AVIAN INFLUENZA-A CURRENT PERSPECTIVE

PRESTAGE DEPT. OF POULTRY SCIENCE

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ECONOMIC IMPACT OF HPAI

• COSTS INCURRED FROM:

- BORDER CONTROL AND PERMITTING TO EXCLUDE AI.
- PREPAREDNESS, TRAINING, INFRASTRUCTURE, R & D.
- OUTBREAK CONTROL AND ERADICATION GOVERNMENT SECTOR PRIVATE SECTOR CONSUMERS
- DISRUPTION IN TRADE OF BREEDING STOCK, CHICKS AND PRODUCTS

FINANCIAL IMPACT OF HPAI

- 1924 U.S.
- 1983 PA.

• 1999 ITALY

- \$ 10 m (2000 VALUE)
- \$ 110m (USDA-APHIS)
- \$ 25m (PRODUCERS)
- \$ 350m (CONSUMERS)
- **\$ 600m**
- 1985 Australia \$ 2m (SINGLE COMPLEX)

HISTORY OF HPAI

- 1878 and 1894 Italy and other European countries (where recognized)
- 1901 Germany and neighboring nations, Chickens "Fowl plague"
- 1924 and 1929 U.S., Chickens (H7)
- 1959 Scotland, Chickens H5N1
- 1961 South Africa, Terns H5N3
- 1976 Australia Chickens H7N7
- 1983 U.S. (PA, MD) Chickens , Turkeys H5N2
- 1995 Pakistan, Chickens H7N3
- 1997 Hong Kong, Chickens etc. H5N1
- 1999 Italy, Chickens etc H7N1
- 2001 China and other Asian nations, Chickens etc H5N1
- 2003 Mexico, Chickens etc. H5N2

RECENT HPAI OUTBREAKS

- 2003 Holland, Chickens H7N7
- 2012 Mexico, Chickens etc. H7N3
- 2013 Australia, Chickens H7N2
- 2014 Canada, (BC) Chickens etc. H5N2
- 2014 Russia, Chickens etc. H5N1
- 2014 India, Ducks and Chickens H5N8
- 2014 Korea, Japan, Ducks and chickens H5N8
- 2014 EU, Chickens, Turkeys and Ducks H5N8
- 2014 Taiwan, Ducks, Geese, Chickens H5N2 and H5N8 plus H5N3
- 2014 Canada, Chickens and Turkeys H5N2
- 2014/5 U.S. (OR, UT, WA, ID), wild birds, backyard flocks H5N2 & H5N8, H5N1
- 2015 Nigeria, Chickens H5N?
- 2015 Israel, Palestine, Bulgaria, Turkeys and Chickens, H5N1

HUMAN HEALTH SIGNIFICANCE

- MAMMALS GENERALLY REFRACTORY TO AVIAN STRAINS
- Asia H5N1 from 2003 onwards
- Holland H7N7 in 2003
- China H7N9 2013 onwards
- Egypt H5N1 mid 2000's onwards
- Mild cases (conjunctivitis)
- Severe cases (respiratory complications and death)
- Marked decline in egg and chicken consumption

CLOSE CONTACT WITH INFECTED FLOCKS RESULTSED IN H5N1 INFECTION IN GENETICALLY PREDISPOSED HUMANS



MOST OF THE 500 CASES OF HUMAN H5N9 AI INFECTION HAVE BEEN DOCUMENTED IN CHINA WITH HIGH FATALITY RATES



AVIAN INFLUENZA VIRUS

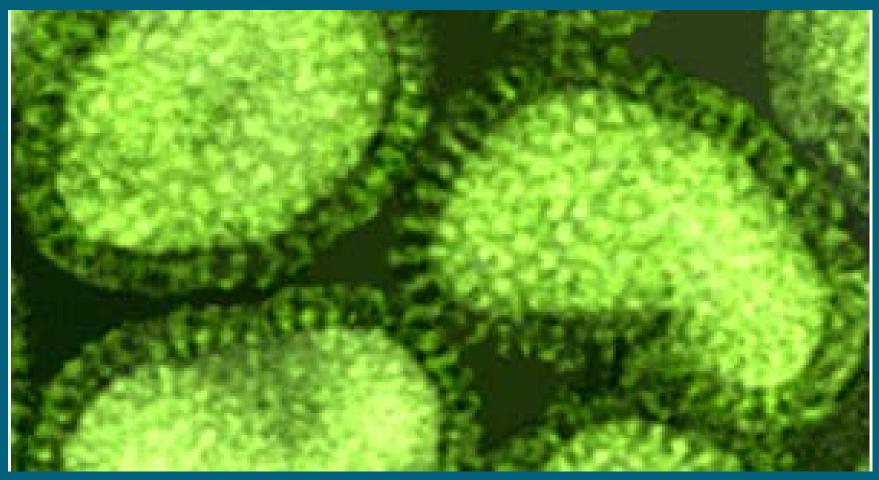
- Family: Orthomyxoviridae
- Genus: Influenzavirus A
- RNA virus with enveloped virion, 80-120 nm
- Eight single RNA strands coding for:-
- ✤ 1. PB 1 transcriptase
- **♦ 2. PB 2** endonuclease
- * 3. PA **RNA replication**
- 🚸 4. HA attachment, envelope fusion, neutralization
- * 5. NP **vRNA** synthesis
- ✤ 6. NA

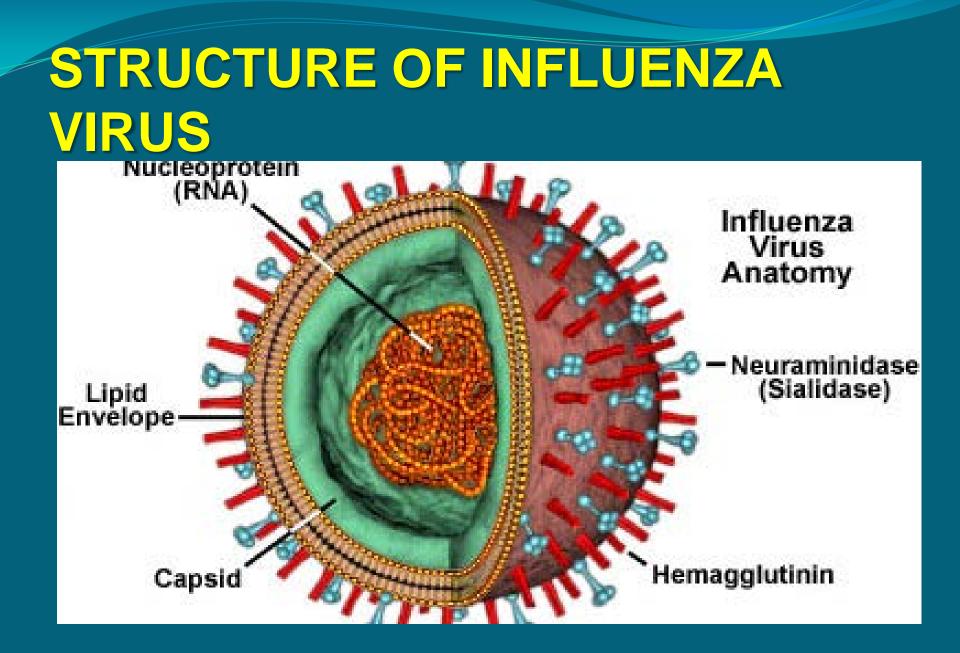
virus elution,

✤ 7. M 1 & 2 virus budding

✤ 8. NS 1 & NS 2 translation of viral mRNA

EM VIEW OF AI ORTHOMYXOVIRUS SHOWING SURFACE HA AND NA GLYCOPROTEIN ANTIGENS





AVIAN INFLUENZA NOMENCLATURE

A / chicken/ Indonesia/ 4 / H7N8/09
A/ chicken /Pennsylvania/ 1370/83
15 HA (hemagglutinin) serotypes
9 NA (neuraminidase) serotypes
"Highly pathogenic avian influenza" H5 and H7
"Low pathogenic avian influenza" 13 other HAs

ANTIGENIC VARIATION

 Surface HA and NA glycoproteins undergo frequent changes.

Antigenic Drift

- Arises by point mutation. (vaccination pressure and
 - population density?)
- H5 and H7 strains of LPAI becoming HPAI
- Antigenic Shift
- Arises from genetic reassortment

SIGNIFICANCE OF VIRAL SHIFT REASSORTMENTS

- A/goose/Taiwan/??/2015 H5N3
- Identified mid-January 2015
- H5 99% similar to 2014 H5N8 isolate involved in extensive outbreaks in S.Korea and Japan.
- N3 98% similar to 2010 H2N3 Taiwan isolate migratory ducks
- 2011 H1N3 Thailand isolate in waterfowl
- 2013 H5N3 Taiwan isolate migratory ducks

OIE CRITERIA FOR HPAI

- 1. Al isolate lethal to +6/8 5-week SPF chickens receiving 0.2 ml 10⁻¹ allantoic fluid iv.
- 2. Any H5 or H7 isolate with a preponderance of basic amino acids at the HA cleavage site.
- 3. Any isolate other than an H5 or H7 lethal to 1 to 5 chickens and can be grown in cell culture without trypsin

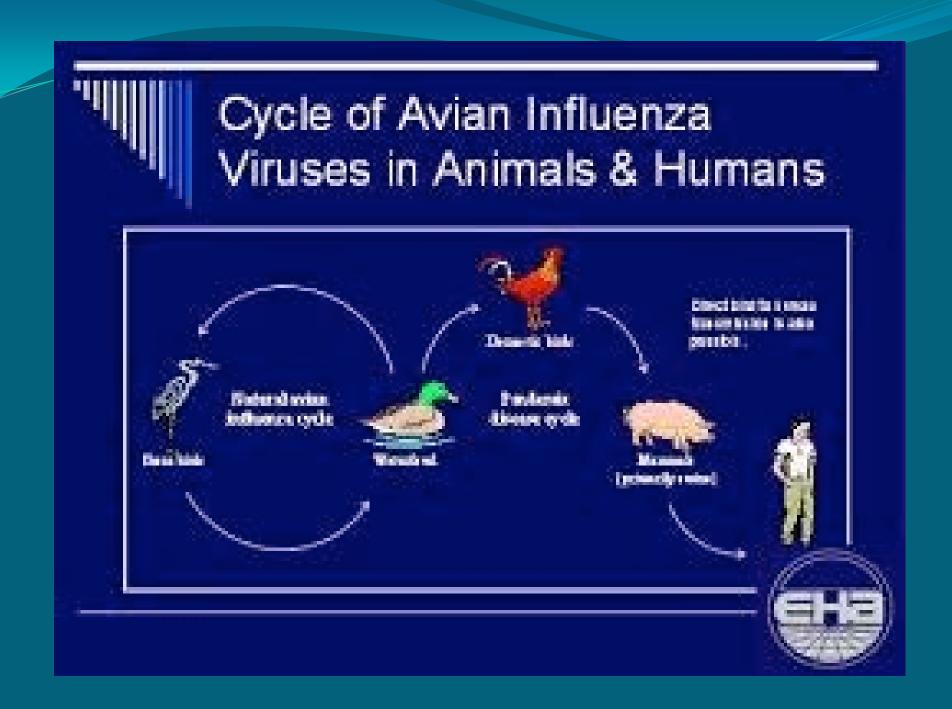
SENSITIVITY OF AI VIRUS

Inactivated by:-

- solvents and detergents
- aldehydes (formalin and gluteraldehyde)*
- oxidizing agents (sodium hypochlorite 5%)*
- chemical disinfectants (phenolics, QACs)*
- * ONLY if not protected by organic matter.
- Al virus can persist in liquid manure for 100 days in NE U.S winter

in feces for 30 days at 4° F

Susceptible to 90° F in "cleaned" houses for 1 week



ORIGIN OF AI VIRUS STRAINS

- Role of Asia in evolution of AI strains
- Migratory waterfowl to domestic waterfowl
- Spread to chickens and mixing in hogs and other mammalian hosts in rural locations.
 Reassortment events ("shifts") occur to produce pandemic strains infecting humans. Can be induced under laboratory conditions.
- Endemic infection can lead to mutations ("drift") when introduced into areas with high population density-LPAI to HPAI.

COHABITATION OF DOMESTIC AND MIGRATORY WATERFOWL RESULTS IN TRANSMISSION AND DISSEMINATION OF HPAI



DISSEMINATION OF AI

- Intercontinental and international:
- Migratory waterfowl and shore birds
- Uncooked poultry products
- Contaminated personnel
- Regional and local
- Movement of live poultry (LBM systems)
- Movement of contaminated personnel and
- equipment (fomites)
- Virus entrained on dust particles in air.
- Contaminated housing and waste

LIVE BIRD MARKETS IN ASIA ARE ARE A SOURCE OF AI VIRUS FOR CONSUMERS AND A RESERVOIR FOR POULTRY FLOCKS



CLINICAL PRESENTATION OF HPAI

- SHARP DROP IN WATER AND THEN FEED INTAKE
- CONCURRENT RAPID ASCENDING MORBIDITY
- RATE (10%; 40 % 80% CUMULATIVE ON SUCCESSIVE DAYS)
- LAYERS AND BREEDERS CEASE PRODUCTION (OVER 2 TO 3 DAYS, PRESENCE OF SHELL-LESS EGGS)
- SIMULTANEOUS RAPID ASCENT IN MORTALITY RATE (5%; 25%; 50% CUMULATIVE OVER SUCCESSIVE DAYS)
- TYPICAL SIGNS (PROSTRATION, RESPIRATORY DISTRESS, DIARRHEA, SWOLLEN CYANOTIC HEADS, SKIN HEMORRHAGES)

EXTENSIVE MORBIDITY IN FLOCK INFECTED WITH HPAI. RECUMBENCY, RESPIRATORY DISTRESS EVIDENT IN SMALL FLOCK



RAPIDLY ASCENDING MORTALITY FOLLOWING HPAI INFECTION LEADS TO A "CARPET OF DEAD BIRDS"



CUTANEOUS LESIONS HPAI



SWOLLEN WATTLES, NECROSIS OF COMB WITH HPAI



CHARACTERISTIC SUBCUTANEOUS HEMORRHAGES ON THE SHANKS OCCUR FOLLOWING HPAI INFECTION



LESIONS OF HPAI

- EDEMATOUS, HEMORRHAGIC AND NECROTIC CHANGES IN ADNEXA (skin, wattles, comb, shanks)
- VISCERAL SEROSAL HEMORRHAGES
- FIBRINOUS PERITONITIS (NON-PERACUTE CASES)
- NOTE: GROSS LESIONS ARE NOT PATHOGNOMONIC. D/D INCLUDES vvND (END),
 COMBINATIONS OF vvIBD/ILT/LENTOGENIC ND

DIAGNOSIS OF HPAI

- VIRAL ISOLATION:
- SPF CHICKEN EMBRYOS ALLANTOIC ROUTE WITH HI
- ANTIGEN-CAPTURE ASSAY (DIRECTIGEN®)
- VIRAL RNA:
- PCR ASSAY
- SEROLOGY:
- HEMAGGLUTINATION INHIBITION
- AGAR-GEL WELL AGGLUTINATION
- ELISA

ELISA AI ANTIBODY TEST KITS AVAILABLE COMMERCIALLY

Avian Influenza Virus Antibody Test Kit

in Piter Bargenetik für Denselsen 19 konnt Antibade in Anlan Anfansen Verse (1999) Sam for under 2 Novel (Sami un Antiba v Novel oppik) 21 sementing 22 1999) An enteren in sought pression 2 Ny Samine, 29 Sai Piter 2 Piter 2 Piter 2 Aug.

CHICKEN or

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PREVENTION AND CONTROL OF HPAI

• EXOTIC TO NATION

EXCLUSION BY RESTRICTING IMPORTS RAPID DIAGNOSIS AND ERADICATION

(QUARANTINE, COMPENSATION, SURVEILLANCE, FLOCK DEPLETION AND DISPOSAL) REGIONAL CONTROL OF MOVEMENT (REGIONALIZATION AND COMPARTMENTALIZATION)

FARM BIOSECURITY

ENDEMIC TO NATION

REGIONALIZE AND RESTRICT INTERZONE MOVEMENT VACCINATE, VACCINATE, VACCINATE BIOSECURITY AS FAR AS PRACTICAL

MASS DEPLETION OF FLOCKS IMPOSES RISKS OF INFECTION FOR WORKERS AND ALSO PROBLEMS OF DISPOSAL



VACCINATION AGAINST HPAI

- INTRODUCTION OF AN EXOTIC INFECTION
- VACCINATION NOT RECOMMENDED.
- (PRESENCE OF ANTIBODIES PRECLUDE TRADE)
- PRESENCE OF ENDEMIC INFECTION
- INACTIVATED EMULSIONS (STABLE, HIGH ANTIGEN TITER)
- DIVA APPLIED IN ITALY H7N3 VACCINE AGAINST H7N1
- •
- **RECOMBINANT rHVT H5 PRODUCT DEVELOPED (CEVA)**
- NO LIVE ATTENUATED VACCINES!!!
- \bullet

I/M VACCINATION OF CHICKS WITH INACTIVATED EMULSION SUPPRESSES MORTALITY BUT DOES NOT ERADICATE INFECTION. LOW SHED RATE OF VIRUS FOLLOWING VACCINATION.



VACCINATION OF INDIVIDUAL HENS IN THE FACE OF INFECTION IS LABORIOUS



TRADE CONSIDERATIONS

- TO REGULATE TRADE, THE WTO (world trade organization) RECOGNIZES STANDARDS DEVELOPED BY THE OIE (world organization for animal health).
- CANNOT BAN TRADE IF SAME DISEASE OCCURS IN IMPORTING NATION
- •
- ALL H5 AND H7 AND HIGH-PATH ISOLATES TO BE REPORTED TO OIE 1 DAY FOLLOWING CONFIRMATION. OTHER AI ISOLATES AT ANNUAL INTERVALS.
 BILATERAL AGREEMENNTS EXIST BETWEEN NATIONS MANDATING DISCLOSURE OF LPAI
- SOME NATIONS (RUSSIA) USE AI AS AN EXCUSE FOR POLITICAL EMBARGOS
- MANY NATIONS (INDIA) USE AI TO PROTECT LOCAL PRODUCTION

QUESTIONS TO PONDER 1. CAN I APPLY THE BASICS OF AI VIRUS MOLECULAR BIOLOGY TO AN UNDERSTANDING OF THE DISEASE? 2. DO I UNDESTAND THE EPIDEMIOLOGY OF AI AND THE SIGNIFICANCE IN PREVENTION AND CONTROL?

3. AM I AWARE OF THE SOCIOECONOMIC IMPACT OF AI?

4. COULD I DEVELOP A PROGRAM TO ERADICATE EXOTIC AI?

5. COULD I APPLY THE PRINCIPLES ACQUIRED IN THIS REVIEW TO LIMIT INTRODUCTION AND DISSEMINATION OF AI?

THANK YOU

QUESTIONS?

COMMENTS!