

In-Silico Evaluation of the Capsid Proteins of FMDV as Potential Vaccine Candidates

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Abstract: In this study, the capsid proteins of four major serotypes of Foot and Mouth Disease Virus (FMDV) were assessed as the vaccine candidates. Different protein sequences regarding FMDV capsid of O, A, Asia 1 and C type were identified from NCBI Genome Database and UniprotKB. Phylogenetic tree of the four serotypes was developed using ClustalW software. HMMTOP, RANKPEP, Swiss-Model and Vaxign software were used for comparing the capsid proteins in terms of their feasibility as vaccine candidates. The virus and viral serotype were identified from the cultured disease sample using RT-PCR. Our results revealed that different capsid proteins of the four serotypes vary in their suitability to be considered as peptide vaccine components. Viral protein 1 (VP1) for Asia 1 serotype represented the best result as a vaccine candidate. The VP1 region of Asia 1 serotype amplified based on the result of dry lab analysis. Our findings provide a future indication of multivalent vaccine development against FMDV.

Keywords: Reverse Vaccinology, FMDV, Capsid Protein, Viral Protein 1, Vaccine Candidate

1. Introduction

Vaccines have been developed for generating effective immunogenicity to prevent diseases throughout previous decades [1-2]. Different methodologies have been exploited so far to develop vaccines against life threatening diseases of human and animals [2]. Reverse vaccinology has come out as one of the most modern system of vaccine development in recent years [3]. It takes advantages of genomic and proteomic data regarding pathogenic organisms already available in the databases. Antigenic materials of certain pathogens can be analyzed by using different computational resources and tools. Results from the computational approach can be utilized to develop vaccine within 1-2 years through further experimental approaches.

However, foot and mouth disease (FMD) is an economically important and highly contagious viral disease.

The FMDV viral particle (25-30 nm) contains an icosahedral capsid consisting of proteins and no envelope [4-5]. The virus possesses a positive-sense single stranded RNA (SS RNA) (about 8.3 kb) genome that encodes a polyprotein which is subsequently processed to yield structural and non-structural proteins [8-9]. Globally, the virus exists in seven immunologically distinct serotypes; the Southern African Territories [SAT] types 1-3 and Eurasian types namely O, A, C and Asia 1, with multiple subtypes within each serotype [10]. Among these serotypes show some regionality; the O serotype is the most common while four serotypes (O, A, Asia 1, C) are available in south Asian countries [4]. The RNA genome of FMDV goes through a high rate of mutation because of error prone replication by the RNA polymerase which results in high genetic diversity [11-12]. Additionally, persistent infection, recombination, and quasi-species dynamics have also been reported as contributing factors to

the genetic variation [12-13]. One major concern is that immunity to one serotype of FMDV does not confer protection against another. The complex intra-serotypic variation coupled with the presence of multiple serotypes has complicated disease control, specifically in case of vaccination. The most common forms of vaccines against FMDV are killed or attenuated vaccines [14]. The major problems regarding such vaccination are requirement of high specificity, attainment of temporary immunity (months to years), requirement of revaccination for prophylactic control, reversal effects of vaccine components [14]. Moreover, vaccination against FMDV lacks induction of rapid protection against challenge or prevention of the development of the carrier state. Furthermore, it is evidential that the clinical protection depends upon the span of immunization and the period of exposure/challenge methods [15]. All these aspects have created great challenges in development of vaccine against FMDV. Subunit vaccine or peptide vaccine for FMDV prevention has been suggested by several authors in this regard [16]. Subunit vaccine is a vaccine that contains viral antigens made free of viral nucleic acid. It is less possibility to cause adverse reactions than a vaccine containing the whole virion.

The aim of our study was to compare and analyze the capsid proteins (VP0, VP1, VP2, VP3 and VP4) of different Eurasian serotypes (O, A, Asia 1, C; UniprotKB entry P03305 P49303, E9KMQ6, P15072 respectively) of FMDV through computational approach in order to evaluate their feasibility as vaccine candidates. In our dry lab approach, FMDV genome polyprotein sequences of different serotypes

have been identified, and similarity and dissimilarity among the sequences were analyzed. Different bioinformatics tools were utilized to analyze the capsid proteins of four serotypes for their antigenic and immunogenic property. In a wet lab study, we conducted molecular characterization for particular serotypes from tissue samples of suspected FMDV infected cattle of Savar Military Firm, Dhaka Bangladesh.

2. Materials and Methods

2.1. Dry Lab Study

At first the genome polyprotein sequences of four serotypes of FMDV were identified from NCBI Genome Database. Multiple sequence alignment of the genome polyproteins of four serotypes of FMDV was done by ClustalW [17] to determine the mutations among the serotypes (Supplementary file 1). Maximum likelihood method was used for phylogenetic tree construction from MEGA software in which boot strapped value was 1000. The serotypes under study show very close phylogenetic relationship with approximately 90% sequence similarity among them and some observable point mutations (Figure:1A and 1B). Phylogenetic tree acts as harbinger to find out the ancestral relationship between different types of viruses. In this research Phylogenetic tree analysis depicts that four types of virus are closely related to each other. So it will act as quintessence to develop single type of vaccine to unfold the intricateness of these viruses.

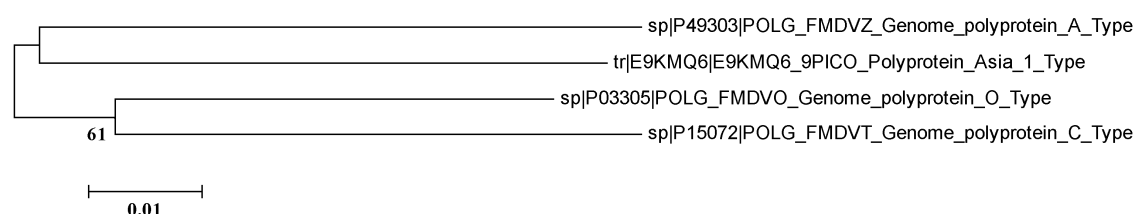


Fig. 1A. Original phylogenetic tree for O, A, Asia 1 and C serotypes of FMDV.

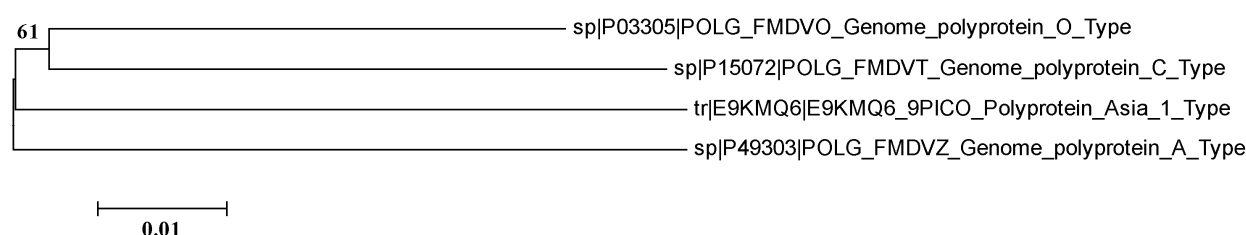


Fig. 1B. Bootstrapped consensus phylogenetic tree.

Amino acid sequence of the proteins of FMDV was identified from UniprotKB [18]. HMMTOP [19-20] was used to determine the number of transmembrane (TM) helix, topology of the antigenic proteins and feasibility of molecule cloning of the antigenic proteins. Epitope binding sites against specific MHC (major histocompatibility complex) for each antigen were determined by RANKPEP [21-22]. Determination of the adhesion probability (which denotes to

binding capacity of antigen to host surface) against specific MHC molecule of each antigen was done by Vaxign [23]. The antigenic proteins were compared based on all the parameters under study (related to immunogenicity and antigenicity) to identify the better vaccine candidates. At last the structures of each antigen were also predicted based on homology modeling by SWISS-MODEL (structure not shown in this article) [24- 26].

All protein ID and sequence information shown in supplementary file 1 and 2.

2.2. Wet Lab Study

Cell culture

Baby hamster kidney cells (BHK-21) (American Type Culture Collection, ATCC, Rockville, MD, USA) were cultured and maintained in Dulbecco's modified Eagle's medium (DMEM) supplemented with 10% heat-inactivated fetal bovine serum (Gibco Life Technologies, USA), 100 U/ml penicillin, 2 mM L-glutamine, and 100 µg/ml streptomycin.

Determination of cellular morphology upon FMDV infection

BHK-21 cells (1.6×10^5) were grown onto 60 mm tissue

culture plate. Cells were challenged with the viral sample collected from suspected FMD Virus infected cattle from Savar Military Farm, Dhaka Bangladesh. After 72 hours, images were taken under an inverted microscope with a magnification of 10X.

Reverse transcription-polymerase chain reaction (RT-PCR) assay

BHK-21 cells were challenged with the viral sample using established protocol of Virology Lab of Animal Health Research Division of Bangladesh Livestock Research Institute (BLRI). RNA was extracted by using Qiagen RNeasy kit and RT-PCR was done by Superscript III RT-PCR kit. The oligonucleotide primers for the detection of FMDV and FMDV serotypes were used from the 2B and VP1 (1D) regions of the viral genome as published (Table: 1) [27].

Table 1. List of the primers, their sequences and size of PCR amplicon were used for the diagnosis of FMD virus from the field samples

FMDV serotype	Primer label	Sequence(5'-3')	Location	PCR products (bp)
All Serotypes	P32	CAGATGCAGGAGGACATGTC	2B	131 bp
	P33	AGCTTGTACCAGGGTTTGGC	2B	
	P74	GACACCACTCAGGACCGCCG	VP1(1D)	
Asia 1	P75	GACACCACCCAGGACCGCCG	VP1(1D)	292 bp
	P76	GACACCACACAAGACCGCCG	VP1(1D)	
	P77	GACACGACTCAGAACCGCCG	VP1(1D)	

2.3. Results and Discussions

In the present study, the capsid proteins of four major serotypes (O, A, Asia 1, C) of Foot and Mouth Disease Virus (FMDV) were evaluated as the vaccine candidates. Using computational approach, we found no transmembrane helix of the viral proteins of the mentioned serotypes but there was

an entropy variation among them. VP4 region of all the mentioned FMDV serotypes showed the lowest entropy value while VP2 exhibited highest entropy for O type FMDV serotype and VP3 showed highest entropy for Asia 1, A and C type FMDV serotypes (Table 2-5). In addition, VP1 region demonstrated moderate entropy for Asia 1 serotype (Table 2).

Table 2. Dry lab analysis results of A type (FMDV)

Protein	HMMTOP Analysis		Swiss model analysis		RANKPEP for MHC I			RANKPEP for MHC II			Vaxign analysis
	No. of TM helix	Entropy	QMEAN Score	Z Score	Optimal score	Score	% OPT	Optimal	Score	% OPT	Adhesin Probability
VP1	0	17.0120	0.61	-2.53	128	85.0	66.41	48.247	17.528	36.33	0.282
VP2	0	17.0135	0.75	-0.53	128	83.0	64.84	48.247	24.199	50.16	0.467
VP3	0	17.0136	0.67	-1.62	128	78.0	60.94	48.247	13.852	28.71	0.445
VP4	0	17.0079	0.08	-4.33	128	57.0	44.53	48.247	24.199	50.16	0.580

As entropy means disorder, it will be difficult to clone a protein with higher entropy [19] and proteins with lower entropy are more feasible for molecular cloning.

In case of O type FMDV, VP4 showed highest percentage

(%) of optimal (OPT) antigenicity among these proteins (VP1, VP2, VP3 and VP4), and VP3 exhibited the lowest % OPT (Table 3)

Table 3. Dry lab analysis results of O type (FMDV)

Protein	HMMTOP analysis		Swiss model analysis		RANKPEP for MHC I			RANKPEP for MHC II			Vaxign analysis
	No. of TM helix	Entropy	QMEAN Score	Z score	Optimal score	Score	% OPT	Optimal Score	Score	% OPT	Adhesin Probability
VP1	0	17.0119	0.63	-2.26	128.0	87.0	67.97	48.247	17.528	36.33	0.352
VP2	0	17.0133	0.66	-1.79	128.0	90.0	70.31	48.247	24.199	50.16	0.558
VP3	0	17.0125	0.69	-1.34	128.0	81.0	63.28	48.247	13.852	28.71	0.519
VP4	0	17.0080	0.08	-4.33	128.0	90.0	70.31	48.247	24.199	50.16	0.580

With the highest % OPT antigenicity, VP4 would show the highest epitope binding capacity with mammalian MHC I molecule. VP1 region of A type FMDV showed highest %OPT among these proteins while VP4 represented the lowest %OPT (Table 2). Consequently, VP1 would show

the highest epitope binding capacity with mammalian MHC I molecule. In addition, VP2 and VP3 of Asia 1 type FMDV exhibited the same highest %OPT among these proteins (Table 4).

Table 4. Dry lab analysis results of Asia1 type (FMDV)

Protein	HMMTOP Analysis		Swiss model analysis		RANKPEP for MHC I			RANKPEP for MHC II			Vaxign analysis
	No. of TM helix	Entropy	QMEAN Score	Z Score	Optimal score	Score	% OPT	Optimal Score	Score	% OPT	Adhesin Probability
VP1	0	17.0093	0.72	-1.04	128.0	88.0	68.75	65.642	19.542	29.77	0.273
VP2	0	17.0136	0.69	-1.42	128.0	97.0	75.78	65.642	13.432	20.46	0.588
VP3	0	17.0145	NA	NA	128.0	97.0	75.78	65.642	8.128	12.38	0.534
VP4	0	17.0086	0.08	-4.35	128.0	57.0	44.53	65.642	6.686	10.19	0.571

As expected, these two proteins should have highest epitope binding capacity with mammalian MHC I molecule. On the other hand, VP4 showed the lowest % OPT and VP1 showed a standard value %OPT (Table 4). VP1 region of C

type FMDV, revealed the highest % of OPT among these proteins while VP4 displayed the lowest OPT (%) (Table 5). Thus, VP1 should have the highest epitope binding capacity with mammalian MHC I molecule.

Table 5. Dry lab analysis results of C type (FMDV)

Protein	HMMTOP Analysis		Swiss model analysis		RANKPEP for MHC I			RANKPEP for MHC II			Vaxign analysis
	No of TM helix	Entropy	QMEAN Score	Z Score	Optimal score	Score	% OPT	Optimal score	Score	% Opt	Adhesin Probability
VP1	0	17.0094	0.63	2.12	128.0	85.0	66.41	65.642	11.9	18.13	0.435
VP2	0	17.0136	0.6	-2.6	128.0	80.0	62.50	65.642	13.432	20.46	0.613
VP3	0	17.0144	0.61	-2.51	128.0	82.0	64.06	65.642	6.056	9.23	0.493
VP4	0	17.0079	0.08	-4.33	128.0	57.0	44.53	65.642	6.726	10.25	NA

The study of epitope binding capacity of the viral proteins with mammalian MHC II molecule, we found that VP2 and VP4 region of O type FMDV represented the same highest OPT (%) among these proteins (Table 3). These two proteins should have highest epitope binding capacity with mammalian MHC II molecule. Additionally, VP1 of A, Asia 1, and C type FMDV demonstrated the highest OPT (%) among these proteins (Table 2-4). Consequently, VP1 should have the highest epitope binding capacity with mammalian MHC II molecule for the above mentioned three types of FMDV.

Previous studies have been suggested that the higher epitope binding capacity of particular proteins denotes to their viability as vaccine components than the proteins of lower epitope binding efficiency [21-22, 28]. Based on the binding capacity of antigenic proteins to mammalian MHC I and MHC II molecules, we concluded that VP1 region can be a novel vaccine components for A, Asia 1 and C types of FMDV. Although, the results have been based on the sequences of highest possible OPT (%) for each capsid protein, we therefore further studied the 3D structure and adhesion probability of the antigenic proteins from their sequence.

Adhesion probability represents the binding capacity of antigens to the host cell. It has been reported that highest adhesion probability corresponds to the highest binding

capacity [23, 31]. We found that VP4 of O and A type FMDV showed the highest adhesion probability and VP1 displayed the lowest value (Table 3, 2). In case of Asia 1 and C type FMDV, VP2 showed the highest adhesion probability and VP1 demonstrated the lowest value (Table 4-5).

Different parameters regarding the feasibility of vaccine candidates showed different results for different serotypes (in some cases HMMTOP and Vaxign analysis do not show results due to short sequence). There was no single capsid protein that showed top results in case of all studies. This actually indicates that no single capsid protein can be treated as the best vaccine component against FMDV. Different antigenic proteins can be suitable candidates from different point of view. So peptide vaccine based on a single protein for all types of FMDV will not provide the best results. So far VP1 has been regarded as the best antigen for producing peptide vaccine against FMDV [16]. Moreover, the structural protein coding region VP1 has been shown to vary significantly between strains and serotypes indicating the higher mutation rate than the structural protein coding gene of FMDV [32-33]. This reduces the suitability of VP1 for being the monovalent peptide vaccine. The overall study leaves the chance to think about the production of multivalent vaccine against FMDV instead of monovalent one. This may combine the prominent capsid proteins across different serotypes of FMDV to prepare a novel

vaccine against multi-serotypes of FMDV [14].

According to the results of dry lab analysis, we tried to find out the vaccine candidate in wet-lab research. The findings of the wet-lab study confirmed that the virus

infected samples contained FMDV and the serotype was Asia-1. We identified Vp1 region of Asia 1 region which showed best result in dry lab as vaccine candidate (fig:2)

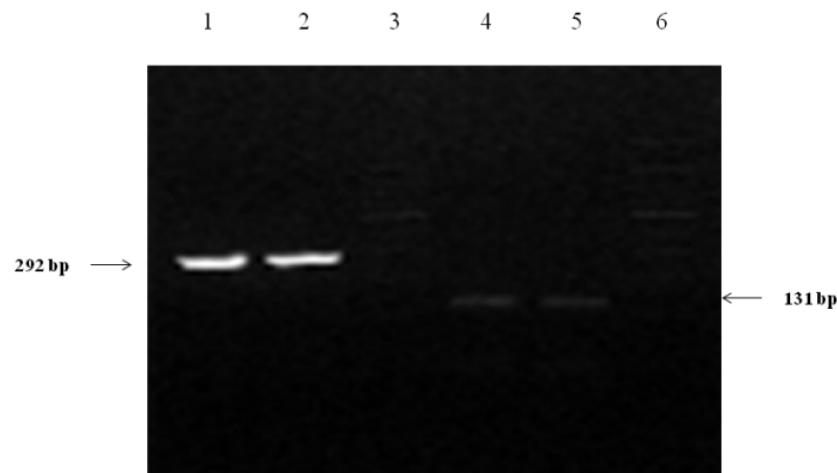


Fig. 2. Confirmation of FMDV and VP1 in tissue samples using RT-PCR. Total RNA was extracted from FMDV-infected vascular fluid of tongue epithelium tissue and reverse-transcribed using two specific primers designated as 2B and VP1 (1D) regions of the viral genome for the detection of FMDV and FMDV serotypes. Lane 1-2, VP1 region of Asia 1 serotype (292 bp); lane 3, 100 bp DNA marker; lane 4-5, B1 region of all serotypes; lane 6, 100 bp DNA marker.

We also observed the cytopathic effect of virus sample (Fig 3). Further study is needed to be done for VP1 region of different serotypes that can be assembled as a multivalent vaccine. Mainly four serotypes (O, A, Asia 1 and C type) of

FMDV are predominant in Bangladesh and other countries of South Asia. Among them Asia-1 has been mostly reported in Bangladesh.

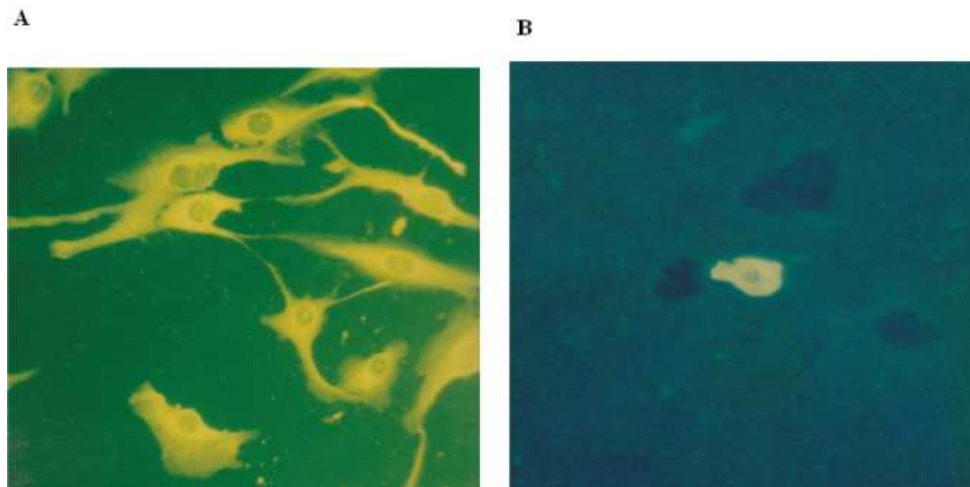


Fig. 3. Cytopathic effects of FMDV in BHK-21 cells. About 1.6×10^5 BHK-21 cells were grown onto 60 mm cell culture plate and infected with FMDV at multiplicity of infection (MOI) of 1. After 72 hrs cells were rinsed with 1 ml of 1X PBS buffer and the cellular morphological appearance was observed using inverted microscopy (Olympus, Canada) with a magnification of 10X. (A) Mock. (B) Infection with FMDV.

In conclusion, the present study mainly focused on genomic based approach of vaccine development known as reverse vaccinology. In the dry lab study, the capsid proteins of different serotypes of FMDV showed different levels of feasibility that are to be considered as peptide vaccine components. Since the idea of producing single peptide vaccine can be replaced by the concept of multivalent vaccine. The wet lab study identified the Asia 1 serotype of FMDV in the samples of suspected animals. The results can be further validated in the laboratory. Our study enhances our knowledge for the possibility of producing novel vaccine

based on VP1 sequence of multiple serotypes.

Author Contributions

FMNH has contributed to idea development, wet lab and dry lab experimentation, and data generation. KMTR, SSS, MSR, MFI and MBA have contributed to data analysis, literature mining and interpretation of results. MSR and MFI has contributed to data maintenance and handling. KMH and MG have supervised the whole work. All the authors have contributed equally to the writing of the paper.

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Genome Polyprotein Sequences of Different Serotypes of FMDV

O Type

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>sp|P03305|POLG_FMDVO Genome polyprotein
OS=Foot-and-mouth disease virus (isolate
Bovine/Germany/O1Kaufbeuren/1966 serotype
O) PE=1 SV=1
MNTTDCFIALVQAIKALFLSRTTGKMELTLYNGEKKTF
YSRPNNHNCWLNAILQLF
RYVEEPFFDWWYSSPENLTLEAIKQLEDLTGLELHEGGPPA
LVIWNIKHLHTGIGTASR
PSEVCMVDGTMCLADFHAGIFLKGQEHAVFACVTSNGWYA
IDDEDFFYPWTPDPSDVLVF
VPYDQEPLNGEWKAKVQRKLKGAGQSSPATGSQNQSGNTGS
IINNYMQQYQNSMDTQLG
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GALLADKKTEETTLLEDRI
LTTRNGHTTSTTQSSVGVTYGYATAEDFVSGPNTSGLETRV
VQAERFFKTHLFDWVTSDS
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RELYQLTLFPHQFINPRTNMTAHITVPFVGVNRYDQYKVHK
PWTLVVMVAPLTVNTEGA
PQIKVYANIAPTNVHVAGEFFPSKEGIFPVACSDGYGGLVTT
DPKTADPVYGVKVFNPFRNQ
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FSIPYLSAADYAYTASGVAETTNVQGWVCLFQITHGKADGD
ALVVLASAGKDFELRLPVD
ARAETTSAGESADPVTTTVENYGGGETQIQRRQHTDVSFIMD
RFVKVTPQNQINILDLMOI
PSHTLVGALLRASTYYFSDLEIAVKHEGDLTWVPNGAPEKA
LDNTTNPTAYHKAPLTRLA
LPYTAPHRVLATVYNGECRYNRNAVPNLRGDLQVLAQKVAR
TLPTSFNYGAIKATRVTEL
LYRMKRAETCYPRLLAIHPTEARHKQKIVAPVKQTLNFDL
LKLADGVESNPGPFFFSVD
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AIRTGLDEAKPWYKLIKLL
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DSLSSLFHVPAPVFSFGAP
VLLAGLVKVASSFFRSTPEDLERAEKQLKARDINDIFAILK
NGEWLVKLILAIRDWIKAW
IASEEKFTMTDLVPGILEKQORDLNDPSKYKEAKEWLDNAR
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KPFNSKVIIATTNLYSGFT
PRTMVCPCDALNRRFHFDDIDVSAKDGYKINSKLDIIKALED
HANPVAMFQYDCALLNGMA
VEMKRMQQDMFKPQPPLQNVYQLVQEVDRVELHEKVSSH
IFKQISIPSQKSVLYFLIE
KGQHEAAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLK
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IRETRKRQKMVDVAVNEYIEKANITDDDKTLDEAEKSPLET
SGASTVGFRERTLPGQKAC
DDVNSEPAQPVEEQPQAEQPYAGPLERQKPLKVRALPQQE
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MGNTKPVLELIDGKTVAIC
CATGVFGTAYLVPRHLFAEKYDKIMVDGRAMTDSYRVFEF
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RGNRVRDITKHFRDTARMKKGTTPVGVINNADVGRILFISGE
ALTYKDIVVCMGDGTMPLG
FAYRAATKAGYCGGAVLAKDGADTFIVGTHSAGGNGVGYCS
CVSRSMMLKMAHIDPEPH
HEGLIVDTRDVEERVHVMRKTKLAPTVAHGVFNPEFGPAAL
SNKDPRLNEGVLDEVIFS
KHKGDTKMSEEDKALFRCAADYASRLHSLVGTANAPLSIY
EAIKGVLDGLDAMEPDTAPG
LPWALQKRRGALIDFENGTVGPEVEAALKLMEKREYKFVC
QTFCLKDEIRPLEKVRAGKT
RIVDVLPVEHILYTRMMIGRCAQMHSSNNGPQIGSAVGCNP
DVDWQRFQTHFAQYRNVD
VDYSAFDANHCSDAMNIMFEEVFRTEFGFHPNAEWILKTLV
NTEHAYENKRITVGGGMP
GCSATSIINTILNIIYVLYALRRHYEGVELDITYTMISYDD
IVVASDYDLDFEALKPHFK
SLGQITITPADKSDKGFVLGHSITDVTFLKRHFHMDYGTGFY
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TIQEKLISVAGLAVHSGPDEYRRLFEFPQGLFEIPSYRSLY
LRWVNAVCGDA
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A type

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>sp|P49303|POLG_FMDVZ Genome polyprotein
OS=Foot-and-mouth disease virus (isolate
-/Azerbaijan/A22-550/1965 serotype A)
PE=1 SV=1
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YSRPNNHNCWLNLTILQLF
RYVDEPFFDWWYDSPAENLTCEAIRQLEEITGLELHEGGPPA
LVIWNIKHLHTGIGTASR
PSEVCMVDGTMCLADFHAGIFLKGQEHAVFACVTSNGWYA
IDDEDFFYPWTPDPSDVLVF
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IINNYMQQYQNSMDTQLG
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DNAISGGSNEGSTDTTSTHTTNTQNDWFSKLASSAFSGLF
 GALLADKKTEETTLLEDRI
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 FGHLEKLELPTDHKGVYGLVDSFAYMRNGWDVEVSAVGNQ
 FNGGCLLVAMVPEWKELTP
 REKYQLTLFPHQFISPRTNMTAHIVVPYLGVNRYDQYKKHK
 PWTLVVMVVSPLTTNTVSA
 GQIKVYANIAPTHVHVAGELPSKEGIVPVACSDGYGGLVTT
 DPKTADPVYGMVYNPPTN
 YPGRFTNLLDVAEACPTFLCFDDGKPYVVTTRTDEQRLLAKF
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 YYAQYSGTINLHFMFTGSTDSKARYMVAYVPPGVETPPDTP
 EKAACIHAEDWTGLNSKF
 TFSIPYVSAADYAYTASDVAETTQVQWVCIIYQITHGKAEQ
 DTLVSVSAGKDFELRLPI
 DPRSQTTSTGESADPVTITVENYGETQVQRRQHTDVTFIM
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 THQHGLVGLLRAATYFSDLEIVVRHDGNLTWVPNGAPEA
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 QLPTSFNFGAIQATTIHEL
 LVRMKRAELYCPRLLAVEVSSQDRHKQKIAPAKQLLNFD
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 VRSNFSKLVTINQMQEDMSTKHGPDFNRLVSAFEELATGV
 KAIRTGLDEAKPWYKLIK
 LSRLSCMAAVALSKDPVLVAIMLADTGLEILDSTFVVKI
 SDSLSSLFHVPAVPSFGA
 PILLAGLVKVASSFFRSTPEDLERAQKQKARDINDIFAIL
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 WIASEEKFTVMTDLVPGILEKQRDNDPSKYKEAKEWLDSA
 RQACLKNGNVHIANLCKVV
 TPAPSKSRPEPVVCLRGKSGQGSFLANVLAQAISTHFTG
 RIDSVWYCPDPDHFHDGYN
 QQTVVVMDLGLQNPDKDFKYFAQMVSSTGFIPPMASLEDK
 GKPFNSKVIITTNLYSGF
 TPRTMVCPCDALNRRFHFDDIVSAKDGKYNKLDITKALED
 THTNPVAMFKYDCALLNGM
 AVEMKRMQDMFKPQPLQNVYQLVQEVIERVELHEKVSSH
 QIFKQISIPSQKSVLYFLI
 EKGQHEAAIEFFEGLVHDSIKEELRPLIQQTSFVKRAFRL
 KENFEIVALCLTLLANIVI
 MIRETRKRQMQMDDAVNEYIEKANITDDKTLDEAEKNPLE
 TSGVSIVGFRERTLPGHRA
 SDDVNSEPARPVVEEQPAEGPYTGPLERQKPLKVKAKLPQQ
 EGPYAGPMERQKPLKVKVK
 APVVKEGPYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKM
 VMGNTKPVLEILDGKTVAI
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 FEIKVKGQDMLSDAALMVL
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 LFAYKAATKAGYCGGAVLAKDGADTFIVGTHSAGGNGVGYC
 SCVSRSMMLKMAHIDPEP
 HHEGLIVDTRDVEERVHVMRKTCLAPTVAHGVEFNPEFGPAA
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 SKHKGDTKMTEEDKALFRRCAADYASRLHNVLTANAPLSI
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 GLPWALQGKRRGTLIDFENGTVGPEVASALELMEKRQYKFT
 CQTFCLKDEVPRMEKVRAGK

TRIVDVLPEVHILYTRMMIGRFCAQMHSNNGPQIGSAVGCN
 PDVDWQRFQTHFAQYKNVW
 DVDYSAFDANHCSDAMNIMFEEVFRTEFGFHPNAEWILKTL
 VNTEHAYENKRITVEGGMP
 SGCSATSIINTILNNIYVLYALRRHYEGVELDITYTMISYGD
 DIVVASDYDLDFEALKPHF
 KSLGQITITPADKSDKGFVLGQSITDVTFLKRHFRMDYGTGF
 YKPVMAKSTLEAILSFAARR
 GTIQEKLISVAGLAVHSGPDEYRRLFEPFQGLFEIPSYRSL
 YLRWVNAVCGDAQSL

Asia 1 Type

>tr|E9KM06|E9KM06_9PICO Polyprotein
 OS=Foot-and-mouth disease virus - type
 Asia 1
 PE=3 SV=1
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 YSRPNHNDNCWLNTILQLF
 RYVDEPFDDWVYDSPENLTLEAIRQLEEVGTGLELHEGGPPA
 LVIWNIKHLHTGVGTASR
 PSEVCMVDGTMCLADFHAGIFLKGQEHAVFACVTSNGWYA
 IDDEDYFPWTPDPSDVLVF
 VPYDQEPNGEWEKAKVQKRLKGAGQSSPATGSQNSGNTGS
 IINNYMQYQNSMDTQLG
 DNAISGGSNEGSTDTTSTHTTNTQNDWFSRLASSAFSGLF
 GALLADKKTEETTLLEDRI
 LTRNGHTTSTTQSSVGVTYGYAVAEDAVSGPNTSGLETRV
 TQAERFFKKHLFDWTPDLS
 FGHCHYLELPSEHKGVFGSLMSSYAYMRNGWDVEVTAVGNQ
 FNGGCLLVALVPELKELDT
 RQKYQLTLFPHQFINPRTNMTAHINVPYGVNRYDQYELHK
 PWTLVVMVAPLTVKTGGS
 EQIKVYMNAAPTYVHVAGELPSKEGIVPVACVDGYGNMVT
 DPKTADPVYGVSNPPTS
 FPGFRFTNFDVAEACPTFLRFGEVFPVKTVNSGDRLLAKFD
 VSLAAGHMSNTYLAGLAQY
 YTQYSGTMNIHFMFTGPTDAKARYMVAYIPPGMTPTDPER
 AAACIHEWDTGLNSKFTF
 SIPYLSAADYAYTASDVAETTSVQWVCIIYQITHGKAEGDA
 LVVSVSAGKDFEFLRPVDA
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 FVKLTQPKSTQTLDMQIP
 SHTLVGALLRSATYFSDLEVALVHTGPVTWVPNGAPKTAL
 NNHTNPTAYQKQPITRLAL
 PYTAPHRVLSTVYNGKTTYGEESSRRGDLAALARRVSNRLP
 TSFNYGAVKADTITELLIR
 MKRAETYCPRPLLALDTTQDRRKQEIIPAPEKQTLNFDLLKL
 AGDVESNPGPFFFSVRSN
 FTKLVDTINQMQEDMSTKHGPDFNRLVSAFEELATGVKAIR
 TGLDEAKPWYKLIKLLSRL
 SCMAAVALSKDPVLVAIMLADTGLEILDSTFVVKIISDSL
 SSLFHVPAVPSFGAPVLL
 AGLVKVASSFFRSTPEDLERAQKQKARDINDIFAILKNGE
 WLKILILAIRDWIKAWIAS
 EEKFTVMTDLVPGILEKQRDNDPSKYEEAKEWLDNARQAC
 LKSGNVHIANLCKVAPAP
 SKSRPEPVVCLRGKSGQGSFLANVLAQAISTHFTGRDTS
 VWYCPDPDHFHDGYNQQT
 VVMDLGLQNPDKDFKYFAQMVSSTGFIPPMASLEDKKGPF
 NSKVIIATTNLYSGFTPRT

MVCPDALNRRFHFDDIDVSAKDGKINNKLDIIKALEDTHTN
 PVAMFYQDCALLNGMAVEM
 KRMQQDMFKPQPPLQNVYQLVQEVIDRVELHEKVSSHPIFK
 QISIPSQKSVLYFLIEKGQ
 HEAAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLKENF
 EIVALCLTLLANIVIMIRE
 TRKRQQMVNDVANEYIDKANITDDDKTLDEAEKNPLETSGA
 STVGFRERTLPGRKTSDDV
 YSEPVKPVVEEQPAEGPYAGPLERQKPLEVRAKLPQQEGPY
 AGPMERQKPLKVKAKAPVV
 KEGPYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKMVMGN
 TKPVELILDGKTVAICCAT
 GVFGTAYLVPRHLFAEKYDKIMLDGRAMTDSYRVFEFEIK
 VKGQDMLSDAALMVLHRGN
 RVRDITKHFRDVAKMKKGTTPVVGVINNADVGRILFSGEALT
 YKDIVVCMGDGTMPGLFAY
 KAVTRAGYCGGAVLAKDGAETFIVGTHSAGNGVGYCSCVS
 RSMLLKMAHIDPEPHHEG
 LVVDTRDVEERVHVMRKTKLAPTVAHGVEFNPEFGPAALSNK
 DPRLNEGVLDEVIFSKHK
 GDTKMSEEDKALFRRCAADYASRLHSLGTANAPLSIYEAI
 KGVDGLDAMEPDTAPGLPW
 ALQKRRGALIDFENGTVGPEVEAALKLMEKREYKFACQTF
 LKDEIRPMEKVRAGKTRIV
 DVLPEHILYTRMMIGRFCAQMHSNNGPQIGSAVGCNPDID
 WQRFGTHFAQYRNVDVDY
 SAFDANHCSAMNIMFEEVFRTEFGFHPNAEWILKTLVNTE
 HAYENKRIVVEGGMPSGCS
 ATSIINTILNNIYVLYALRRHYEGVELDITYTMISYGDDIVV
 ASDYDLDFEALKPHFKSLG
 QTITPADKSDKGFLGHSITDVTFLKRHFHMDYGTGFYKPV
 MASKTLEAILSFAARRGTIQ
 EKLTSVAGLAVHSGPDEYRRLFEPFQGLFEIPSYRSLYLWR
 VNAVCGDA

C Type

>sp|P15072|POLG_FMDVT Genome polyprotein
 OS=Foot-and-mouth disease virus (isolate
 -/Germany/C1Oberbayen/1960 serotype C)
 PE=1 SV=2
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 YSRPNNDNCWNLNLTILQLF
 RYVDEPFDDWVYNSPENLTLEAIKQLEELTGLELREGGPPA
 LVIWNKHLHGTGIGTASR
 PSEVCMVDGTMCLADFHAGIFMKGQEHAVFACVTSNGWYA
 IDDEDYFPWTPDPSDVLVF
 VPYDQEPLNEGWKANVQRKLKGAGQSSPATGSQNQSGNTGS
 IINNYMQQYQNSMDTQLG
 DNAISGGSNEGSTDTTSTHTTNTQNDWFSKLASSAFSGLF
 GALLADKKTEETTLEDRI
 LTTRNGHTTSTTQSSVGVTFGYATAEDSTSGPNTSGLETRV
 HQAERFFKMALFDWVPSQN
 FGHMHKVVLPHEPKGVYGGVLKSYAYMRNGWDVEVTAVGNQ
 FNGGCLLVALVPEMGDISD
 REKYQLTLYPHQFINPRTNMTAHITVPYVGVNRYDQYKQHR
 PWTLVVMVVAPLTTNTAGA
 QQIKVYANIAPTNVHVAGELPSKEGIFPVACSDGYGNMVT
 DPKTADPAYGKVINPPRTA
 LPGRFTNYLDVAEACPTFLMFENVPYVSTRTDGQRLLAKFD
 VSLAAKHSNTYLAGLAQY

YTQYTGTLNLHFMFTGPTDAKARYMVAYVPPGMDAPDNPEE
 AAHCIHAEWDTGLNSKFTF
 SIPYISAADYAYTASHEAETTCVQGWVCVYQITHGKADADA
 LVVSASAGKDFELRLPVDA
 RQQTATGESADPVTTTVENYGGGTQVQRRHHTDVAFLDR
 FVKVTVSGNQHTLDVMQAH
 KDNIVGALLRAATYFSDLEIAVTHTGKLTWVPNGAPVSAL
 DNTTNPTAYHKGPLTRLAL
 PYTAPHRVLATAYTGTTTYTASTRGDSAHLTATRARHLPTS
 FNFVAVKAETITELLVRMK
 RAELYCPRPILPIQPTGDRHKQPLVAPAKQLLNFDLLKLAG
 DVESNPGPFFFFSDVRSNFS
 KLIVETINQMQEDMSTKHGPDFNRLVSAFEELASGVKAIRTG
 LDEAKPWYKLIKLLSRLSC
 MAAVAARSKDPVLVAIMLADTGLEILDSTFVVKKISDSLSS
 LFHVPAFAFSFGAPILLAG
 LVKVASSFFRSTPEDLERAQKQKARDINDIFAILKNGEWL
 VKLILAIRDWIKAWIASEE
 KFTVMTDLVPGILEKQRDNDPSKYKDAKEWLDNTRQACLK
 SGNVHIANLCKVVAAPPSK
 SRPEPVVCLRGKSGQKSFANVLAQAISTHLTGRDTSVW
 YCPDPDHFHDGYNQQTVVV
 MDDLQNPDKGDFKYFAQMVSTTGFIPPMASLEDKGKPFSS
 KVIIATTNLYSGFTPKTMV
 CPDALNRRFHFDDIDVSAKDGKINNKLDIIKALEDTHTNPV
 AMFYQDCALLNGMAVEMKR
 LQQDMFKPQPPLQNVYQLVQEVIERVELHEKVSSHPIFKQI
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 AAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLKENFEI
 VALCLTLLANIVIMIRETH
 KRQKMVDDAVNEYIEKANITDDDKTLDEAEKNPLETSGAST
 VGFRERTLPGQKARDDVNS
 EPAQPTTEEQPAEGPYAGPLERQKPLKVRKLPQQEGPYAG
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 GPYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKMVMGNTK
 PVELILDGKTVAICCATGV
 FGTAAYLVPRHLFAEKYDKIMLDGRALTDSDYRVFEFEIKVK
 GQDMLSDAALMVLHRGNRV
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 MLLKMAHIDPEPHHEGLI
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 RLNEGVLDEVIFSKHKG
 TKMSEEDKALFRRCAADYASRLHSLGTANAPLSIYEAIKG
 VDGLDAMEPDTAPGLPWAL
 QGKRRGALIDFENGTVGPEVEAALKLMEKREYKFACQTFK
 DEIRPMEKVRAGKTRIVDV
 LPVEHILYTRMMIGRFCAQMHSNNGPQIGSAVGCNPDVDWQ
 RFGTHFAQYRNVDVDYSA
 FDANHCSAMNIMFEEVFRTEFGFHPNAEWILKTLVNTEHA
 YENKRITVEGGMPSGCSAT
 SIINTILNNIYVLYALRRHYEGVELDITYTMISYGDDIVVAS
 DYDLDFEALKPHFKSLGQT
 ITPADKSDKGFLGHSITDVTFLKRHFHMDYGTGFYKPVMA
 SKTLEAILSFAARRGTIQEK
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 AVCGDA

Capsid Protein Sequences of Different Serotypes of FMDV

O Type

vp0

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LTRNGHTTSTTQSSVGVTY
YATAEDFVSGPNTSGLETRVVQAERFFKTHLFDWVTSDFS
RCHLLELPTDHKG VYGS
DSYAYMRNGWDVEVTAVGNQFNGGCLLVAMVPELYSIQKRE
LYQLTLFPHQFINPRTNMT
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IKVYANIAPTNVHVAGEFFPSKE

vp1

>sp|P03305|725-935
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LVGALLRASTYYFSDLEIAVKHEGDLTWVPNGAPEKALDNT
TNPTAYHKAPLTRLALPYT
APHRVLATVYNGECRYNRNAVPNLRGDLQVLAQKVARTLPT
SFNYGAIKATRVTELLYRM
KRAETCYCPRLLAHPTEARHKQKIVAPVKQ

vp2

>sp|P03305|287-504
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FVSGPNTSGLETRVVQAER
FFKTHLFDWVTSDFS
RCHLLELPTDHKG VYGS
LTD SYAYMRNGWDVEVTAVGNQFNGGC
LLVAMVPELYSIQKRELYQLTLFPHQFINPRTNMTAHITVP
FVGVNRYDQYKVHKPWTLLV
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vp3

>sp|P03305|505-724
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SGTINLHFMFTGPTDAKAR
YMVAYAPPGMEPPKTPAAAAHCIAEWDGTLNSKFTFSIPY
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QGWVCLFQITHGKADGDALVVLASAGKDFELRLPVDARAE

vp4

>sp|P03305|202-286
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A Type

vp0

>sp|P49303|202-504
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LTRNGHTTSTTQSSVGVTY
YSTQEDHVS
GPNTSGLETRVVQAERFFKKYLFDWTPDKAFG
HLEKLELPTDHKG VYGH

DSFAYMRNGWDVEVSAVGNQFNGGCLLVAMVPEWKELTPRE
KYQLTLFPHQFISPRNTMT
AHIVVPYLGVNRYDQYKKHKPWTLLVVMVVSPLTTNTVSAGQ
IKVYANIAPTHVHVAGELP
SKE

vp1

>sp|P49303|726-936
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LVGALLRAATYYFSDLEIVVRHDGNLTWVPNGAPEAALS
NMGNPTAYPKAPFTRLALPYT
APHRVLATVYNGTGKYSAGGMGRRGDLEPLAARVAAQLPTS
FNFGAIQATTIHELLVRMK
RAELYCPRPLLAVEVSSQDRHKQKIAPAKQ

vp2

>sp|P49303|287-504
DKKTEETTLLLEDRI
LTRNGHTTSTTQSSVGVTYGYSTQED
HVS
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LVDSFAYMRNGWDVEVSAVGNQFNGGC
LLVAMVPEWKELTPREKYQLTLFPHQFISPRNTMTAHIVVP
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VMVVSPLTTNTVSAGQIKVYANIAPTHVHVAGELPSKE

vp3

>sp|P49303|505-725
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SGTINLHFMFTGSTD
SKARYMVAYVPPGVETPPDTPKAAHCIAEWDGTLNSKFTFSIP
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vp4

>sp|P49303|202-286
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Asia 1 Type

vp1

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TNPTAYQKQPITRLALPYT
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FNYGAVKADTITELLIRMKR
AETCYCPRLALDTTQDRRKQEIIAPEKQ

vp2

>tr|E9KM06|287-504
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FFKKHLDWTPDLSFGHCHYLELPSEHKGVFGSLMSSYAYM
RNGWDVEVTAVGNQFNGGC
LLVALVPELKELDTRQKYQLTLFPHQFINPRTNMTAHINVP
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vp3

>tr|E9KM06|505-723

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GTMNIHFMFTGPTDAKARY
MVAYIPPGMTPPTDPERAAHCIIHSEWDTGLNSKFTFSIPYL
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vp4

>tr|E9KM06|202-286
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C Type**vp0**

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LTRNGQTTSTTQSSVGVTFG
YATAEDSTSGPNTSGLETRVHQAERFFKMAFDWVPSQNF
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KYQLTLYPHQFINPRTNMT
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IKVYANIAPTNVHVAGELP
SKE

vp1

>sp|P15072|724-930
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TNPTAYHKGPLTRLALPYT
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LYCPRPILPIQPTGDRHKQPLVAPAKQ

vp2

>sp|P15072|287-504
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RNGWDVEVTAVGNQFNGGC
LQAALVPEMGDISDREKYQLTLYPHQFINPRTNMTAHITVP
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vp3

>sp|P15072|505-723
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GTINLHFMFTGPTDAKARY
MVAYVPPGMADPNPEEAHCIIHAEWDTGLNSKFTFSIPYI
SAADYAYTASHEAETTCVQ
GWVCVYQITHGKADADALVVSASAGKDFELRLPVDARQQ

vp4

>sp|P15072|202-286
GAGQSSPATGSQNSGNTGSIINNYMQQYQNSMDTQLGDN
AISGGSNEGSTDTTSTHTT
NTQNNDFWFSKLASSAFSGLFGALLA

Genome Polyprotein Sequences of Different Serotypes of FMDV

O Type

>sp|P03305|POLG_FMDVO Genome polyprotein
OS=Foot-and-mouth disease virus (isolate
Bovine/Germany/O1Kaufbeuren/1966 serotype
O) PE=1 SV=1
MNTTDCFIALVQAIKALFLSRTTGKMELTLYNGEKKTF
YSRPNNDNCWLNAILQLF
RYVEEPFFDWWYSSPENLTLEAIKQLEDLTGLELHEGGPPA
LVIWNIKHLHTGIGTASR
PSEVCMVDGTDMLADFHAGIFLKGQEHAVFACVTSNGWYA
IDDEDFYPWTPDPDVLVF
VPYDQEPNGEWKAKVQRKLKGAGQSSPATGSQNSGNTGS
IINNYMQQYQNSMDTQLG
DNAISGGSNEGSTDTTSTHTTNTQNNDFWFSKLASSAFSGLF
GALLADKKTEETTLLEDRI
LTTNRGHTTSTTQSSVGVTYGYATAEDFVSGPNTSGLETRV
VQAERFFKTHLFDWVTS
FGRCHLLELPTDHKGVYGS LTD SYAYMRNGWDVEVTAVGNQ
FNGGCLLVAMVPELYSIQK
RELYQLTLFPHQFINPRTNMTAHITVPFVGVNRYDQYKVHK
PWTLVVMVAPLTVNTEGA
PQIKVYANIAPTNVHVAGEFSPKEGIFPVACSDGYGGLVTT
DPKTADPVYGVFNPPRNQ
LPGRFTNLLDVAEACPTFLRFEGGVPIVTTKTDSRVLQAQF
DMSLAAKQMSNTFLAGLAQ
YYTQYSGTINLHFMFTGPTDAKARYMVAYAPPGMEPPKTP
AAAHCIHAEWDTGLNSKFT
FSIPYLSAADYAYTASGVAETTNVQGWVCLFQITHGKADGD
ALVVLASAGKDFELRLPVD
ARAETTSAGESADPVTTTVENYGGGETQIQRRQHTDVSFIMD
RFVKVTPQNQINILDLMI
PSHTLVGALLRASTYFSDLEIAVKHEGDLTWVPNGAPEKA
LDNTTNPTAYHKAPLTRLA
LPYTAPHRVLATVYNGECRYNRNAVPNLRGDLQVLAQKVAR
TLPTSFNYGAIKATRVTEL
LYRMKRAETVCPRLLAHPTEARHKQKIVAPVKQTLNFDL
LKLADVESNPGPFFFSV
RSNFSKLVTETINQMQEDMSTKHGPDFNRLVSAFEELAIGVK
AIRTGLDEAKPWYKLIKLL
SRLSCMAA VAARS KDPVLVAIMLADTGLEILDSTFVVKIS
DSLSSLFHVPAVPVFSFGAP
VLLAGLVKVASSFFRSTPEDLERAEKQLKARDINDIFAILK
NGEWLVKLILAIRDWIKAW
IASEEKFVTMTDLVPGILEKQRDLNDPSKYKEAKEWLDNAR
QACLSGNVHIANLCKVVA
PAPSKSRPEPVVCLRGKSGQKGSFLANVLAQAISTHTGR
IDSVWYCPPDPDHFHDGYNQ
QTVVVMDDLQGNPDGKDFKYFAQMSTTGFIPPMASLEDKG
KPFNSKVIIATTNLYSGFT
PRTMVCPDALNRRFHFIDIVSAKDGKINSKLDIIKALED
HANPVAMFQYDCALLNGMA
VEMKRMQQDMFKPQPPLQNVYQLVQEVIDRVELHEKVSSHP
IFKQISIPSQKSVLYFLIE
KGQHEAAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLK
ENFEIVALCLTLLANIVIM
IRETRKRQKMVDDAVNEYIEKANITDDDKTLDEAEKSPLET

SGASTVGFRERTLPGQKAC
 DDVNSEPAQPVEEQPQAEQPYAGPLERQKPLKVRAKLPQQE
 GPYAGPMERQKPLKVKAKA
 PVVKEGPEYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKMV
 MGNTKPVLELILDGKTVAIC
 CATGVFGTAYLVPRHLFAEKYDKIMVDGRAMTDSYRVFEF
 EIKVKGQDMLSDAALMVLH
 RGNRVRDITKHFRDTARMKKGTPVVGVINNADVGRILFSGE
 ALTYKDIVVCMGDGTMPLG
 FAYRAATKAGYCGGAVLAKDGADTFIVGTHSAGGNGVGYCS
 CVSRSMMLKMKAHIDPEPH
 HEGLIVDTRDVEERVHVMRKTCLAPTVAHG VFNPEFGPAAL
 SNKDPRLNEGVLDEVIFS
 KHKGDTKMSEEDKALFRRCAADYASRLHSLVLTANAPLSIY
 EAIKGV DGLDAMEPDTAPG
 LPWALQGKRRGALIDFENGTVGPEVEAALKLMEKREYKFC
 QTFCLKDEIRPLEKVRAGKT
 RIVDVLPVEHILYTRMMIGRFAQMHNNGPQIGSAVGCNP
 DVDWQRF GTHFAQYRNVWD
 VDYSAFDANHCSDAMNIMFEEVFRTEFGFHPNAEWILKTLV
 NTEHAYENKRITVGGGMP
 GCSATSIINTILNNIYVLYALRRHYEGVELDTYTMISYGDD
 IVVASDYDLDFEALKPHFK
 SLGQTITPADKSDKGFLVGHSTIDVTFLKRHFHMDYGTGFY
 KPVMSKTLAAILSFARRG
 TIQEK LISVAGLAVHSGPDEYRRLFEFPQGLFEIPSYRSLY
 LRWVNAVCGDA

A type

>sp|P49303|POLG_FMDVZ Genome polyprotein
 OS=Foot-and-mouth disease virus (isolate
 -/Azerbaijan/A22-550/1965 serotype A)
 PE=1 SV=1
 MNTTDCFIALLYALREIKAFLLSRTQGMELTLYNGEKKTF
 YSRPNNHDCNLNTILQLF
 RYVDEPFFDWVYDSPENLTCEAIRQLEEITGLELHEGGPPA
 LVIWNKHLHTGIGTASR
 PSEVCMVDGTMCLADFHAGIFLKGQEHAVFACVTS DGWYA
 IDDEDYFPWTPDPSDVLVF
 VPYDQEP LNGEWAQKQKRLKGAGQSSPATGSQNSGNTGS
 IINYYMQQYQNSMDTQLG
 DNAISGGSNEGSDTSTHTTNTQNDWFSKLASSAFSGLF
 GALLADKKTEETLLEDRI
 LTTRNGHTTSTTQSSVGVTYGYSTQEDHVS GPNTSGLETRV
 VQAERFFKKYLFWDWTPDKA
 FGHLEKLELPTDHKG VYGHVLSDFAYMRNGWDVEVSAVGNQ
 FNGGCLLVAMVPEWKELTP
 REKYQLTLFPHQFISPRTNMTAHIVVPYLG VNRIDQYKKHK
 PWTLVVMVVSPLTNTTVSA
 GQIKVYANIAPTHVHVAGELPSKEGIVPVACSDGYGGLVTT
 DPKTADPVYGMVYNPPRTN
 YPGRFTNLLDVAEACPTFLCFDDGKPYVVTTRTDEQRLLAKF
 DLSLAAKHMSNTYLSGIAQ
 YYAQYSGTINLHFMFTGSTDSKARYMVAYVPPGVETPPDTP
 EKAACIHAEDWTGLNSKF
 TFSIPYVSAADYAYTASDVAETTNNVQGWVCIYQITHGKAEQ
 DTLVSVSAGKDFELRLPI
 DPRSQTTSTGESADPVTTTVENYGGGETQVQRRQHTDVTFIM
 DRFVKIQNLNPIHVIDLMQ
 THQHGLV GALLRAATYFSDLEIVVRHDGNLTWVPNGAPEA

ALSNMGNPTAYPKAPFTRL
 ALPYTAPHRVLATVYNGTGKYSAGGMGRRGDLEPLAARVAA
 QLPTSFNFGAIQATTIHEL
 LVRMKRAELYCPRLLAVEVSSQDRHKQKI IAPAKQLLNFD
 LLKLAGDVESNPGPFFFS
 VRSNFSKLIVETINQM QEDMSTKHGPDFNRLVSAFEELATGV
 KAIRTGLDEAKPWYKLIK
 LSRLSCMAAVAARSKDPVLVAIMLADTGLEILDSTFVVKKI
 SDSLSSLFHVPAVPVFSFGA
 PILLAGLVKVASFFRSTPEDLERAEKQLKARDINDIFAIL
 KNGEWLVKLILAIRDWIKA
 WIASEEKFVTMTDLVPGILEKQRD LNDPSKYKEAKEWLDSA
 RQACLKNGNVHIANLCKVV
 TPAPSKSRPEPVVCLRGKSGQGSFLANVLAQAISTHFTG
 RIDSVWYCPDPDHFHDGYN
 QQTVVVMDLGLQNPDKGDFKYFAQMVSTTGFI PPMAILEDK
 GKPFNSKVIITTTNLYSGF
 TPRTMVC PDALNRRHFHDIDVSAKDGYKVNKL DITKALED
 THTNPVAMEFYDCALLNGM
 AVEMKRMQQDMFKPQPPLQNVYQLVQEVIERVELHEKVSSH
 QIFKQISIPSQKSVLYFLI
 EKGQHEAAIEFFEGLVHDSIKEELRPLIQQTSFVKRAFRL
 KENFEIVALCLTLLANIVI
 MIRETRKRQQMVDDAVNEYIEKANITDDKTLDEAEKNPLE
 TSGVSIVGFRERTLPGHRA
 SDDVNSEPARPVVEEQPQAEQPYTGPLERQKPLKVKAKLPQQ
 EGPYAGPMERQKPLKVKVK
 APVVKEGPEYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKM
 VMGNTKPVLELILDGKTVAI
 CCATGVFGTAYLVPRHLFAEKYDKIMLDGRAMTDSYRVFE
 FEIKVKGQDMLSDAALMVL
 HRGNRVRDITKHFRDTARMKKGTPVVG VINNADVGRILFSG
 EALTYKDIVVCMGDGTMPLG
 LFAYKAATKAGYCGGAVLAKDGADTFIVGTHSAGGNGVGYC
 SCVSRSMMLKMKAHIDPEP
 HHEGLIVDTRDVEERVHVMRKTCLAPTVAHG VFNPEFGPAA
 LSNKDPRLNEGVLDEVIF
 SKHKGDTKMTEEDKALFRRCAADYASRLHNVLTANAPLSI
 YEAIKGV DGLDAMEPDTAP
 GLPWALQGKRRGTLIDFENGTVGPEVASALELMEKRQYKFT
 CQTFCLKDEV RPMEKVRAGK
 TRIVDVLPVEHILYTRMMIGRFAQMHNNGPQIGSAVGCN
 PDVDWQRF GTHFAQYKNVW
 DVDYSAFDANHCSDAMNIMFEEVFRTEFGFHPNAEWILKTL
 VNTAHAYENKRITVEGGMP
 SGCSATSIINTILNNIYVLYALRRHYEGVELDTYTMISYGD
 DIVVASDYDLDFEALKPHF
 KSLGQTITPADKSDKGFLVQGSITDVTFLKRHFHMDYGTGF
 YKPVMSKTLAAILSFARR
 GTIQEK LISVAGLAVHSGPDEYRRLFEFPQGLFEIPSYRSL
 YLRWVNAVCGDAQSL

Asia 1 Type

>tr|E9KMQ6|E9KMQ6_9PICO Polyprotein
 OS=Foot-and-mouth disease virus - type
 Asia 1
 PE=3 SV=1
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 YSRPNNHDCNLNTILQLF
 RYVDEPFFDWVYDSPENLTLEAIRQLEEVTGLELHEGGPPA

LVIWNIKHLHTGVGTASR
 PSEVCMVDGTMCLADFHAGIFLKGQEHAVFACVTSNGWYA
 IDDEDYFPWTPDPSDVLVF
 VPYDQEPLNGEWKAKVQKRLKGAGQSSPATGSQNQSGNTGS
 IINNYMQQYQNSMDTQLG
 DNAISGGSNEGSTDTTSTHTNNTQNDWFSRLASSAFSGLF
 GALLADKKTEETTLLDRI
 LTTRNGHTTSTTQSSVGVTYGYAVAEDAVSGPNTSGLETRV
 TQAERFFKKHLFDWTPDLS
 FGHCHYLELPSEHKGVFGSLMSSYAYMRNGWDVEVTAVGNQ
 FNGGCLLVALVPELKELDT
 RQKYQLTLFPHQFINPRTNMTAHINVPYVGVNRYDQYELHK
 PWTLVVMVAPLTVKTTGS
 EQIKVYMNAAPTYVHVAGELPSKEGIVPVACVDGYGNMVT
 DPKTADPVYGVKVSNPPTS
 FPGFRFTNFDVAEACPTFLRFGEVFPVKTVNSGDRLLAKFD
 VSLAAGHMSNTYLAGLAQY
 YTQYSGTMNIHFMFTGPTDAKARYMVAYIPPGMTPTDPER
 AAHCIIHSEWDTGLNSKFTF
 SIPYLSAADYAYTASDVAETTSVQGWVCIYQITHGKAEGDA
 LVVSASAGKDFEFLRPVDA
 RQQTSTTTGESADPVTSTVENYGETQTARRLHTDVAFLDR
 FVKLTQPKSTQTLDMQIP
 SHTLVGALLRSATYFSDLEVALVHTGPTVWPNGAPKTAL
 NNHTNPTAYQKQPIRLAL
 PYTAPHRVLSTVYNGKTTYGEESSRRGDLAALARRVSNRLP
 TSFNYGAVKADTITELLIR
 MKRAETYCPRLALDTTQDRRKQEI I APEKQTLNFDLLKL
 AGDVESNPGPFFFSVDRSN
 FTKLVDITINQMQEDMSTKHGPDFNRLVSAFEELATGVKAIR
 TGLDEAKPWYKLIKLLSRL
 SCMAAVAARSKDPVLVAIMLADTGLEILDSTFVVKKISDSL
 SSLFHVPAFVFSFGAPVLL
 AGLVKVASSFFRSTPEDLERAEKQLKARDINDIFAILKNGE
 WLVLKILAIRDWIKAWIAS
 EEKFVTMTDLVPGILEKQRDLDNPSKYEEAKEWLDNARQAC
 LKSGNVHIANLCKVAPAP
 SKSRPEPVVCLRGKSGQGSFLANVLAQAISTHFTGRDTS
 VWYCPDPDHFHDGYNQQT
 VVMDLQGNPDGKDFKYFAQMSTTGFIIPPMASLEDKKGPF
 NSKVIIATTNLYSGFTPT
 MCVCPDALNRRFHFIDIVSAKDGKINNKLDIIKALEDHTN
 PVAMFYQYDCALLNGMAVEM
 KRMQDMFKPQPPLQNVYQLVQEVIDRVELHEKVSSHPIFK
 QISIPSQKSVLYFLIEKGQ
 HEAAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLKENF
 EIVALCLTLLANIVIMIRE
 TRKRQQMVNDVAVNEYIDKANITDDKTLDEAEKNPLETSGA
 STVGFRERTLPGRKTSDDV
 YSEPVKPVVEEQPAEGPYAGPLERQKPLEVRAKLPQQEGPY
 AGPMERQKPLKVKAAPV
 KEGPYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKMVMGN
 TKPVELILDGKTVAICCAT
 GVFGTAYLVPRHLFAEKYDKIMLDGRAMTDSYRVFEFEIK
 VKGQDMLSDAALMVLHRGN
 RVRDITKHFRDVAKMKGTVPVGVINNADVGRILFSGEALT
 YKDIVVCMGDGTMPLGFAY
 KAVTRAGYCGGAVLAKDGAETFIVGTHSAGNGVGVCSCVS
 RSMMLKMKAHIDPEPHHEG
 LVVDTRDVEERVHVMRKTKLAPTVAHGTVFNPEFGPAALSNK

DPRLNEGVLDEVIFSKHK
 GDTKMSEEDKALFRRCAADYASRLHSVLGTANAPLSIYEAI
 KGVDGLDAMEPDTAPGLPW
 ALQKRRGALIDFENGTVGPEVEAALKMEKREYKFACQTF
 LKDEIRPMEKVRAGKTRIV
 DVLPVEHILYTRMMIGRFCAQMHSNNGPQIGSAVGCNPDI
 WQRFGTHFAQYRNVDVDY
 SAFDANHCS DAMNIMFEEVFRTEFGFHPNAEWILKTLVNTE
 HAYENKRIVVEGGMPSGCS
 ATSIINTILNNIYVLYALRRHYEGVELDTYTMISYGDDIVV
 ASDYDLDFEALKPHFKSLG
 QTITPADKSDKGFVLGHSITDVTFLKRHFHMDYGTGFYKPV
 MASKTLEAILSFAARRGTIQ
 EKLTSVAGLAVHSGPDEYRRLFEFPQGLFEIPSYRSLYLRW
 VNAVCGDA

C Type

>sp|P15072|POLG_FMDVT Genome polyprotein
 OS=Foot-and-mouth disease virus (isolate
 -/Germany/C1Oberbayen/1960 serotype C)
 PE=1 SV=2
 MNTTDCFIADVNAIREIRALFLPRTTGKMEFTLHDGEKKVF
 YSRPNNDNCWLNLTILQLF
 RYVDEPFDFWVYNPENLTLEAIKQLEELTGLELREGGPPA
 LVIWNIKHLHTGIGTASR
 PSEVCMVDGTMCLADFHAGIFMKGQEHAVFACVTSNGWYA
 IDDEDYFPWTPDPSDVLVF
 VPYDQEPLNEGWKANVQKRLKGAGQSSPATGSQNQSGNTGS
 IINNYMQQYQNSMDTQLG
 DNAISGGSNEGSTDTTSTHTTNTQNDWFSKLASSAFSGLF
 GALLADKKTEETTLLDRI
 LTTRNGHTTSTTQSSVGVTFGYATAEDSTSGPNTSGLETRV
 HQAERFFKMALFDWVPSQN
 FGHMHKVLPHEPKGVYGLVKSAYMRNGWDVEVTAVGNQ
 FNGGCLLVALVPEMGDISD
 REKYQLTLFPHQFINPRTNMTAHITVPYVGVNRYDQYKQHR
 PWTLVVMVAPLTTNTAGA
 QQIKVYANIAPTNVHVAGELPSKEGIFPVACSDGYGNMVT
 DPKTADPAYGVKNPPTA
 LPGRFTNYLDVAEACPTFLMFENVYPVSTRTDGQRLAKFD
 VSLAAGHMSNTYLAGLAQY
 YTQYTGTLNLHFMFTGPTDAKARYMVAYVPPGMDAPDNPEE
 AAHCIIHSEWDTGLNSKFTF
 SIPYLSAADYAYTASHEAETTCVQGWVCYQITHGKADADA
 LVVSASAGKDFELRLPVDA
 RQQTATGESADPVTSTVENYGETQVQRRHHTDVAFLDR
 FVKVTVSGNQHTLDVMQAH
 KDNIVGALLRAATYFSDLEIAVTHTGKLTWVPNGAPVSAL
 DNTTNPTAYHKGPLTRLAL
 PYTAPHRVLATAYTGTTTYTASTRGDSAHLTATRARHLPTS
 FNFVAVKAETITELLVRMK
 RAEVLCPRPILPIQPTGDRHKQPLVAPAKQLLNFDLLKLAG
 DVESNPGPFFFSVDRSNFS
 KLVETINQMQEDMSTKHGPDFNRLVSAFEELASGVKAIRTG
 LDEAKPWYKLIKLLSRLSC
 MAAVAARSKDPVLVAIMLADTGLEILDSTFVVKKISDSLSS
 LFHVPAFAFSFGAPILLAG
 LVKVASSFFRSTPEDLERAEKQLKARDINDIFAILKNGEWL
 VKLILAIRDWIKAWIASEE
 KFTVTMTDLVPGILEKQRDLDNPSKYKDAKEWLDNTRQACLK

SGNVHIANLCKVVPAPPSK
 SRPEPVVCLRGKSGQGKSFLANVLAQAISTHLTGRTDSVW
 YCPDPDHFHDGYNQQTVVV
 MDDLQONPDGKDFKYFAQMVSSTGFIIPMASLEDKKGPFSS
 KVIIATTNLYSGFTPKTMV
 CPDALNRRFHFDDIVSAKDGYKINNKLDIKALEDTHTNPV
 AMFQYDCALLNGMAVEMKR
 LQQDMFKPQPPLQNVYQLVQEVIERVELHEKVSSHPIFKQI
 SIPSQKSVLYFLIEKGQHE
 AAIEFFEGMVHDSIKEELRPLIQQTSFVKRAFKRLKENFEI
 VALCLTLLANIVIMIRETH
 KRQKMVDDAVNEYIEKANITDDKTLDEAEKNPLETSGAST
 VGFRERTLPGQKARDDVNS
 EPAQPTTEEQPQAEQPYAGPLERQRPVKRAKLPQQEGPYAG
 PMERQKPLKVKARAPVVKE
 GPYEGPVKKPVALKVKAKNLIVTESGAPPTDLQKMVMGNTK
 PVELILDGKTVAICCATGV
 FGTAYLVPRHLFAEKYDKIMLDGRALTDSDYRVFEFEIKVK
 GQDMLSDAALMVLHGRNRV
 RDITKHFRDVARMKKGTPVVGVINNADVGRLIFSGEALTYK
 DIVVCMGDGTMPLFAYKA
 ATKAGYCGGAVLAKDGADTFIVGTHSAGNGVGVCSCVSR
 MLLKMKAHIDPEPHHEGLI
 VDTRDVEERVHVMRKTCLAPTVAHGVEFNPEFGPAALSNDP
 RLNEGVLDEVIFSKHKG
 TKMSEEDKALFRRCAADYASRLHSLVLTANAPLSIYEAIK
 VDGLDAMEPDTAPGLPWAL
 QGKRRGALIDFENGTVGPEVEAALKLMEKREYKFACQTFK
 DEIRPMEKVRAGKTRIVDV
 LPVEHILYTRMMIGRFCAQMHSNNGPQIGSAVGCNPDVDWQ
 RFGTHFAQYRNVDVDYSA
 FDANHCS DAMNIMFEEVFRTEFGFHPNAEWILKTLVNTEHA
 YENKRITVEGGMPSGCSAT
 SIINTILNNIYVLYALRRHYEGVELDTYTMISYGDDIVVAS
 DYDLDFEALKPHFKSLGQT
 ITPADKSDKGFLGHSITDVTFLKRHFMDYGTGFYKPVMA
 SKTLEAILSFAARRGTIQEK
 LISVAGLAVHSGPDEYRRLFEPFQGLFEIPSYRSLYLWVN
 AVCGDA

Capsid Protein Sequences of Different Serotypes of FMDV

O Type

vp0

>sp|P03305|202-504
 GAGQSSPATGSQNSGNTGSIINNYMQQYQNSMDTQLGDN
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 LTRNGHTTSTTQSSVGVTYG
 YATAEDFVSGPNTSGLETRVVQAERFFKTHLFDWVTSDFS
 GRCHLLELPTDHKGVSGLT
 DSYAYMRNGWDVEVTAVGNQFNGGCLLVAMVPELYSIQKRE
 LYQLTLFPHQFINPRNTMT
 AHITVPFVGVNRYDQYKVKHPWTLVVMVAPLTVNTEGAPQ
 IKVYANIAPTNNHVHAGEFPSKE

vp1

>sp|P03305|725-935
 TTSAGESADPVTTTVENYGGGETQIQRRQHTDVSFIMDRFVK

VTPQNQINILDLMQIPSH
 LVGALLRASTYYFSDLEIAVKHEGDLTWVPNGAPEKALDNT
 TNPTAYHKAPLTRLALPYT
 APHRVLATVYNGECRYNRNAVPNLRGDLQVLAQKVARTLPT
 SFNYGAIKATRVTELLYRM
 KRAETCYCPRLLAIHPTEARHKQKIVAPVKQ

vp2

>sp|P03305|287-504
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 FVSGPNTSGLETRVVQAER
 FFKTHLFDWVTSDFSGRCHLLELPTDHKGVSGLTDSYAYM
 RENGWDVEVTAVGNQFNGG
 CLLVAMVPELYSIQKRELYQLTLFPHQFINPRNTMTAHITVP
 FVGVNRYDQYKVKHPWTLV
 VMMVAPLTVNTEGAPQIKVYANIAPTNNHVHAGEFPSKE

vp3

>sp|P03305|505-724
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 VPYVTTKTDSDRVLAQFDMSLAAKQMSNTFLAGLAQYYTQY
 SGTINLHFMFTGPTDAKAR
 YMVAYAPPGMEPPKTPEAAAHCIHAEDWTGLNSKFTFSIPY
 LSAADYAYTASGVAETTNV
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vp4

>sp|P03305|202-286
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A Type

vp0

>sp|P49303|202-504
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 LTRNGHTTSTTQSSVGVTYG
 YSTQEDHVS
 GPNNTSGLETRVVQAERFFKKYLF
 DWTPDKAFG
 HLEKLELPTDHKGVSGLV
 DSFAYMRNGWDVEVSAVGNQFNGGCLLVAMVPEWKELTPRE
 KYQLTLFPHQFISPRNTMT
 AHIVVPYLGVNRYDQYKVKHPWTLVVMVVSPLTNTVSAGQ
 IKVYANIAPTHVHVAGELP
 SKE

vp1

>sp|P49303|726-936
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 NMGNPTAYPKAPFTRLALPYT
 APHRVLATVYNGTGKYSAGGMGRGDLEPLAARVAAQLPTS
 FNFGAIQATTIHELLVRMK
 RAELYCPRLLAVEVSSQDRHKQKIIAPAKQ

vp2

>sp|P49303|287-504
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 RENGWDVEVSAVGNQFNGG
 CLLVAMVPEWKELTPREKYQLTLFPHQFISPRNTMTAHIVVP

YLGVNRYDQYKKHKPWTLV
VMVVSPLTTNTVSAGQIKVYANIAPTHVHVAGELPSKE

vp3

>sp|P49303|505-725
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KPYVVTRTDEQRLAKFDLSLAAKHMSNTYLSGIAQYYAQY
SGTINLHFMFTGSTDASKAR
YMVAYVPPGVETPPDTPKAAHCHAEWDTGLNSKFTFSIP
YVSAADYAYTASDVAETTN
VQGWCYIYQITHGKAEQDQLVVSVSAGKDFELRLPIDPRSQ

vp4

>sp|P49303|202-286
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NTQNNDWFSKLASSAFSGLFGALLA

Asia 1 Type

vp1

>tr|E9KM06|724-932
TTTTGESADPVTTTVENYGGGETQTARRLHTDVAFLDRFVK
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TNPTAYQKQPITRLALPYT
APHRVLSTVYNGKTTYGEESRRGDLAALARRVSNRLPTSF
NYGAVKADTITELLIRMKR
AETCYCPRLALDQDQDRRQKEIIAPEKQ

vp2

>tr|E9KM06|287-504
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RNGWDVEVTAVGNQFNGGC
LLVALVPELKELDTRQKYQLTLFPHQFINPRTNMTAHINVP
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VMVVAPLTVKTGGSEQIKVYMNAAPTYVHVAGELPSKE

vp3

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PFVKTVNSGDRLLAKFDVSLAAGHMSNTYLAGLAQYYTQYS
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MVAYIPPGMTPTDPERAAHCHSEWDTGLNSKFTFSIPYL
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vp4

>tr|E9KM06|202-286
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C Type

vp0

>sp|P15072|202-504
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TRNGQTTSTTQSSVGVTFG
YATAEDSTSGPNTSGLETRVHQAERFFKMALFDWVPSQNF
HMHKVLPHEPKGVYGGGLV
KSYAYMRNGWDVEVTAVGNQFNGGCLQAALVPEMGDISDRE
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AHITVPYVGVNRYDQYKQHRPWTLVVMVVAPLTTNTAGAQQ
IKVYANIAPTNNHVAGELP
SKE

vp1

>sp|P15072|724-930
TTTTGESADPVTTTVENYGGGETQVQRRHHTDVAFLDRFVK
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IVGALLRAATYYFSDLEIAVTHTGKLTWVPNGAPVSALDNT
TNPTAYHKGPLTRLALPYT
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vp2

>sp|P15072|287-504
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RNGWDVEVTAVGNQFNGGC
LQAALVPEMGDISDREKYQLTLYPHQFINPRTNMTAHITVP
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vp3

>sp|P15072|505-723
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GTINLHFMFTGPTDAKARY
MVAYVPPGMDAPDNPEEAAHCHAEWDTGLNSKFTFSIPYI
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vp4

>sp|P15072|202-286
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