## Supplementary information, Table. S3 | Effects of residue mutation in the ligand-binding pocket on GLP-2R-induced cAMP accumulation<sup>a</sup>.

Receptor mutant	Human GLP-2(1-33)	
	pEC <sub>50</sub> ±SEM	Emax (% WTb)
GLP-2R (1-553) (WT)	10.4±0.05	100.9±2.4
Y182 <sup>1.43</sup> A	9.6±0.08	101.8±3.9
Y186 <sup>1.47</sup> A	9.2±0.06**	100.3±2.8
K231 <sup>2.67</sup> A	9.9±0.07	105.1±3.0
R242 <sup>ECL1</sup> E	9.1±0.03**	108.2±1.9
R242 <sup>ECL1</sup> A	9.8±0.07	106.7±3.4
W249 <sup>ECL1</sup> A	7.2±0.7***	127.9±161.5
Y252 <sup>ECL1</sup> A	9.8±0.06	103.1±2.6
H268 <sup>3.37</sup> A	7.8±0.2***	118.4±22.6
N334 <sup>ECL2</sup> A	9.6±0.06	102.7±2.9
W340 <sup>5.36</sup> A	8.9±0.06***	106.8±3.6
R344 <sup>5.40</sup> A	9.2±0.05**	108.6±2.8
K414 <sup>7.35</sup> A	9.8±0.07	110.0±3.3

<sup>&</sup>lt;sup>a</sup>All data were fitted with a three-parameter logistic curve to obtain pEC<sub>50</sub> values. Data represent means  $\pm$  S.E.M. of at least three independent experiments performed duplicate. One-way ANOVA and Dunnett's post-test were used to determine statistical difference. \*P<0.05, \*\*P<0.01, \*\*\*P<0.001.

<sup>&</sup>lt;sup>b</sup> WT, wild-type.