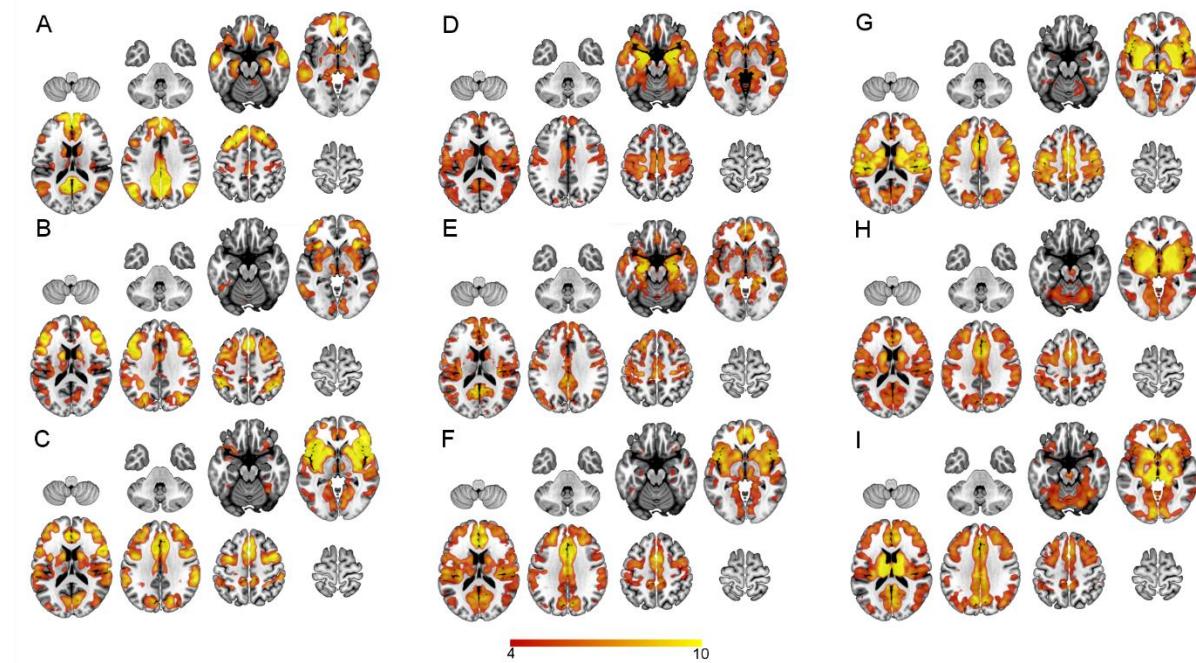
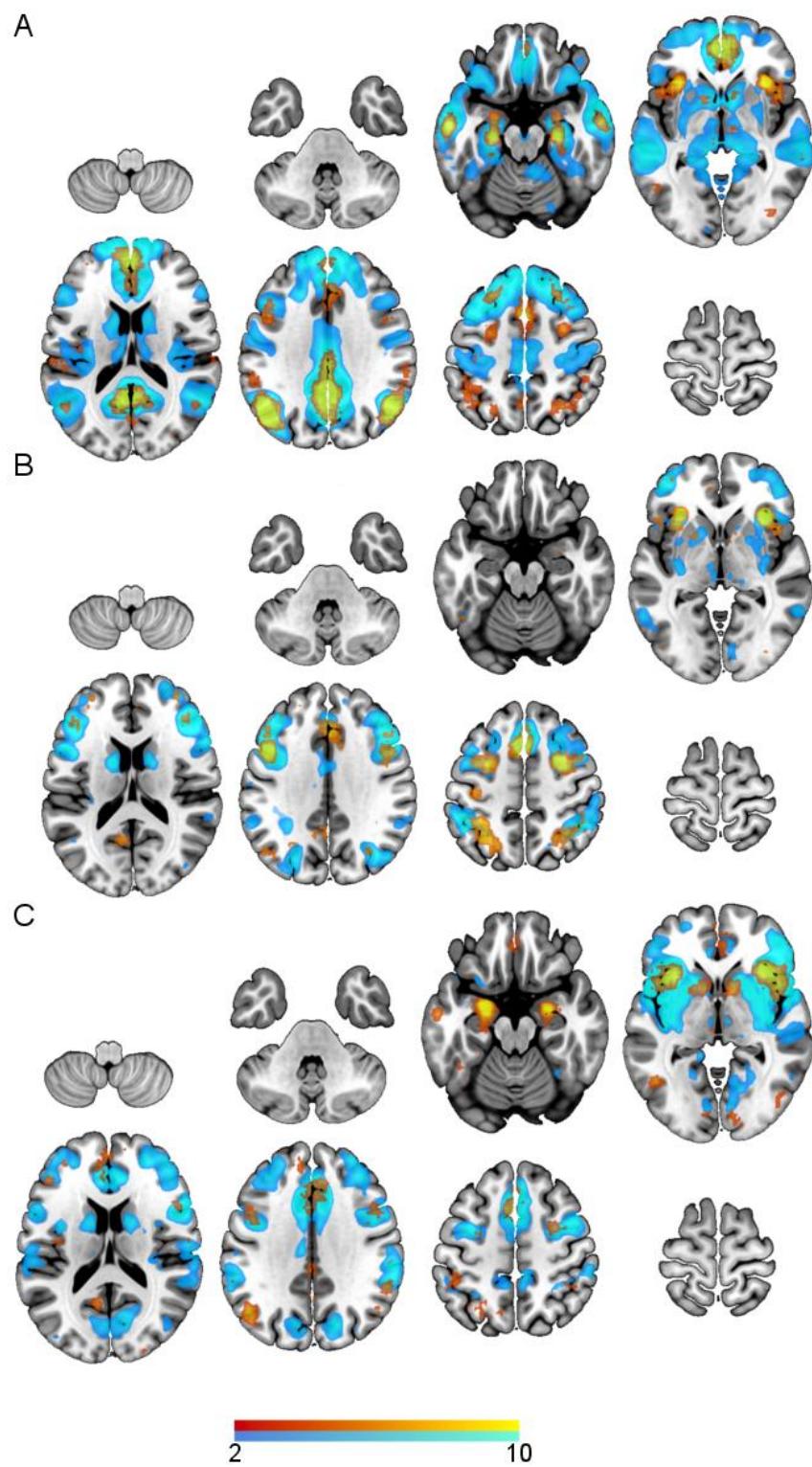


**Supplemental Figure 1.** Group mean spatial maps. Group mean for all participants ( $n=29$ ) for both visits (kisspeptin and vehicle) for the following seed-based resting state networks: (A) Default mode (DMN), (B) Executive control (ECN), (C) Salience (SAL), (D) Amygdala, (E) Hippocampus, (F) Anterior cingulate, (G) Putamen, (H) Globus pallidus, (I) Thalamus. All analyses performed using cluster-based thresholding ( $Z = 2.3$ ,  $P < 0.05$ ) for the purposes of inference. Spatial maps thresholded at  $Z = 4$  here for the purposes of clarity of display.



**Supplemental Figure 2.** Overlap between seed-based and Neurosynth networks. Group mean spatial maps for all participants ( $n=29$ ) for both visits (kisspeptin and vehicle) using seed-based analysis (in blue) overlaid on the respective networks from Neurosynth meta-analysis tool (<http://www.neurosynth.org/>; in red-yellow) for (A) Default mode (DMN), (B) Executive control (ECN) and (C) Salience (SAL). Spatial maps are thresholded for clarity of display ( $Z = 2$ ).



Supplemental Data

**Supplemental Table 1.** Correlation analyses between kisspeptin's modulation of the Default Mode Network (DMN) connectivity with psychometric outcomes. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                        | Correlation Coefficient ( $r$ ), $P$ Value |
|------------------------|--|
| BAS Drive              | -0.489, 0.008**                            |
| BAS Fun                | -0.176, 0.368                              |
| BAS Reward             | -0.079, 0.691                              |
| BIS                    | 0.109, 0.582                               |
| SADI Evaluative        | 0.002, 0.993                               |
| SADI Negative          | -0.499, 0.006**                            |
| SADI Physiological     | -0.112, 0.570                              |
| SADI Motivational      | 0.072, 0.716                               |
| Positive Affect Change | -0.072, 0.714                              |
| Negative Affect Change | 0.056, 0.777                               |

## Supplemental Data

**Supplemental Table 2.** Correlation analyses between kisspeptin's modulation of the Default Mode Network (DMN) connectivity with task brain activity in *a priori* limbic and paralimbic brain structures. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                      |  | Correlation Coefficient (r), P Value                            |
|----------------------|--|---|
| Amygdala             | Sexual<br>Bonding<br>Negative<br>Neutral | 0.273, 0.151<br>-0.181, 0.347<br>0.253, 0.185<br>0.106, 0.585   |
| Hippocampus          | Sexual<br>Bonding<br>Negative<br>Neutral | 0.275, 0.148<br>0.041, 0.832<br>0.279, 0.142<br>0.346, 0.065    |
| Thalamus             | Sexual<br>Bonding<br>Negative<br>Neutral | 0.313, 0.097<br>-0.115, 0.552<br>0.232, 0.224<br>0.219, 0.253   |
| Putamen              | Sexual<br>Bonding<br>Negative<br>Neutral | 0.406, 0.028<br>-0.063, 0.747<br>0.230, 0.229<br>0.256, 0.179   |
| Globus Pallidus      | Sexual<br>Bonding<br>Negative<br>Neutral | 0.500, 0.005**<br>-0.094, 0.628<br>0.185, 0.335<br>0.145, 0.451 |
| Anterior Cingulate   | Sexual<br>Bonding<br>Negative<br>Neutral | 0.338, 0.072<br>-0.206, 0.283<br>0.139, 0.473<br>0.113, 0.560   |
| Posterior Cingulate  | Sexual<br>Bonding<br>Negative<br>Neutral | 0.475, 0.009**<br>-0.239, 0.211<br>0.271, 0.154<br>0.139, 0.473 |
| Medial Frontal Gyrus | Sexual<br>Bonding<br>Negative<br>Neutral | 0.298, 0.115<br>-0.094, 0.628<br>0.185, 0.335<br>0.062, 0.747   |
| Nucleus Accumbens    | Sexual<br>Bonding<br>Negative<br>Neutral | 0.130, 0.500<br>-0.220, 0.251<br>-0.039, 0.841<br>-0.008, 0.967 |
| Caudate              | Sexual<br>Bonding<br>Negative<br>Neutral | 0.149, 0.441<br>-0.354, 0.059<br>-0.051, 0.794<br>-0.057, 0.771 |

Supplemental Data

**Supplemental Table 3.** Correlation analyses between kisspeptin's modulation of the Executive Control Network (ECN) connectivity with psychometric outcomes. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

| Correlation Coefficient (r), P Value |               |
|--------------------------------------|---------------|
| BAS Drive                            | 0.263, 0.177  |
| BAS Fun                              | 0.032, 0.870  |
| BAS Reward                           | 0.277, 0.153  |
| BIS                                  | -0.005, 0.980 |
| SADI Evaluative                      | 0.148, 0.453  |
| SADI Negative                        | -0.026, 0.897 |
| SADI Physiological                   | 0.047, 0.812  |
| SADI Motivational                    | 0.015, 0.938  |
| Positive Affect Change               | -0.085, 0.668 |
| Negative Affect Change               | 0.031, 0.877  |

## Supplemental Data

**Supplemental Table 4.** Correlation analyses between kisspeptin's modulation of the Executive Control Network (ECN) connectivity with task brain activity in *a priori* limbic and paralimbic brain structures. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                      |  | Correlation Coefficient (r), P Value                             |
|----------------------|--|--|
| Amygdala             | Sexual<br>Bonding<br>Negative<br>Neutral | -0.224, 0.242<br>-0.310, 0.102<br>-0.198, 0.302<br>-0.098, 0.614 |
| Hippocampus          | Sexual<br>Bonding<br>Negative<br>Neutral | -0.304, 0.109<br>-0.398, 0.032<br>-0.294, 0.121<br>-0.132, 0.495 |
| Thalamus             | Sexual<br>Bonding<br>Negative<br>Neutral | -0.286, 0.133<br>-0.325, 0.085<br>-0.339, 0.072<br>-0.204, 0.289 |
| Putamen              | Sexual<br>Bonding<br>Negative<br>Neutral | -0.229, 0.231<br>-0.335, 0.076<br>-0.306, 0.106<br>-0.355, 0.058 |
| Globus Pallidus      | Sexual<br>Bonding<br>Negative<br>Neutral | -0.271, 0.156<br>-0.115, 0.552<br>-0.186, 0.334<br>-0.119, 0.538 |
| Anterior Cingulate   | Sexual<br>Bonding<br>Negative<br>Neutral | -0.155, 0.422<br>-0.067, 0.730<br>-0.166, 0.390<br>0.008, 0.966  |
| Posterior Cingulate  | Sexual<br>Bonding<br>Negative<br>Neutral | -0.230, 0.231<br>-0.170, 0.378<br>-0.001, 0.996<br>-0.166, 0.390 |
| Medial Frontal Gyrus | Sexual<br>Bonding<br>Negative<br>Neutral | -0.233, 0.224<br>-0.284, 0.136<br>-0.346, 0.066<br>-0.196, 0.307 |
| Nucleus Accumbens    | Sexual<br>Bonding<br>Negative<br>Neutral | -0.104, 0.592<br>-0.165, 0.393<br>-0.056, 0.773<br>0.093, 0.631  |
| Caudate              | Sexual<br>Bonding<br>Negative<br>Neutral | -0.295, 0.121<br>-0.334, 0.077<br>-0.318, 0.093<br>-0.255, 0.182 |

Supplemental Data

**Supplemental Table 5.** Correlation analyses between kisspeptin's modulation of the Salience Network (SN) connectivity with psychometric outcomes. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                        | Correlation Coefficient ( $r$ ), $P$ Value |
|------------------------|--|
| BAS Drive              | -0.264, 0.173                              |
| BAS Fun                | 0.020, 0.918                               |
| BAS Reward             | 0.040, 0.841                               |
| BIS                    | -0.143, 0.467                              |
| SADI Evaluative        | 0.191, 0.328                               |
| SADI Negative          | -0.565, 0.002**                            |
| SADI Physiological     | 0.073, 0.712                               |
| SADI Motivational      | 0.267, 0.168                               |
| Positive Affect Change | 0.109, 0.582                               |
| Negative Affect Change | 0.119, 0.545                               |

## Supplemental Data

**Supplemental Table 6.** Correlation analyses between kisspeptin's modulation of the Salience Network (SN) connectivity with task brain activity in *a priori* limbic and paralimbic brain structures. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                      |  | Correlation Coefficient (r), P Value                             |
|----------------------|--|--|
| Amygdala             | Sexual<br>Bonding<br>Negative<br>Neutral | 0.087, 0.654<br>-0.063, 0.747<br>0.107, 0.579<br>-0.018, 0.926   |
| Hippocampus          | Sexual<br>Bonding<br>Negative<br>Neutral | 0.148, 0.442<br>0.124, 0.522<br>0.196, 0.307<br>0.187, 0.332     |
| Thalamus             | Sexual<br>Bonding<br>Negative<br>Neutral | 0.140, 0.469<br>-0.137, 0.479<br>0.072, 0.711<br>0.008, 0.967    |
| Putamen              | Sexual<br>Bonding<br>Negative<br>Neutral | 0.221, 0.248<br>-0.023, 0.904<br>0.114, 0.555<br>0.138, 0.474    |
| Globus Pallidus      | Sexual<br>Bonding<br>Negative<br>Neutral | 0.291, 0.124<br>-0.056, 0.774<br>0.030, 0.878<br>0.003, 0.987    |
| Anterior Cingulate   | Sexual<br>Bonding<br>Negative<br>Neutral | 0.164, 0.394<br>-0.145, 0.453<br>0.158, 0.413<br>0.024, 0.901    |
| Posterior Cingulate  | Sexual<br>Bonding<br>Negative<br>Neutral | 0.301, 0.112<br>-0.063, 0.744<br>0.237, 0.215<br>0.158, 0.413    |
| Medial Frontal Gyrus | Sexual<br>Bonding<br>Negative<br>Neutral | 0.024, 0.900<br>-0.207, 0.281<br>-0.059, 0.761<br>-0.178, 0.354  |
| Nucleus Accumbens    | Sexual<br>Bonding<br>Negative<br>Neutral | -0.107, 0.580<br>-0.144, 0.456<br>-0.139, 0.470<br>0.020, 0.918  |
| Caudate              | Sexual<br>Bonding<br>Negative<br>Neutral | -0.079, 0.685<br>-0.268, 0.159<br>-0.175, 0.362<br>-0.106, 0.583 |

Supplemental Data

**Supplemental Table 7.** Correlation analyses between kisspeptin's modulation of hippocampus-globus pallidus connectivity with psychometric outcomes. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                        | Correlation Coefficient (r), P Value |
|------------------------|--------------------------------------|
| BAS Drive              | 0.245, 0.207                         |
| BAS Fun                | -0.027, 0.892                        |
| BAS Reward             | 0.207, 0.289                         |
| BIS                    | 0.307, 0.111                         |
| SADI Evaluative        | -0.226, 0.246                        |
| SADI Negative          | -0.313, 0.103                        |
| SADI Physiological     | -0.286, 0.138                        |
| SADI Motivational      | -0.055, 0.782                        |
| Positive Affect Change | -0.019, 0.922                        |
| Negative Affect Change | 0.090, 0.647                         |

**Supplemental Table 8.** Correlation analyses between kisspeptin's modulation of hippocampus-globus pallidus connectivity with task brain activity in *a priori* limbic and paralimbic brain structures. Two-tailed partial correlations with associations adjusted for visit order as previously described (1). \*\*  $P < 0.01$ .  $n=29$ .

|                      |          | Correlation Coefficient (r), P Value |
|----------------------|----------|--------------------------------------|
| Amygdala             | Sexual   | 0.134, 0.487                         |
|                      | Bonding  | 0.164, 0.396                         |
|                      | Negative | 0.353, 0.059                         |
|                      | Neutral  | 0.034, 0.862                         |
| Hippocampus          | Sexual   | 0.131, 0.497                         |
|                      | Bonding  | 0.175, 0.362                         |
|                      | Negative | 0.423, 0.021                         |
|                      | Neutral  | 0.158, 0.411                         |
| Thalamus             | Sexual   | 0.159, 0.409                         |
|                      | Bonding  | 0.144, 0.456                         |
|                      | Negative | 0.520, 0.004**                       |
|                      | Neutral  | 0.290, 0.125                         |
| Putamen              | Sexual   | 0.293, 0.122                         |
|                      | Bonding  | 0.269, 0.158                         |
|                      | Negative | 0.485, 0.007**                       |
|                      | Neutral  | 0.360, 0.054                         |
| Globus Pallidus      | Sexual   | 0.209, 0.276                         |
|                      | Bonding  | 0.106, 0.584                         |
|                      | Negative | 0.462, 0.011                         |
|                      | Neutral  | 0.227, 0.236                         |
| Anterior Cingulate   | Sexual   | 0.045, 0.817                         |
|                      | Bonding  | -0.134, 0.487                        |
|                      | Negative | 0.291, 0.125                         |
|                      | Neutral  | 0.058, 0.765                         |
| Posterior Cingulate  | Sexual   | 0.023, 0.906                         |
|                      | Bonding  | -0.060, 0.759                        |
|                      | Negative | 0.484, 0.007**                       |
|                      | Neutral  | 0.086, 0.658                         |
| Medial Frontal Gyrus | Sexual   | 0.087, 0.653                         |
|                      | Bonding  | 0.054, 0.783                         |
|                      | Negative | 0.486, 0.007**                       |
|                      | Neutral  | 0.207, 0.279                         |
| Nucleus Accumbens    | Sexual   | -0.026, 0.892                        |
|                      | Bonding  | -0.096, 0.622                        |
|                      | Negative | 0.475, 0.009**                       |
|                      | Neutral  | 0.325, 0.084                         |
| Caudate              | Sexual   | 0.243, 0.203                         |
|                      | Bonding  | 0.109, 0.575                         |
|                      | Negative | 0.508, 0.005**                       |
|                      | Neutral  | 0.266, 0.162                         |

*Reference:*

1. Comninou AN, Wall MB, Demetriou L, Shah AJ, Clarke SA, Narayanaswamy S, Nesbitt A, Izzi-Engbeaya C, Prague JK, Abbara A, et al. Kisspeptin modulates sexual and emotional brain processing in humans. *The Journal of Clinical Investigation*. 2017;127(2):709-19.