

Supporting Information

Rotation conformational effects of selected cytotoxic cardiac glycosides on their interactions with Na⁺/K⁺-ATPase

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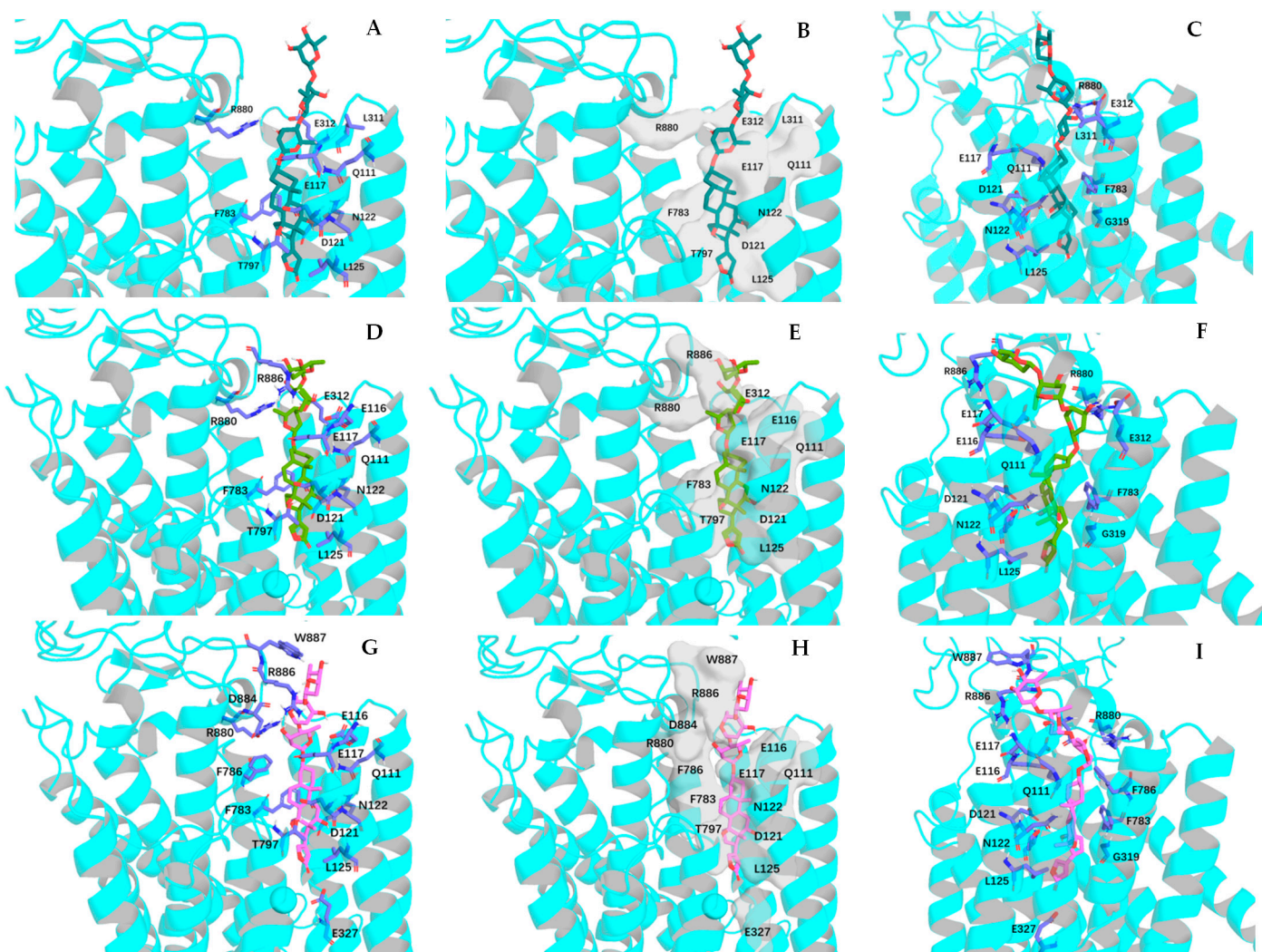


Figure S2. Docking profiles for digoxin-4RENT (deep teal) and **1e** (green) and **1f** (magenta) and NKA. A and D, overlapped docking profiles for digoxin-4RENT, **1e**, or **1f** and NKA, with no interacting residues represented; B and E, overlapped docking profiles for digoxin-4RENT, **1e**, or **1f** and NKA, with the interacting residues represented by gray

surfaces; C and F, docking profiles for **1e** or **1f** and NKA, with the interacting residues represented by slate sticks (digoxin-4RENT, deep teal; **1e**, green; **1f**, magenta).

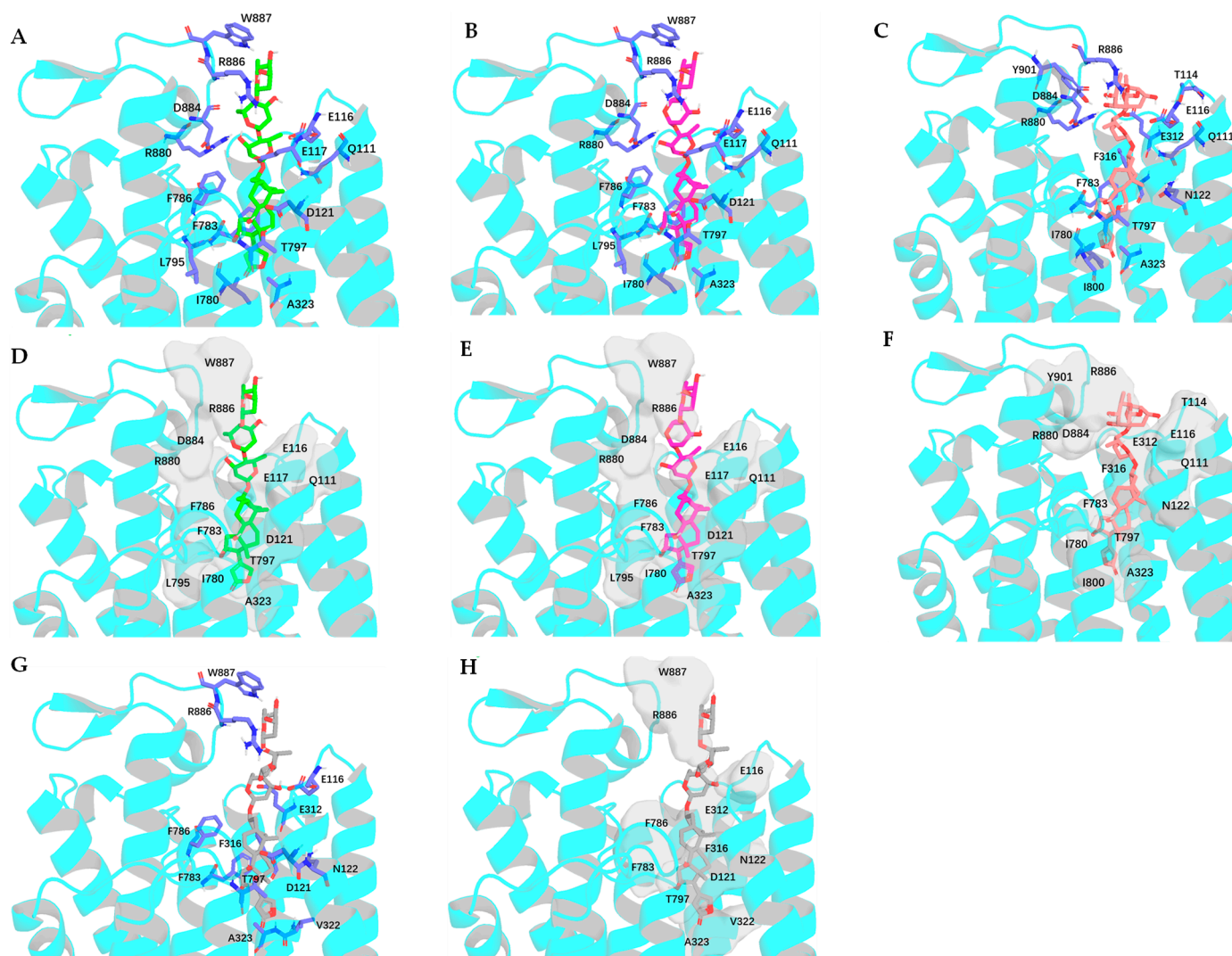


Figure S3. Docking profiles for **2a** (green), **2b** (magenta), **2c** (salmon), and **2d** (gray). A, B, C, and G, the interacting residues are represented by slate sticks; D, E, F, and H, the interacting residues are represented by gray surfaces.

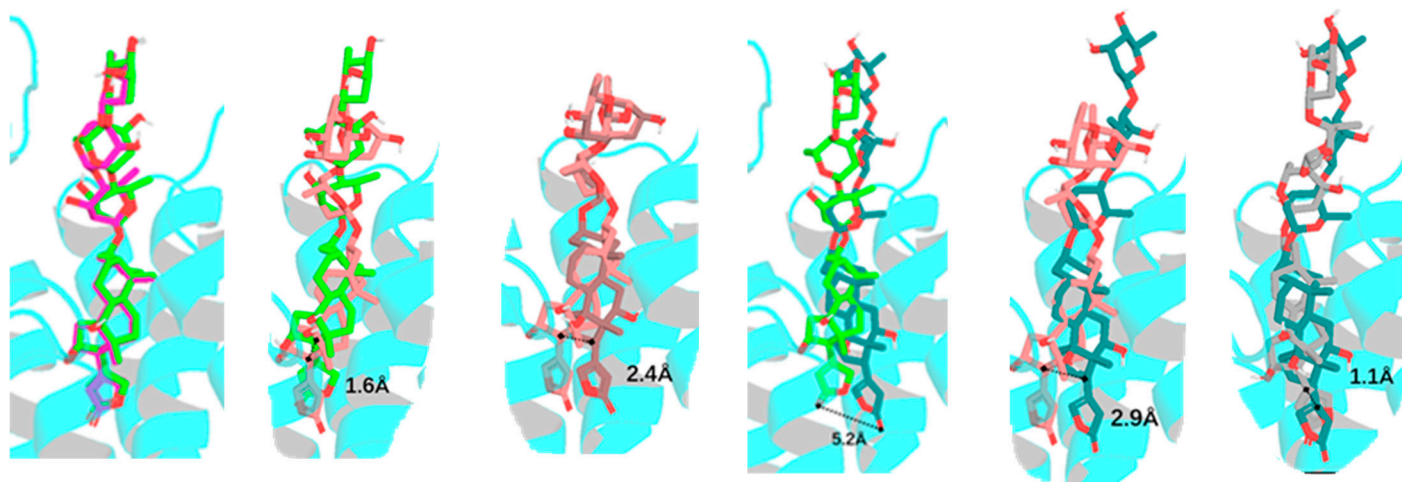
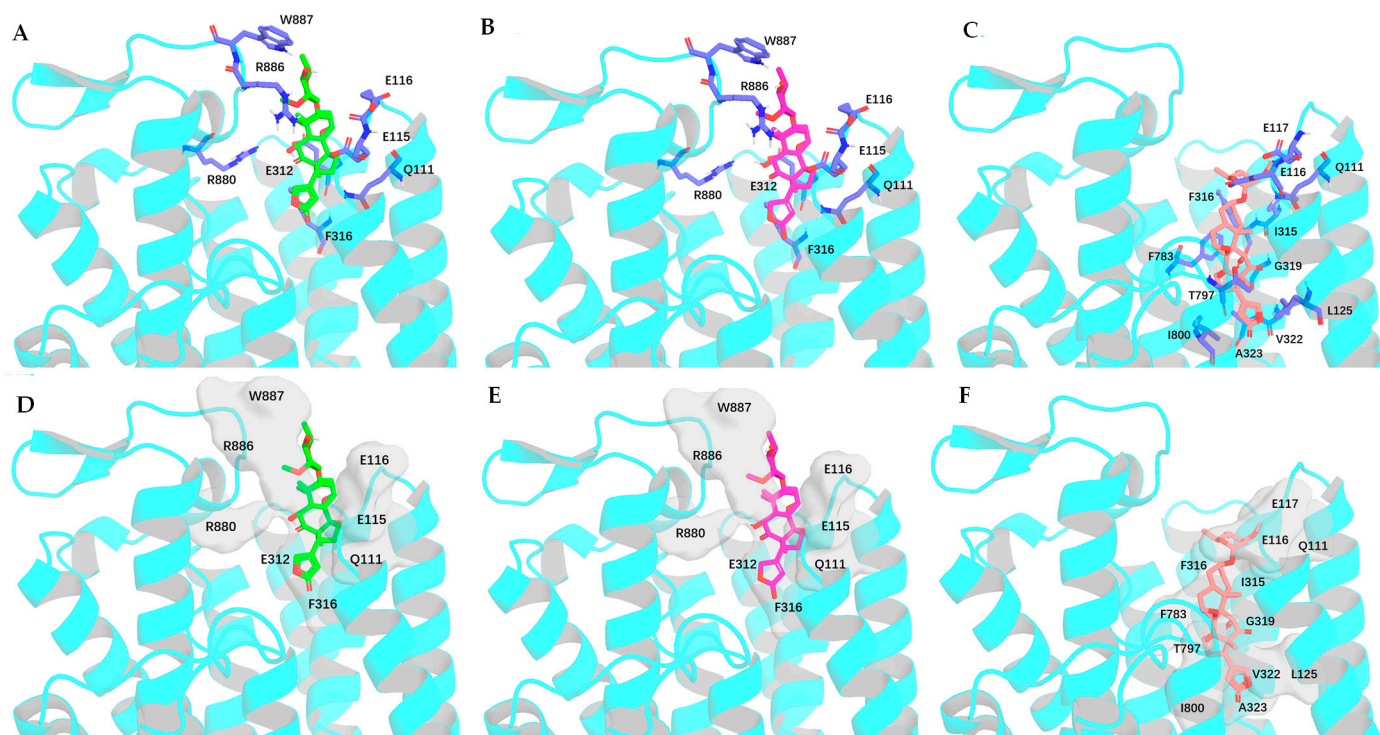


Figure S4. Overlapped docking profiles for digoxin-4RET, **1c**, or **2a–2d** and NKA (**2a**, green; **2b**, magenta; **2c**, salmon; **2d**, gray; **1c**, brown; and digoxin-4RET, deep teal).



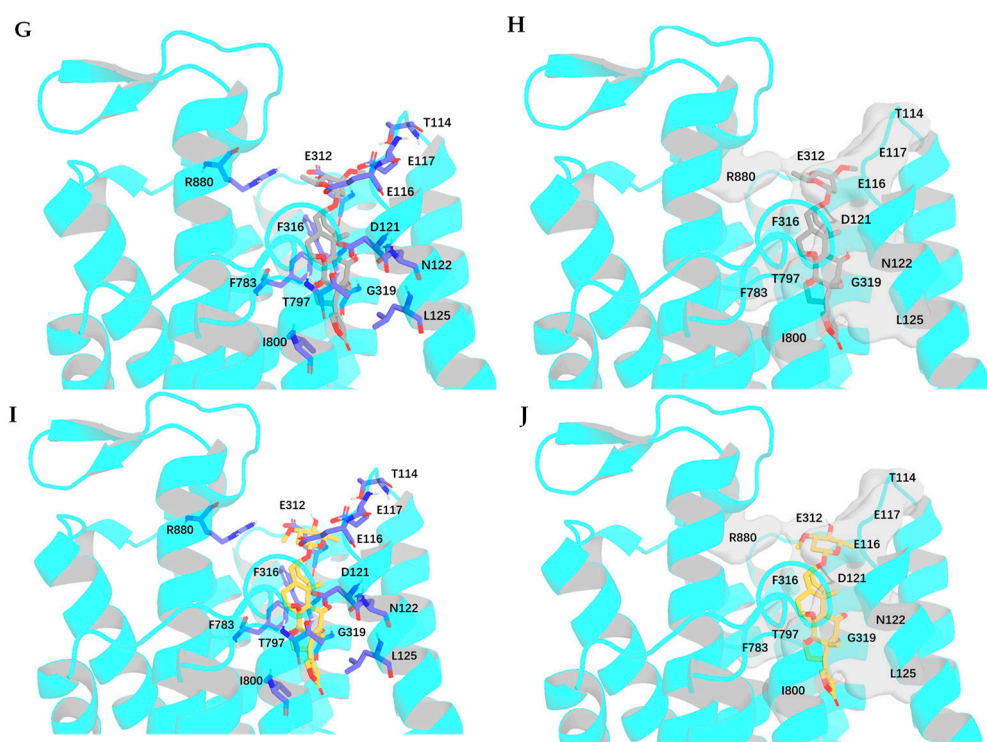


Figure S5. Docking profiles for **3a** (green), **3b** (magenta), **3c** (salmon), **3d** (gray), and **3e** (yellow). A, B, C, G, and I, the interacting residues are represented by slate sticks; D, E, F, H, and J, the interacting residues are represented by gray surfaces.