



**Fig. S22: Cryo-EM map of selected residues at the MC4R-Gs protein interface.**

(a-b) Corresponding cryo-EM densities of residues highlighted in main Fig. 6d (a) and 6e (b) at a contour level of  $1.2 \sigma$ .

(c-d) Corresponding cryo-EM densities of residues highlighted in main Fig. 7a (c) and 7b (d) at a contour level of  $4 \sigma$ . For further details see legend Fig. 7.