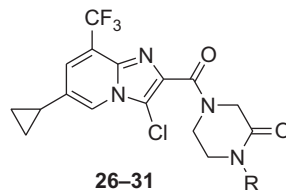


**Table 5.1** Comparison between a series of hydroxylated cyclohexanol analogs (**27–31**) and the unsubstituted derivative **26** with respect to *cLogP*, binding and replicon potency.



|              |                 | <i>EC</i> <sub>50</sub> (nM) |                  |                  |                  |           |           |
|--------------|-----------------|------------------------------|------------------|------------------|------------------|-----------|-----------|
| <i>Assay</i> | <i>Genotype</i> | <b>26</b>                    | (+/-)- <b>27</b> | (+/-)- <b>28</b> | (+/-)- <b>29</b> | <b>30</b> | <b>31</b> |
| <i>cLogP</i> |                 | 5.2                          | 4.3              | 3.3              | 3.3              | 3.3       | 3.3       |
| Replicon     | 1a              | 6.0                          | 120              | 630              | 96               | 78        | 0.8       |
| Replicon     | 1b              | 2.1                          | 50               | 540              | 42               | 120       | 5.4       |
| Binding      | 1b              | 8.3                          | 130              | 580              | 120              | 300       | 53        |

**Table 5.2** Pharmacokinetic parameters of **31** in four species.

| <i>Species</i> | <i>i.v.</i> (1 mg kg <sup>-1</sup> )               |                                  |                           |   | <i>p.o.</i> (5 mg kg <sup>-1</sup> )               |              |
|----------------|--|----------------------------------|---------------------------|---|--|--------------|
|                | <i>Cl</i> (mL min <sup>-1</sup> kg <sup>-1</sup> ) | <i>Cl</i> (% hepatic blood flow) | <i>T</i> <sub>½</sub> (h) | <i>V</i> <sub>dss</sub> (L kg <sup>-1</sup> ) | <i>AUC</i> <sub>0-∞</sub> (μg h mL <sup>-1</sup> ) | <i>F</i> (%) |
| Mouse          | 4.7  | 5                                | 1.8                       | 0.5   | 6.6  | 37           |
| Rat            | 20   | 36                               | 4.2                       | 4.1   | 3.8  | 98           |
| Dog            | 1.1  | 3                                | 13                        | 0.9   | 52   | 65           |
| Monkey         | 2.1  | 5                                | 4.9                       | 0.7   | 50   | > 100        |