Japan-U.S. Brain Research Cooperation Program Group Joint Study Project Program FY20<u>10</u>- FY20<u>12</u>: Report

Field: \_\_\_\_\_2\_\_\_\_

- Principal Researcher Name Tomomi Shimogori Title Team Leader Affiliation Brain Science Institute RIKEN
- 2. Project Title: Genomic analysis of mouse hypothalamus and nuclei specific gene expression.
- 3. Japanese Group Tomomi Shimogori Team Leader Brain Science Institute RIKEN

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- 4. U.S. Group Seth Blackshaw Associate Professor Johns Hopkins University School of Medicine
- 5. Research Period, from/to (mm/dd/yyyy) and total number of years. Apr. 1. 2010 To Mar. 31. 2013 (3 Years)
- 6. Abstract, Results, and Research Significance (300 words):

The mammalian hypothalamus controls a large range of physiological processes, but the mechanism by which it is patterned during development is poorly understood. We have used microarray-based expression profiling and large-scale two-color in situ hybridization to conduct a detailed characterization of gene expression during mouse hypothalamic neurogenesis. We have determined that a combination of transcription factors define unique domains along the anterior-posterior axis of the developing prethalamus and hypothalamus, implying that these diencephalic regions form a single developmental compartment patterned by common differentiation factors. Furthermore, we have determined that developing hypothalamic nuclei selectively express different Lhx family transcription factors, which are known to direct cell fate specification. Using both targeting *in* utero electroporation and knockout mice, together with the large collection of nuclear and subtype-specific molecular markers identified in our screen, we will investigate cell whether candidate morphogens such as Shh and Wnt3a control anterior-posterior patterning of and hypothalamus, and investigate whether the transcription factors also the prethalamus directly regulate this process. We will likewise use similar techniques to determine whether more transcription factors direct development of the dorsomedial and posterior hypothalamic nuclei.

7. Other (Research-related concerns, particular points of note):

\*Please attach any reference materials as necessary.