Emerging pathogens and possible threats to blood services

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New pathogens emerge regularly. These new agents include both bacterial and viral pathogens. Many viral agents have a viremic phase during the course of infection and are transmitted by blood transfusion. New infectious agents provide a major challenge for blood services. The extend and mode of transmission through blood is initially unclear. Serological tests and molecular tools are used to detect infectious donors. A residual risk remains due to the diagnostic window period and sensitivity limits of the tests systems. Some detection systems proved to be extremely helpful, e.g. for HIV, HCV or WNV, but sometimes provide in certain periods of the infection false-negative results due to sensitivity problems or to escape mutants (e.g. Hepatitis B). WNV provides an excellent example, how rapid and efficient blood donation services coped with this new threat for blood safety. Not all emerging infectious agents transmissible by blood represent a threat for the health. A number of putative pathogens turned out later not to be associated with pathogenesis. Prion diseases represent a particular challenge for detection in blood donors. Since the ‘normal’ cellular form the host protein is widely distributed, the usual nucleic acid amplification systems or serological tests are not feasible. Extremely sensitive tests (picogram range) are required for detection of the pathological protein in the asymptomatic period. Variants of existing viruses e.g. newly emerging Influenza A subtypes with the potential for pandemic spread represent an enormous threat for public health and for the safety of blood. Blood services should take precautions for an influenza pandemic. The mosquito borne Chikungunya virus turned out to be transmitted by blood transfusion and caused many infections in the islands of the Indian Ocean and in India. Newly emerging infectious agents require attention and rapid actions in order to guarantee a safe and reliable blood supply.

In the last three decades, on average roughly once a year a new pathogen emerged in one or the other area of the world. A substantial number of these new or re-emerging pathogens continues to remain a threat for the health of the population and may remain with mankind for the foreseeable future. These new agents include both bacterial and viral pathogens like Borrelia, human immunodeficiency virus (HIV), helicobacter, hepatitis C or the agents causing transmissible spongiform encephalopathies. Some of these newly emerging agents were successfully contained and do presently not cause disease, for example, the severe acute respiratory syndrome Corona virus. One may predict that also in the future new pathogens will emerge. Some of these agents were never recognized previously, like severe acute respiratory syndrome or HIV. Various other agents show suddenly a new distribution or a new form of disease manifestation, for example, West Nile virus. A substantial number of pathogens are transmitted from animals to humans, like monkey pox or Nipah virus. In addition, some well known agents cause previously unknown disease, for example, Parvovirus B19, and new forms of previously known agents appear that differ in their properties from previously identified forms, as HIV 1 group O or variant Creutzfeldt–Jakob disease. Finally, one cannot exclude the deliberate release of infectious agents through bioterroristic actions. Many viral agents have a viraemic phase during the course of infection and are therefore transmitted by blood transfusion or cellular blood components and sometimes also plasma products. New infectious agents provide a major challenge for blood services. Initially, there is poor information about the properties of the individual agent and the course of the disease. The extent and mode of transmission through blood is initially unclear and sensitive and specific tests are missing. It requires a substantial amount of time to establish robust tests for screening of blood donations in large numbers. Such tests are required during the asymptomatic period of the disease where seemingly healthy patients still donate blood but are infectious (e.g. West Nile virus). Donor selection criteria are not seldom only partially helpful (e.g. residence in the UK for variant Creutzfeldt-Jakob disease) and inactivation procedures are only useful for a limited number of cellular blood components in contrast to plasma products where effective inactivation procedures exist. The tragic experience with HIV sensitized the population and in many countries a zero risk is – unrealistically – expected. Serological tests and molecular tools are prerequisites to detect infectious donors. A residual risk remains, due to the diagnostic window
The importance of proper hygiene measures and barrier systems in blood services is evident. These systems require detailed training of personnel to ensure the safety of blood and blood products. Such services develop a concept for maintaining their functions in a pandemic situation. Proper planning is crucial to prevent disruptions in blood supplies.

Some pitfalls in the detection of emerging infectious agents can be significant. Despite these challenges, tests may still be necessary. For example, the protein-only hypothesis suggests that serological and molecular tools may not provide a sufficiently reliable screening system. In such cases, alternative methods like protein misfolding cyclic amplification might be considered. The ability to amplify the pathological protein could provide a critical diagnostic tool.

Influenza, a virus that can spread rapidly and has the potential for pandemic spread, offers an example of the need for proactive measures. The ability of a virus to spread from human to human in larger scale provides pandemic capacity. Such a virus could – but must not – originate from the presently circulating H5N1 virus. Blood services should take precautions for an influenza pandemic and also exclusion of donors due to antiviral treatment. The time course and extent of viraemia in an influenza infection remain to be determined for the emerging pandemic virus. The duration of the asymptomatic period should be short, that is, only a few days. The period when a blood donation is infectious remains then to be analysed.

The successful spread of West Nile virus (WNV) within 5 years from the US East coast all the way across the continent to the West coast is an excellent example. WNV spread dramatically due to infections or fears of infection. There might be shortage of blood during the pandemic and also exclusion of donors due to antiviral treatment. The time course and extent of viraemia in an influenza infection remains to be determined for the emerging pandemic virus. The duration of the asymptomatic period should be short, that is, only a few days. The period when a blood donation is infectious remains then to be analysed. Experience from other virus subtypes or the regular seasonal influenza may not apply for the pandemic virus and may not be informative for newly emerging subtypes. The properties of a pandemic virus will only be known when the pandemic does indeed appear. H5N1 virus was found in plasma in a child by reverse transcriptase polymerase chain reaction. The plasma of this severely ill patient was obtained in a terminal period of the disease. During an influenza pandemic, a viraemia in the pre-symptomatic phase of an influenza infection has to be assumed. Due to the short incubation period, the number of influenza transmissions through blood components may remain lower than frequently feared. It is important that blood services develop a concept for maintaining their functions in a pandemic situation. Proper hygiene measures and barrier systems are important and require detailed training of the personnel.
Prevention of infections may prove to be more efficient than the use of antiviral drugs. In a pandemic situation, shortages of blood donations may frequently be compensated by postponement of elective surgery. Finally, the Chikungunya virus (Alpha virus), transmitted similarly to Dengue virus by mosquitoes, turned out to be transmitted by blood transfusion. This virus caused a substantial number of infections in the islands of the Indian Ocean (La Reunion, Mauritius, Seychelles, Madagascar) and in India. In La Reunion, 20–30% of the population were infected. Transport of blood from France was necessary to maintain blood supply. Chikungunya cases in the Italian region Emilia Romana caused concern about the spread of Chikungunya to other parts of Europe. Newly emerging infectious agents require particular attention and rapid actions in order to guarantee a safe and reliable blood supply.