonotic pathogens that infect a wide range of avian species and occasionally spill over into mammalian species, including humans [2][3][4]. IAVs contain a negative-sense segmented RNA genome. Their eight genome segments encode for at least ten classical influenza proteins: Polymerase basic 2 (PB2), Polymerase basic 1 (PB1), Polymerase acidic (PA), Hemagglutinin (HA), Nucleoprotein (NP), Neuraminidase (NA), Matrix 1 (M1), Matrix 2 (M2), Nonstructural 1 (NS1), and Nonstructural 2 or Nuclear Export Protein (NS2/NEP), as well as, dependent on the strain, a variable number of accessory proteins (e.g., PB1-F2, PB1-N40, PA-X, PA-N182, PA-N155) through frame-shifts and by use of complementary sequences [5][6][7]. AIVs are classified into subtypes based on their surface glycoproteins, HA and NA; there are currently 16 HA and 9 NA subtypes identified in avian species. Wild aquatic birds are the natural reservoirs of AIVs [1][9]. AIVs are further categorized by their pathogenicity in chickens into high and low pathogenicity avian influenza viruses (HPAIV). An intravenous pathogenicity index (IVPI) in chickens is used for biological pathotype characterization by experimental infection; alternatively, the sequence of an endoproteolytic cleavage site (CS) in the HA protein (HACS) can be used as a molecular marker of pathogenicity [10]. HPAI viruses exhibit high mortality in chickens and contain a polybasic HACS, which is recognized by endogenous and ubiquitous host cellular proteases, like furin, therefore predisposing these viruses to cause systemic, often lethal, infections. In contrast, LPAI viruses invoke mild respiratory illness but also run asymptomatic courses, especially in wild bird species. The LPAIV HACS consists of a mono-, di-, or tri-basic motif which restricts proteolytic cleavage activation to extracellular trypsin-like host proteases confined to the intestinal and respiratory epithelia, respectively [11]. LPAIVs are therefore incapable of inducing systemic infection.

2. Introduction

Aquatic birds are the natural hosts of IAV. Sporadically, viruses cross from aquatic wild birds to poultry or mammals, and new, adapted viruses may become established in these spill-over hosts. AIVs of subtypes H5 and H7 are the most important ones that cross to terrestrial poultry. These viruses have proven ability to mutate in poultry from low pathogenicity (LP) precursors circulating in wild birds into high pathogenicity (HP) viruses that multiply systemically in chickens, often causing very high mortality in infected flocks [12]. The factors governing such molecular mutation events are not fully understood. Therefore, infections of poultry with AIV of subtypes H5 and H7 are notifiable and require obligatory restriction measures [12][13]. The zoonotic A/goose/Guangdong/1/96 (gs/GD) lineage of H5N1 HPAI viruses, along with the G1 lineage of H9N2 and the Chinese H7N9 AIV, became well adapted to poultry and endemically circulate in many countries and in China, respectively [13][14][15][16]. Multiple clades/lineages and sub-lineages within these subtypes have been recognized, indicating ongoing evolution with significant genetic drift [17][18][19].

Worldwide, Bangladesh is among the countries with the highest number of reported HPAI outbreaks in poultry [20]. This is due to repeated incursions and endemic spread in poultry of HPAIV H5N1 of the gs/GD lineage since 2007 [21]. Sporadically, these HPAIVs are also detected in wild birds in Bangladesh [22][23]. LPAIV H9N2 was first detected in the country in 2006 and has likewise become endemic in poultry and is co-circulating in the country, together with HPAIV [24] [25]. Further AIV subtypes were isolated intermittently from domestic free-range birds and, more rarely, from aquatic wild

birds [22][23][26]. The widespread continuous co-circulation of HPAIV H5N1 and LPAIV H9N2 bears increased risks for the potential generation of new sub- and genotypes of AIVs which constitute additional obstacles to virus eradication. Both viruses cause significant economic damage in poultry production and threaten public health by their zoonotic propensity [27]. AIV surveillance studies in Bangladesh [22][26][28][29] have shown that domestic ducks play an important role in the transmission and emergence of new AIV sub- and genotypes.

3. Ecology and Epidemiology of AIVs in Bangladesh

3.1. Geographical and Ecological Frameworks

Bangladesh is a low to middle-income country in South Asia agriculturally characterized by rich water environments, paddy rice farming, and poultry production. The economy is heavily dependent on agriculture and livestock production. The country consists of a broad, deltaic plain with many tributaries and a sea coast with an extensive mangrove belt. It is at high risk of frequent flooding by three major rivers, the Ganges, Jamuna, and Brahmaputra. In addition, there is annual flooding from the seaside due to cyclones in the Bengal Bay of the Indian Ocean. Bangladesh is also an attractive and important wintering site for wild migratory birds, in particular, of the order *Anseriformes*, which breed in arctic and palearctic regions of Russia. Moreover, two major migratory bird flyways, the Central Asian and East Asian-Australian, are crossing Bangladesh [30][31][32]. The abundance of shallow coastal wetlands and vast inland wetlands (so-called haors) provide a large reservoir for wildlife, especially waterfowl, which migrate from many parts of Russia and Central Asia during winter [22][33]. This creates an ecological scenario where wild aquatic birds, domestic ducks and galliform poultry intermingle and in which pathogens, like AIV, can be easily exchanged (Figure 1). According to farmers' complains, the outbreaks are more common during the autumn and winter; however, studies were unable to identify any distinct seasonality for endemic H5N1 and H9N2 virus outbreaks in Bangladesh, and AIVs have been frequently identified from poultry in live bird markets (LBM) throughout the year [34][35][36][37]. This indicates that endemic virus circulation in poultry populations is likely the most important driver in this scenario.

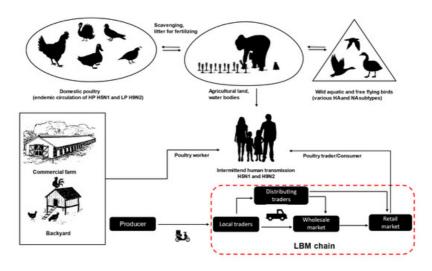


Figure 1. An overview of avian influenza virus transmission and live bird market trading chains (red dashed line) in Bangladesh.

3.2. Poultry Rearing Systems and Trading Chains

Poultry rearing in Bangladesh comprises commercial and backyard poultry production. Commercial farming can be further categorized according to farm sizes following the sectoring approach of the Food and Agriculture Organization of the United States (FAO) [38]: large-scale breeder farming, medium size farming (broiler, layer, and duck), and small size farming (layer, broiler, quail, pigeon, turkey, guinea fowl, etc.). In Bangladesh, more than 60% of the people live in rural villages [39], and it is estimated that about 90% of the rural households raise poultry (chicken, duck, pigeon) in traditional backyard settings [40]. The main poultry types reared and traded in the country include: Industrial white-feathered broiler chickens [38], Sonali (cross-breed of Rhode Island Red cocks and Fayoumi hens) [41], Deshi (back-yard chickens) [42], and ducks. Broiler, Sonali, and few ducks are raised in commercial settings, while Deshi and most of the ducks are reared in traditional scavenging systems [43][44]. Many further commercial poultry breeds, such as hybrid layer, have successfully been established in recent years in Bangladesh and are being profitably utilized by different entrepreneurs [45].

At the beginning of 21st century, the Bangladeshi poultry industry expanded rapidly. The Government of Bangladesh has given top priority to livestock development to meet the growing demand for high quality animal protein in the human diet and to create employment opportunities and generate income for the low-income rural population. As such, both industrial

poultry production and family poultry rearing are supported. Bangladesh currently raises an estimated 282 million of chickens and 55 million of ducks [46]. Improper management and biosecurity practices in poultry rearing have fostered the emergence and re-emergence of economically important infectious disease, like HPAI, leading to endemic spread of HPAIV H5N1 and LPAIV H9N2 [27][47]. During the first and second wave of HPAI H5N1 in 2007 and 2008, approximately 547 poultry farms had been affected that forced the authority to cull nearly 1.7 million birds, resulting in substantial financial losses [48]. Against the general market rules where shortage of products induces rising prices, the price of poultry meat and eggs in Bangladesh declined by 27% as a majority of the consumers desisted from consumption of potentially unsafe broiler meat and chicken eggs [48]. Finally, the poultry market collapsed, and many farm owners lost all capital.

LBMs have been incriminated in the dynamics of AIVs transmission, dissemination, and persistent circulation, thus facilitating the reassortment between different virus strains in many countries [49][50]. In LBMs, mixing of different species of birds (chickens, ducks, geese, pigeons, etc.) from different sources (wild birds, backyards, and commercial farms) creates a suitable niche for persistence and perpetuation of AIVs. In Bangladesh, live bird trading is ubiquitous, and LBMs distribute 95% of the total poultry meat and egg retails [38]. Birds are mostly traded alive because of cultural and religious preferences for consuming freshly slaughtered poultry. There is an apparent lack of processed meat marketing facilities and cold chains, particularly among rural households [51]. LBMs in rural Bangladesh often do not provide even a minimum level of biosecurity and lack proper disinfection and sanitation procedures. Before finally sold in LBMs, birds often move through a complex trading network of peripheral rural markets via several transshipment stations to wholesalers in city markets, thereby increasing risks for AIV spread and transmission to humans (Figure 1). Since 2008, several subtypes of AIVs, including, predominantly, HPAIV H5N1 and LPAIV H9N2, have been isolated from LBMs in Bangladesh [34][35][36][50]. Unhygienic slaughtering processes at LBMs bear increased risks of zoonotic transmissions [34][50], but actual virus transmission to LBM workers as evidenced by seroconversion was scarce [52]. Yet, LBMs remain a very important target to understand and intercept the local circulation of AIVs in domestic poultry in Bangladesh and to identify and combat risk factors in zoonotic transmission. The outbreak frequency of endemic HPAI H5N1 and LPAI H9N2 has increased due to increased poultry production, characterized by a mélange of various, highly fragmented rearing systems and marketing chains [53], as elucidated above.

3.3. Virus Transmission and Risk Factors

Droplet, aerosol, faeco-oral, and direct or indirect contact with contaminated materials are widely described modes of transmission of AIVs [54]. Transmission risks increase depending on host susceptibility and viral load in the environment, as well as on distance between and frequency of contacts [55]. In Bangladesh, the domestic duck is currently considered as the most important epidemiological factor associated with AIV transmission between wild birds and other poultry [22][56]. Low awareness among the raisers of backyard poultry of the zoonotic properties of AIV, neglected practice of biosecurity measures, and the close living arrangements of poultry and rural human populations lodge them at the highest risk for zoonotic transmission [27]. HPAI H5N1 and LPAI H9N2 outbreaks cause enormous losses to small scale poultry producers [57]. Safe disposal of litter and fallen animals pose further severe problems. Untreated poultry litters are being used as fertilizer on agricultural lands and as fish feed in water bodies, which may contaminate the environment and further trigger viral spreading [58]. However, large-scale commercial farms are relatively better managed and follow biosecurity recommendations, which reduces huge financial losses due to Al outbreaks [27]. Moreover, LBMs provide foraging opportunities for peri-domestic birds, such as crows, sparrows, and starlings [59]. Consequently, house crows have been found positive for HPAIV H5N1 on several occasions [60]. Moreover, no closure days for LBMs with H5 AIV-positive birds are usually decreed. Possible risk factors stratified according to poultry rearing and marketing systems in Bangladesh are listed in Table 1. Effective human-human transmission of HPAI or LPAI virus is not evident in the country, but avianhuman transmission follows the close proximity of each other [27][50].

Table 1. Risk factors identified in the Bangladeshi poultry farming and marketing systems that may promote spread of avian influenza virus (AIV).

Sectors	Possible Risk Factors	References
Backyard poultry	 Frequent close contact with neighboring poultry flocks due to free scavenging 	
	Co-rearing of different species (chicken, duck, parrot, pigeon)	
	 Handling and slaughtering of sick birds, including putting sick poultry together 	
	Uncontrolled movement and traffic	[27][40][43][61][62][63][64] [65][66][67]
	 Lack of regular cleaning, safe disposal and disinfection, including feeding of slaughter remnants of purchased chickens to backyard chickens 	
	 Ubiquitous water bodies, rice paddies, and bushland fostering contact with wild bird populations 	
	Commercial poultry	
Small and medium enterprises	Farm location near public places and high density of farms	
	 Lack of isolation facilities and improper disposal of sick or dead birds 	
	 Lack of biosecurity (fencing and improper using of footbaths) leading to contact with free flying wild and backyard birds and other animals 	[21][39][43][57][68][69]
	Exchange of farm equipment	
	Uncontrolled vehicle movement	
	Ubiquitous water bodies, rice paddies, and bushland	
	Mixed origin of restocking poultry	
Large holding (specifically, commercial layer farms)	Limited biosecurity, including	
	-Limited control of vehicle movementsExtreme environmental -Improper management of entry of farm personnelconditions (heat and -Failure to impose unauthorized movementcold stress, humidity) -Exchange of farm equipment -Roaming of village chickens around the farmVaccination failures -Unsafe sources of litter/bedding materials	<u>[21][70]</u>
Live bird markets (LBM)	Low level of biosecurity, including	
	-Lack of market infrastructure Taking back poultry at the end of -Inadequate hygienic measures: sanitation and disinfectionmarket days -No use of personal protection equipment -Lack of awareness about AIV transmission	[21][27][39][57][59][61][71] [72]
	-Keeping chickens and ducks together in the stalls -Exchange of egg-trays and vehicles between LBMs and farms	

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