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Viral respiratory infections and their role as public health problem in tropical countries (Review)

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Summary

Acute respiratory infections (ARI) are a major cause of morbidity and mortality throughout the world. Data from the World Health Organization indicate that there are at least 2.2 million deaths from ARI throughout the world each year. A considerable number of study have been performed in different countries to assess the etiological role of viruses in ARI and is now clear that the majority of infections of the respiratory tract are caused by viruses.

In tropical countries information on the viral etiological agents of ARI is rather scanty. Nevertheless data from Papua New Guinea, Polynesia, India, Pakistan and Singapor show that influenza occurs frequently in tropical countries. The other respiratory viruses especially respiratory syncytial virus, parainfluenza viruses and adenoviruses also play significant role as etiological agents in many tropical countries as Panama, Jamaica, Brazil, Colombia, Trinidad, Uganda, India and Nigeria. The data concerning seasonal prevalence of viral ARI in tropics are contradictory.

Résumé

Les infections respiratoires aigües (IRA) sont la cause majeure de la morbidité et de la mortalité à travers le monde. Des données de l'Organisation Mondiale de la Santé, il résulte qu'il y a au moins 2,2 millions de décès causés chaque année, par les IRA, à travers le monde.

D'importantes études ont été menées dans différents pays pour évaluer le rôle étiologique des virus dans les IRA, et aujourd'hui, il est clair que la majorité des infections des voies respiratoires, est causée par des virus.

Dans les pays tropicaux, les informations sur les agents étiologiques, sont peu abondantes. Toutefois,

des données en provenance de Papouasie Nouvelle Guinée, Polynésie, Inde, Pakistan et Singapour, montrent que la grippe frappe souvent dans les régions tropicales.

Les autres virus respiratoires, notamment: le virus respiratoire syncytial, les virus paragrippaux et les adénovirus jouent aussi un rôle important tout comme les agents étiologiques, dans de nombreux pays tropicaux tels que la Panama, la JamaIque, le Brésil, la Colombie, Trinidad, l'Ouganda, L'Inde et le Nigéria. Les données relatives à la fréquence saisonnière des IRA dans les zônes tropicales, sont contradictoires.

Introduction

Acute respiratory infections (ARI) are a major cause of morbidity and mortality throughout the world. ARI include infections of viral and bacterial origin, such as the common cold, pharyngitis, laryngitis, tracheitis, bronchitis, pneumonia and bronchopneumonia. Data from the World Health Organization (WHO) indicate that there are at least 2.2 millions deaths from ARI throughout the world each year[1,2]. In developed countries among infants and those over 65 years of age, such infections are responsible for 10-15% of deaths. For developing countries, available data indicate that although the morbidity from ARI is roughly the same as in developed countries, the associated mortality may be 30 or more times higher[3], however, for most developing countries data of mortality and causes of death are not available.

Most data on ARI come from studies conducted in temperate areas. The little information that is available shows a leading role of ARI in the tropics and subtropics. In a review of infectious diseases in Angola[4] influenza was found to be the third major cause of illness, preceded by malaria and "acute

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intestinal disease of unknown etiology"

About one-third of all paediatric admissions to hospitals in Panama City (Panama) are due to AR diseases[5]. From 1970 to 1979 at Goroka Base Hospital, in Papua New Guinea, pneumonia was responsible for 29% of the paediatric admissions, and for 31% of the deaths of children in the hospital[6].

Information concerning the situation in Nigeria shows a similar picture. Data from University College Hospital, Ibadan show that 10.4% of all admissions for one year period are patients with respiratory diseases excluding tuborculosis[7]. Osubor and Etta[8] carried out a morbidity survey in Samaru, Zaria, among 399 children under 5 years of age. The 3 commonest ailments were respiratory diseases - 25.1%, protein-calorie-malnutrition -21.1%, and gastroenteritis - 15%. The majority of the infections were in infants - 57%, while only 8% were aged 3-5 years. Obi[9] analysed paediatric medical cases admitted to the Children's Clinic, Benin City. The commonest infections were enteritis - 23%, respiratory infections - 23.3%, and malaria - 21.7%. Warrel[10] reported lobar pneumonia as the commonest cause of hospital admissions in Zaria. The analysis of medical admissions to paediatric ward of the General Hospital Lagos[11] during one year period shows that the largest group, accounting for 20.9% of all admissions and 19.3% of all deaths, suffered from diseases of the respiratory tract, excluding tuberculosis and measles bronchopneumonia. A study carried out in Adeoyo Hospital, Ibadan[12] shows that 23% of total admissions of the children under the age of 10 years were patients with lower respiratory tract infections (LRTI), with mortality rate 32.2%.

Odiase[13] investigated the leading causes of death among in-patients of the University of Benin Teaching Hospital in the year 1974, pneumonia was into the 10 diseases responsible for 73.3% of the deaths. Sofowora and Onadeko[14] reported a death rate of 20% from Nigerian patients with pneumonia. In another study[15] the mortality due to respiratory infections in Nigerian children was found to be 19% of all admissions in Lagos.

Etiological agents of viral respiratory infections

Many studies have been carried out to establish the relative importance of the microbial agents involved in respiratory tract infections. The 3 major types of respiratory pathogens are bacteria, viruses and mycoplasmas. Although bacteria can produce significant diseases of the respiratory tract, it has been possible to establish them as causative agents in only small proportion of such illnesses. A number of studies have been performed in different countries[16,17,18,19,20,21,22] assess the to etiological role of viruses in ARI. The majority of infections of the respiratory tract are caused by viruses. Almost all studies support this premise[23,24]. Viral agents have been estimated to be responsible for over 90% of cases of upper respiratory tract infections (URTI), and a considerable, if lesser proportion of cases of LRTI[25]. Respiratory syncytial virus (RSV). parainfluenza (PI) viruses, adenoviruses and influenza viruses have been found to be the most important agents[26,27,28,29]. Rhinoviruses have also been found to be important pathogens although they normally cause mild diseases[30,31,32]. The role of coronaviruses and reoviruses is not so important. Generally they cause URTI[33,34,35].

Respiratory viruses in the tropics

Most of our knowledge on the etiological agents of viral ARI is derived from observations in the temperate zone, where viral infections predominate. In tropical countries, information on the viral etiological agents of ARI is rather scanty, because of the lack of laboratory facilities and epidemiological data. It is of interest to compare various aspects of viral ARI in temperate areas and the tropics.

Incidence of viral ARI

Data from Papua New Guinea[36], Polynesy[37], India[38,39], and others show that influenza occurs frequently in tropical countries where the prevalence of individual virus variants is not substantially different from that in the countries with temperate climates[40]. Serological survey of influenza in Pakistan during period 1976-1980[41] established that 36% of the population had antibodies to influenza B virus and 89% to influenza. A virus.

Influenza surveillance in Singapore, during 1972-1986[42] shows that influenza is one of the major viral diseases on the island of Singapore and outbreaks due to influenza A virus occurred every year and to influenza B virus at intervals of 16 to 24 months.

During the course of continuous surveillance of

influenza in 1980 in Pune, India[43] three outbreaks of ARI were noted. The first and second outbreaks were associated with influenza A and B viruses.

In a study carried out in Ahmadu Bello University Teaching Hospital, Zaria[44] sera from 13 adult patients with lobar pneumonia were tested for antibody to several respiratory viruses. Influenza A infection was confirmed in 3 patients (23%).

The other respiratory viruses also play significant role as etiological agents of ARI in the tropics[45,46,47]. In a serological survey in the Caribbean Island of Jamaica[48] during a 30 month study period, specimens from 552 patients of all age groups, suffering from some from of respiratory illness were examined. Serologic findings indicated influenza A virus in 5.7% of the cases, influenza B virus -3.2%, influenza C virus -2%, PI virus type 1 -8.3%, PI virus type 3 -7.8%, adenoviruses -4.1% and RSV -8.6%.

During a 4 year period of observation ARI was the commonest cause of infirmary admission among students at the University of Philippines -64.2%[49]. In 27.9% of cases with ARI tested, virus was isolated.

A representative investigation of ARI was carried out in Panama Canal Zone during a 4 year period[50]. The age-specific prevalence of antibodies against a number of respiratory viruses was examined. Generally the curves of prevalence were similar to those observed with the appropriate antigens in Germany[51], United-States[29,52,53], Bulgaria[32] and USSR[54].

During the same investigation carried out in Panama Canal Zone, serologic evidence of viral etiology was demonstrated for RSV in 14.7%, PI virus type 1 -8%, PI virus type 2 -0.7%, PI virus type 3 -8%, adenoviruses -4%. These results are comparable to that observed in Bulgaria in 1984[55] where antibody responses to RSV were also detected more frequently (10.3%) than to adenoviruses -6.3%, PI virus type 1 - 3.3%, PI virus type 2 - 1.9% and PI virus type 3 - 2.9%. An increasing number of studies from tropical countries show the similarity in the incidence and the pattern of viral respiratory diseases between countries with temperate climates and those with warm climates[45].

Viral ARI in childhood

The respiratory viruses were said to have an important role particularly in childhood pathology[19,56,57]. However most of the information originates from studies carried out in countries with temperate climates. In many tropical countries little is known on the subject. Nevertheless the last twenty years has provided some data. In a study organised by WHO[58], paired sera from 528 children up to 5 years old admitted to hospital with severe respiratory illness were collected in 10 tropical countries and were tested by complement fixation test. In most of the countries the pattern of infection was similar. RSV was the most important respiratory tract pathogen of early life (19%), particularly in the first year of life and in cases of bronchiolitis and pneumonia. The PI viruses were next in importance (16%) particularly in cases of croup, but they were older children (1-2 years). commoner in Adenoviruses (6%), influenza A virus (4%) and influenza B virus (2%) were of moderate importance. This pattern is similar to that which has been observed in temperate climates[19,56,57].

A 2-year study was undertaken to establish the incidence and possible viral etiology of ARI among the child population in Rio de Janeiro, Brazil[59]. Viruses were isolated from 20.4% of the throat swabs collected. Of the viruses identified, 47% were adenoviruses, 9% were influenza A virus, 7% - PI viruses, 3% - RSV, and 1% - influenza B virus.

A prospective study of acute LRTI in children under 15 years old was undertaken in Cali, Colombia[60]. A viral agent was identified in 29% of the patients with croup, 20% with tracheobronchitis, 22% with bronchiolitis and 17% with pneumonia. RSV was found in 9% of the cases being the commonest viral pathogen in young infants with bronchiolitis and pneumonia. The other 3 viruses, with which RSV accounted for 94% of all virus identification, were adenoviruses, PI viruses types 1, 2, and 3 and influenza A virus.

Similar results were obtained during an investigation of patients with ARI in Trinidad[61]. 94 cases were found to have RSV infection. Eighty-four per cent of them were less than three years of age.

In a study from Uganda[62] the important role of viruses in the etiology of ARI in infants and young children were confirmed. More than one-third of the cases of respiratory tract infection were of viral origin. The most important of the viruses were RSV (17%) and PI viruses (9%). Adenoviruses were found to be less important and were etiologically related to only 4% of respiratory disease cases.

A study carried out in Kuala Lumpur,

Malaysia[63] showed that in 28.9% of the cases with ARI, viruses were isolated. The most common viruses isolated were RSV and rhinoviruses. Other viruses isolated were PI and influenza A viruses. Mycoplasma Pneumoniae and respiratory viruses are common precipitants of wheezing episodes in asthmatic children in Singapore[64], where PI virus infection was accounting for 9.9% of admissions, followed by the RSV - 9.1%. Influenza A virus and rhinoviruses were found in 3.3% of admissions each and adenoviruses in 2.5% or viruses were accounting for 28.1% of admissions. Mycoplasma Pneumoniae infection was associated with 16.5% of admissions. In only 5% of children wheezing episodes were related to bacterial infection.

The data among Nigerian children, although scanty, also confirm the role of respiratory viruses. Ogunbi[65] reported a serological study of 103 children with LRTI seen at the Lagos University Teaching Hospital. In 53 cases of bronchiolitis 11.3% were positive for RSV. Adenoviruses, PI viruses and influenza viruses were found in 5.6% of cases each. Also in Lagos Njoku-Obi and Ogunbi[66] observed that 32% of children aged between 1-5 years had RSV complement fixing antibodies.

Several investigations, concerning etiological role of RSV have been made among hospitalised children in Benin City. Dym, Schuit, Nwankwo and Omene[67] examined 80 nasopharyngeal secretions obtained from patients between 6 weeks and 3 years of age admitted with pneumonia or bronchiolitis. Fifty-four per cent of them had a positive ELISA for RSV. Okuonghae[68] studied the nasocomial spread of RSV infection in a newborn nursery in Benin during a 12 week period. A total of 56 babies were studied. 11 babies (20%) acquired the infection nasocomially. The infected babies were all symptomatic and some had significant morbidity.

The results of these studies corroborate the findings of studies carried out in temperate areas of the world that have identified RSV, PI viruses, adenoviruses and influenza viruses as the most frequent viral agents associated with respiratory tract infections in childhood.

Seasonal variations

In temperate climates RSV and influenza viruses cause yearly outbreaks of infection usually during the winter months, whereas PI virus types 1 and 2 characteristically produce epidemics in the fall of every other year. Infections with rhinoviruses, coronaviruses, reoviruses, adenoviruses and PI virus type 3 are endemic, occurring in all seasons of the year. The variation from season to season in the incidence of respiratory disease in the temperate zone suggests that the frequency and distribution of such infections might be different in regions with warm climates year-round. This idea has, from time to time, stimulated investigations into the role of respiratory pathogens in the tropics, but much of the work has been limited to the demonstration of the presence of a single agent[42,45,47,69,70].

According to a study from Rio de Janeiro[59], influenza A viruses were circulating in the late autumn and winter (dry season). Influenza outbreaks in Pune, India are reported to occur in the hot season (March) and rainy season (July-September) [38,43]. But data from Nigeria[70] show that the influenza outbreak in 1974 coincided with the onset of the harmattan season (November). Whereas in a study of paired sera collected in several tropical countries[58] and in a study from Uganda[62] no marked seasonal distribution of influenza virus infections was observed.

PI viruses and adenoviruses were observed to be causing infections at all times of the year [9,58,60,61,62]. Inspite of relatively more observations concerning RSV infection the seasonal pattern of RSV activity in a tropical climate remains unclear. Studies in Trinidad[61], Phillipines[49], Singapour[71] have reported that RSV activity occurs in sharply defined outbreaks during the rainy season of the year. Other projects have failed to demonstrate this pattern consistently[6,59,60,62,72].

Conclusion

Despite the great attention given to investigations of viral respiratory infections in recent years, the available information on their etiology in tropical areas remains incomplete. There are no information about the etiological role of coronaviruses and reoviruses in ARI. The data concerning seasonal prevalence of viral ARI are incomplete and contradictory. In many studies the number of patients investigated were insufficient to allow reliable conclusions to be drawn. Nevertheless the data presented in this review have draw attention to the seriousness of viral respiratory infections in tropical countries. This information indicates the need for long-term observations in differing epidemiological settings to provide a rational basis for prophylaxis, diagnosis and therapy of viral ARI.

References

- Bulla A, Hitze K. Acute respiratory infections: A review. Bull. WHO. 1978; 56: 481-498.
- Pringle CR. Progress towards control of the acute respiratory viral diseases of childhood. Bull. WHO. 1987; 65: 133-137.
- WHO. Global mortality from acute respiratory infections among children aged below 5 years. Bull. WHO. 1987; 65: 114-116.
- Khat'ko NI. Epidemiological characteristics of Angola. J. Microb. Epidem. Immunol. 1985; 2: 81-88.
- Reeves WC, Dillman L, Quiroz, E, et al. Epidemiology of acute respiratory disease at the Pediatric Emergency room of the Social Security Medical Center in Panama City, Panama. Bull. PanAmer. Health Organ. 1985; 19: 221-234.
- Shann F, Hart K, Thomas D. Acute lower respiratory tract infections in children: Possible criteria for selection of patients for antibiotic therapy and hospital admission. Bull. WHO. 1984; 62: 749-753.
- Lauckner JR, Rankin AM, Adi FC. Analysis of medical admissions to University College Hospital, Ibadan — 1958. W. Afr. Med. J. 1961; 10: 3-32.
- Osuhor PC, Etta KM. Morbidity patterns amongst children in a semi-urban community in northern Nigeria. J. Trop. Pediatr. 1980; 26: 99-103.
- Obi JO. Analysis of pediatric medical cases admitted to children's clinic, Benin City. Nig. Med. J. 1976; 6: 69-73.
- Warrell DA. Respiratory tract infections in the tropics. The Practitioner 1975; 215: 740-746.
- Gans B. Paediatric problems in Lagos. W. Afr. Med. J. 1961; 10: 33-46.
- Ogunlesi TO. Respiratory infections in the pre-school child: A review of 435 cases admitted to Adeoyo Hospital, Ibadan. W. Afr. Med. J. 1961; 10: 231-234.
- Odiase GI. The leading causes of death among in-patients of the University of Benin Teaching Hospital in the year 1974. Nig. Med. J. 1978; 8: 242-248.
- 14. Sofowora EO, Onadeko BO. Complications and

prognostic factors in pneumonia among Nigerians. Nig. Med. J. 1973; 3: 144-145.

- Gans B, Mcnamara FN, Morley DC, Thomson SW, Watt A. Some observations on the epidemiology of measles in West Africa. W. Afr. Med. J. 1961; 10: 253-262.
- Lewis FA, Lehmann NI, Ferris AA. The haemadsorption viruses in laryngotracheobronchilis. Med. J. Austr. 1961; 48: 929-932.
- Hilleman MR, Hamparian VV, Ketler, A, et al. Acute respiratory illnesses among children and adults. JAMA 1962; 180: 445-451.
- McLean DM, Bach RD, Larke RPB, McNaughton GA. Myxoviruses associated with acute laryngotracheobronchitis in Toronto, 1962-1963. Can. Med. Assoc. J. 1963; 89: 1257-1259.
- Chanock RM, Parrott RH. Acute respiratory disease in infancy and childhood: Present understanding and prospects for prevention. Pediatrics 1965; 36: 21-39.
- Berglund B, Vihma L, Wickstrom J. Respiratory syncytial virus studies on children hospitalised during an outbreek of respiratory illness in Finland. Am. J. Epidem. 1965; 81: 271-282.
- Dobrev I, Mihailov A, Karaivanova G. The level of complement binding antibodies against some respiratory viruses. Prob. Infec. and Paras. Dis. 1978; 6: 20-27.
- Dobrev I, Karaivanova G, Mihailov A. Etiology of acute respiratory diseases during the period 1976-1978 in Bulgaria. Epidemiol. Mikrobiol. Infekc. Bol. 1980; 1: 40-49.
- Glezen WP, Denny FW. Epidemiology of acute lower respiratory disease in children. N. Engl. J. Med. 1973; 288: 498- 505.
- Parrott RH, Kim HW, Arrobio JO. Epidemiology of respiratory syncytial virus infection in Washington, DC. II. Infection and disease with respect to age, immunologic status, race, and sex. Am. J. Epidem. 1973; 98: 289-300.
- WHO. A programme for controlling acute respiratory infections in children: Memorandum from a WHO meeting. Bull. WHO. 1984; 62: 47-58.
- Bloom HH, Johnson KM, Jacobsen R, Chanock RM. Recovery of parainfluenza viruses from adults with upper respiratory illness. Am. J. Hyg. 1961; 74: 50-59.
- Chanock RM, Kim HW, Vargosko AJ, et al. Respiratory syncytial virus. I. Virus recovery

and other observations during 1960 outbreak of bronchiolitis, pneumonia, and minor respiratory diseases in children. JAMA 1961; 176: 647-653.

- Rosen L. Hemagglutination-inhibition antibody responses in human adenovirus infections. Proc. Soc. Exp. Biol. Med. 1961; 108: 474-479.
- Parrott RH, Vargosko AJ, Kim HW, Bell JA, Chanock RM. Acute respiratory diseases of viral etiology. III. Parainfluenza. Am. J. Pub. Health 1962; 52: 907-917.
- Tyrell DAJ, Bynoe ML. Some further virus isolations from common colds. Br. Med. J. 1961; I. 393-397.
- Tyrrell DAJ, Chanock RM. Rhinoviruses: A description. Science 1963; 141: 152-153.
- Mihailov A, Dobrev I, Karaivanova G. Level of collective immunity to rhinovirus types 1A, 2, 5, 7, 8 and 11 in different age groups. Epidemiolog. Mikrobiolog. Infekc. Bol. 1980; 1: 54-60.
- 33. Karaivanova G, Dobrev I, Mihailov A. Serological and epidemiological data for the spreading of reovirus types 1, 2, 3 in human sera. Letopisy na HES 1976; 5: 132-134.
- Tyrrell DAJ, Alexander DJ, Almeida JD, et al. Coronaviridae: Second report. Int. Virol. 1978; 10: 321-328.
- Dobrev I, Karaivanova G, Mihailov A. Coronavirus 229E in the etiology of acute respiratory diseases in Bulgaria. Prob. Infec. and Paras. Dis. 1981; 9: 28-33.
- Canil KA, Pratt D, Sungu MS, Phillips PA. Influenza surveillance: Alternative laboratory techniques for a developing country. Bull. WHO. 1985; 63: 79-82.
- Taylor R, Nemaia H, Tukuitonga C, et al. An epidemic of influenza in the population of Niue. J. Med. Virol. 1985; 16: 127-136.
- Rao BL. Investigation of 1979 influenza outbreak of types A and B influenza viruses in Pune. India. Ind. J. Chest Dis. and Allied Sciences 1980; 22: 143-145.
- Rao BL. Investigation of the monsoon outbreak of Influenza A (H3N2) virus strain in Pune, India, 1981. Ind. J. Med. Res. 1983; 77: 417-419.
- WHO. Progress in the development of influenza vaccines: Memorandum from a WHO meeting. Bull. WHO. 1987; 65: 289-293.
- Ghafour A, Burney MI. Studies on influenza epidemics on Rawalpindi and Islamabad. Pak. J.

Med. Res. 1981; 20: 36-39.

- Doraisingham S, Goh KT, Ling AE, Yu M. Influenza surveillance in Singapore. 1972-86. Bull. WHO. 1988; 66: 57-63.
- Rao BL, Kadam SS, Pavri KM, Kothavale VS. Epidemiological, clinical, and virological features of influenza outbreaks in Pune, India, 1980. Bull. WHO. 1982; 60: 639-642.
- Macfarlane JT, Adegboye DS, Warrell MJ. Mycoplasma pneumoniae and the aetiology of lobar pneumonia in northern Nigeria. Thorax, 1979; 34: 713-719.
- Bell SD. Jr, McComb DE, Murray ES, Chang WS-M, Synder JC. Adenoviruses isolated from Saudi Arabia. I. Epidemiologic features. Am. J. Trop. Med. Hyg. 1959; 8: 492-500.
- Tai F-H, Grayston JT. Adenovirus neutralising antibodies in persons on Taiwan. Proc. Soc. Exp. Biol. Med. 1962; 109: 881-884.
- Chan VF, Espiritu-Campos L, Cenebre L, Guinto-Famatiga E. Viruses of the respiratory tract in Filipinos. I. Parainfluenza types 1 and 3.
 J. Philipp. Med. Assoc. 1963; 39: 303-306.
- Jennings R, Grant LS. Respiratory viruses in Jamaica: A virologic and serologic study 1. Virus isolations and serologic studies on clinical specimens. Am. J. Epidemiol. 1967; 86: 690-699.
- Evans AS, D'Allessio DA, Espiritu-Campos L, Dick EC. Acute respiratory disease in University of the Philippines and University of Wisconsin students: A comparative study. Bull. WHO. 1967; 36: 397-407.
- Monto AS, Johnson KM. A community study of respiratory infections in the tropics. I. Description of the community and observations of the activity of certain respiratory agents. Am. J. Epidemiol. 1967; 86: 78-92.
- Deibel R, Vivell O, Lips G. Serologische und klinische untersuchungen ueber viruserkrankungen des respirationstrakts. III. Mitteilung. Durchseuchungsstudien mit 12 antigendifferenten viren des respiratiostraktes. Z. Kinderheilk 1962; 86: 543-552.
- Huebner RJ, Rowe WP, Ward TG, Parrott RH, Bell JA. Adenoidal pharyngeal-Conjunctival agents: A new recognised group of common viruses of the respiratory system. N. Engl. J. Med. 1954; 251: 1077-1086.
- Chanock RM, Finberg L. Recovery from infants with respiratory illness of a virus related to chimpanzee coryza agent (CCA). II.

Epidemiologic aspects of infection in infants and young children. Am. J. Hyg. 1957; 66: 291-300.

- Dreizen RS, Zolotarskaia EE, Davidova AA. Immunologic structure of the Moscow populace to adenoviruses. Voprosy Virusol 1962; 7: 85-91.
- 55. Karaivanova G. Results of laboratory research during 1984 in Bulgaria for demonstration the etiology of noninfluenza viral acute respiratory diseases. Official Bull. of Res. Inst. Infec. and Paras. Dis. 1985; 7: 14-18.
- 56. Dobrev UI, Karaivanova G, Mihailov A, Georgeiva G, Kolarova G. On the viral etiology of acute respiratory diseases in a "Mother and Child" home in 1978-1979. I. Laboratoryvirologic and clinical epidemiologic data. Epidemilog. Mikrobiolog. Infekc. Bol. 1981; 3: 245-253.
- Valenti WM, Clarke TA, Hall CB, Menegus MA, Shapiro DI. Concurrent outbreaks of rhinovirus and respiratory syncytial virus in an intensive care nursery: Epidemiology and associated risk factors. J. Pediatr. 1982; 100: 722-726.
- Chanock R, Chambon L, Chang W, et al. WHO respiratory disease survey in children. A serological study. Bull. WHO. 1967; 37: 363-369.
- Sutmoller F, Nascimento JP, Chaves JRS, Ferreira V, Pereira MS. Viral etiology of acute respiratory diseases in Rio de Janeiro: First two years of a longitudinal study. Bull. WHO. 1983; 61: 845-852.
- Berman S, Duenas A, Bedoya A, et al. Acute lower respiratory tract illnesses in Cali, Colombia: A two-year ambulatory study. Pediatrics 1983; 71: 210-218.
- Spence L, Barratt N. Respiratory syncytial virus associated with acute respiratory infections in Trinidad patients. Am. J. Epidemiol. 1968; 88: 257-266.
- 62. Sobeslavsky O, Sebikari SRK, Harland PSEG,

Skrtic N, Fayinka OA, Soneji AD. The viral etiology of acute respiratory infections in children in Uganda. Bull. WHO. 1977; 55: 625-631.

- Ong SB, Lam KL, Lam SK. Viral agents of acute respiratory infections in young children in Kuala Lumpur. Bull. WHO 1982; 60: 137-140.
- Teo J, Vellayappan K, Yip WCL, Doraisingham S. Mycoplasma pneumoniae and viral infections in childhood asthma. J. Trop. Pediatr. 1986; 32: 87-89.
- Ogunbi O. Bacteriological and viral aetiology of bronchiolitis and bronchopneumonia in Lagos children. J. Trop. Med. Hyg. 1970; 73: 138-140.
- 66. Njoku-Obi AN, Ogunbi O. Viral respiratory diseases in Nigeria: A serological survey. II. Complement fixing antibody levels of adenoviruses, respiratory syncytial virus, psittacosis virus. J. Trop. Med. Hyg. 1966; 69: 147-149.
- Dym AM, Schuit KE, Nwankwo MU, Omene JA. Respiratory syncytial virus and acute lower respiratory infections in Benin City, Nigeria. Pediatr. Inf. Dis. 1986; 5: 717-718.
- Okuonghae HO. Nosocomial respiratory syncytial virus infection in a newborn nursery in Benin. Dissertation, April 1989.
- Craighead JE, Shelokov A, Peralta PH, Vogel JE. Croup-associated virus infection in adults: Report of two cases. N. Engl. J. Med. 1961; 264: 135-137.
- David-West TS, Cooke AR. Laboratory and clinical investigation of the 1974 influenza epidemic in Nigeria. Bull. WHO. 1974; 51: 103-105.
- Doraisingham S, Ling AE. Patterns of viral respiratory tract infections in Singapore. Ann. Acad. Med. Singapore. 1986; 15: 9-14.
- Kloene W, Bang FB, Chakraborty SM, et al. A two-year respiratory virus survey in four villages in West Bengal, India. Am. J. Epidemiol. 1970; 92: 307-320.

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