**Virologica Sinica**

**Supplementary Data**

**Coxsackievirus B3 HFMD animal models in Syrian hamster and rhesus monkey**

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**Supplementary materials**

**Supplementary Table 1.** Clinical manifestation scores of CVB3 infected Syrian hamsters.

|  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Group** | **CG** | | | | | | **IG** | | | | | | **NG** | | | | | |
| Number | 1 | 2 | 3 | 4 | 5 | 6 | 1 | 2 | 3 | 4 | 5 | 6 | 1 | 2 | 3 | 4 | 5 | 6 |
| D0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| D1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| D2 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| D3 | 0 | 0 | 0 | 0 | 0 | 0 | 10 | 10 | 5 | 2 | 5 | 10 | 10 | 10 | 10 | 5 | 5 | 0 |
| D4 | 0 | 0 | 0 | 0 | 0 | 0 | 10 | 10 | 10 | 5 | 10 | 10 | 10 | 10 | 5 | 5 | 10 | 5 |
| D5 | 0 | 0 | 0 | 0 | 0 | 0 | 5 | 5 | 5 | 10 | 10 | 5 | 5 | 5 | 10 | 5 | 10 | 5 |
| D6 | 0 | 0 | 0 | 0 | 0 | 0 | 5 | 5 | 0 | 0 | 0 | 5 | 10 | 10 | 5 | 5 | 5 | 5 |
| D7 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 2 | 5 | 2 | 2 | 2 | 5 | 5 | 5 | 0 | 0 | 0 |
| D8 | 0 | 0 | 0 | - | - | - | 2 | 2 | 0 |  |  |  | 2 | 0 | 2 |  |  |  |
| D9 | 0 | 0 | 0 | - | - | - | 0 | 2 | 0 |  |  |  | 2 | 0 | 0 |  |  |  |
| D10 | 0 | 0 | 0 | - | - | - | 0 | 0 | 0 |  |  |  | 2 | 0 | 0 |  |  |  |
| D11 | 0 | 0 | 0 | - | - | - | 0 | 0 | 0 |  |  |  | 2 | 0 | 0 |  |  |  |
| D12 | 0 | 0 | 0 | - | - | - | 0 | 0 | 0 |  |  |  | 0 | 0 | 0 |  |  |  |
| D13 | 0 | 0 | 0 | - | - | - | 0 | 0 | 0 |  |  |  | 0 | 0 | 0 |  |  |  |
| D14 | 0 | 0 | 0 | - | - | - | 0 | 0 | 0 |  |  |  | 0 | 0 | 0 |  |  |  |

Notes: The clinical manifestations of CVB3-infected Syrian hamsters were scored based on the following criteria, and the final score was computed by adding the General performance score and the Food consumption score. General performance: normal, 0; slow/quiet/alert/freedom of movement, 5; quiet/hunched back/alertness/limited movement, 10; lethargy/inability to move/no alertness, 15; Food consumption: normal, 0; decreased appetite, 2; anorexia, 5. In this table, animals with a score of "10" exhibit slow/quiet/alert/free movement, along with anorexia.

**Supplementary Table 2.** Clinical manifestation scores of CVB3 infected rhesus monkey.

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
| Time | A1 | A2 | A3 | B1 | B2 | B3 |
| D0 | 0 | 0 | 0 | 0 | 0 | 0 |
| D1 | 0 | 0 | 0 | 0 | 0 | 0 |
| D2 | 0 | 5 | 5 | 0 | 0 | 0 |
| D3 | 5 | 5 | 5 | 0 | 0 | 0 |
| D4 | 5 | 5 | 5 | 0 | 0 | 0 |
| D5 | 10 | 10 | 10 | 0 | 0 | 0 |
| D6 | 10 | 15 | 10 | 0 | 0 | 0 |
| D7 | 10 | 15 | 10 | 0 | 0 | 0 |
| D8 | 5 | 15 | 5 | 0 | 0 | 0 |
| D9 | 5 | 10 | 5 | 0 | 0 | 0 |
| D10 | 5 | 5 | 5 | 0 | 0 | 0 |
| D11 | 5 | 10 | 0 | 0 | 0 | 0 |
| D12 | 0 | 5 | 0 | 0 | 0 | 0 |
| D13 | 0 | 0 | 0 | 0 | 0 | 0 |
| D14 | 0 | 0 | 0 | 0 | 0 | 0 |

Notes: The clinical manifestations of CVB3-infected rhesus monkeys were scored based on the following criteria, and the final score was computed by adding the General performance score and the Food consumption score. General performance: normal, 0; slow/quiet/alert/freedom of movement, 5; quiet/hunched back/alertness/limited movement, 10; lethargy/inability to move/no alertness, 15; Food consumption: normal, 0; decreased appetite, 2; anorexia, 5.

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**Supplementary Fig.S1** The histopathological assay of oral mucosa, lingual tonsils, pharyngeal tonsils, and palatal tonsils from Syrian hamsters at day 7 post-CVB3 infection.

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**Supplementary Fig.S2** The viral nucleic acid profiles of 18 tissues, including those from 7 d.p.i, brain, intestine, and others, were examined. The findings revealed that all tested tissues contained a certain number of viral copies, indirectly indicating the presence of CVB3 infection in these tissues. **A** Viral load in tissue from intraperitoneal infection (IG) group; **B** Viral load in tissue from nasal drip infection (NG) group.

**Supplementary Video. S1** Control group: Syrian hamsters untreated.

**Supplementary Video. S2** Intraperitoneal group: Clinical signs of Syrian hamsters infected with 105.75 CCID50 CVB3 via intraperitoneal injection.

**Supplementary Video. S3** Nasal drop group: Clinical signs of Syrian hamsters infected with 105.75 CCID50CVB3 via nasal drip.