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Orthopoxviruses belonging to the family of Poxviruses that infect humans are variola (smallpox), vaccinia, monkey pox and cowpox. Poxviruses are large (200–450 nm in size), brick- or ovoid-shaped double-stranded DNA viruses that are especially adapted to epidermal cells. 9 Variola virus, the cause of small pox is the only one that affects humans exclusively, the others are zoonotic infections.

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The genus Orthopoxvirus of the family Poxviridae includes species pathogenic to humans, such as variola (VARV), monkeypox (MPXV), cowpox (CPXV), and vaccinia (VACV) viruses. VARV causes smallpox and is an exclusively anthroponotic agent.

Multiplex PCR detection and species differentiation of ...  
OBJECTIVE: Variola virus, belonging to Orthopoxviridae family, is one of the most dangerous human pathogens that could be used as biological weapon. We have developed a new rapid assay, based upon Real-time PCR and melting temperatures analysis of amplicons, for the contemporary detection of Orthopoxvirus, VZV and HSV1-2, that are the most important infectious agents to be considered for differential diagnosis.

[Rapid differential diagnosis of Orthopoxviruses and ...  
However, other orthopoxviruses (OPVs), such as cowpox virus, buffalopox virus (a subspecies of vaccinia virus), and especially monkeypox virus (1), can also infect humans through contact with wildlife and farm animals, since each of these infectious agents has vast natural reservoirs (7, 31).