**Supplemental Table S1**. Regions of interest involved in the brain processing of itch significantly impacted by scratching.

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| *Brain area* | Itch | Activelyscratching an Itch | Passively scratching an Itch | Correlation with pleasurability of scratching#  | Correlation with itch relief induced by scratching# |
| Primary somatosensory | Activated | Activated | Activated | Inverse BA3; Z=4.8 | InverseBA1,2,3 Z=4.4 |
| Secondary somatosensory | Activated | Activated | Activated | Inverse BA40; Z=2.3 | InverseBA40; Z=3.2 |
| Primary motor | Activated | Activated | Deactivated | Inverse BA4; Z=5.7 | InverseBA4; Z=5.4 |
| Supplementary motor area | Activated | Activated |  | Inverse BA5,6,7 Z=4.3 | InverseBA5,6,7 Z=5.5 |
| **Prefrontal cortex** |  |  |  |  |  |
| Dorsolateral | Activated | Activated |  | Positive BA9,46; Z=10 Inverse BA8 Z=6.9 | InverseBA8;Z=4.8 |
| Frontal pole | Activated | Deactivated | Deactivated | Positiveextensive BA10; Z=10.0 |  |
| Orbitofrontal | Activated | Deactivated | Deactivated | PositiveBA11; Z=8.1 |  |
| Ventrolateral | Activated | Deactivated | Deactivated | Positive BA45; Z=3.2 |  |
| ACC | Activated | Activated or deactivated in different regions | Deactivated | Positive BA32,33 Z=5.7 | PositiveBA24; Z=3.5 |
| PCC | Activated | Activated | Activated | Inverse (extensive) BA23, 31 Z=5.4  | Inverse(extensive) BA23, 31 Z=9.7 |
| Precuneus | Activated | Activated | Activated | PositiveBA31; Z=8.3InverseBA7; Z=9.3 | PositiveBA7; Z=4.2InverseBA31; Z=8.8 |
| Parahippocampus | Activated | Deactivated | Activated | Positive Z=10.8 | Positive Z=9.0 |
| Hippocampus | Activated | Deactivated | Activated | Positive Z=4.7 | Inverse Z=5.2 |
| Amygdala | Activated | Deactivated |  | Inverse Z=5.4 | Inverse Z=4.6 |
| Insula | Activated | Deactivated | Deactivated |  |  |
| **Thalamus** | Activated | Activated | Deactivated |  |  |
| VPL |  |  |  | Positive Z=2.8 |  |
| VPM |  |  |  | Positive Z=2.7 | Positive Z=2.6 |
| LPN |  |  |  | Positive Z=7.2 | Positive Z=3.6 |
| LDN |  |  |  | Positive Z=7.9 | Positive Z=3.5 |
| AN |  |  |  | Positive Z=6.8 | Positive Z=6.3 |
| MD |  |  |  | Positive Z=7.4 | Positive Z=7.7 |
| Pulvinar |  |  |  | Positive Z=4.6 | Positive Z=2.7 |
| **Striatum** |  |  |  |  |  |
| Caudate |  | Activated |  | Positive(caudate head) Z=7.9 | Positive (caudate body) Z=4.2 |
| Putamen | Activated |  | Deactivated | Inverse Z=4.3 | Positive Z=4.9 |
| Nc. accumbens |  | Deactivated\* |  |  |  |
| Globus pallidus | Activated |  |  | Positive Z=6.0 | Positive Z=4.5 |
| Substantia nigra |  |  |  | Positive Z=4.4 | Positive Z=4.8 |
| Subthalamic nucleus |  |  | Activated |  |  |
| **Midbrain** |  |  |  |  |  |
| VTA |  | Deactivated\* |  | Positive Z=4.7 | Positive Z=6.0 |
| Red nucleus |  | Deactivated\* |  |  |  |
| Dorsal nucleus of the raphé |  | Deactivated\* |  | Positive Z=4.1 | Positive Z=6.5 |
| PAG |  | Deactivated\* |  | Positive Z=4.1 |  |

Notes. Brain areas correlated with itch relief and scratching pleasurability are shown. The highest Z scores > 2.3 are presented for most significantly correlated clusters.

BA = Brodmann area. ACC = anterior cingulate cortex; PCC = posterior cingulate cortex; VTA = ventral tegmental area; PAG = periaqueductal gray; VPL= ventroposterolateral nucleus; VPM = ventroposteromedial nucleus; AN=Anterior nucleus; LPN = lateral posterior nucleus; LDN = lateral dorsal nucleus; MD = Mediodorsal nucleus.

\*Deactivated in a higher-level contrast analysis in comparison to itch condition.

# Results for regression analyses performed for active scratching an itch are displayed.