## PROGRAM PROJECT GOAL PHASE 1: DEFINE TRIPARTITE SYNAPSE AT THE GENOMIC LEVEL IN MOUSE

| Level of analysis     | Team 1a             | Team 1b                 | Team 2              | Team 3a         | Team 3b           |
|-----------------------|---------------------|-------------------------|---------------------|-----------------|-------------------|
| 1. define molecular   | Glutamate cycling   | Spatal buffering of K+  | NADH transport      | Glutamine       | Glutamate         |
| exchange              |                     |                         | _                   | transporter     | transporter       |
| 2. ID key gene        | SLC1A3              | KCNJ10                  | ID transporter      | SN1             | VGlut (which one) |
| products              |                     |                         |                     |                 |                   |
| 3. Assess specificity | Astrocyte specific? | ?                       | ?                   | ?               | ?                 |
|                       | High affinity?      |                         |                     |                 |                   |
| 4. genomic analysis   | Accession #?        | Accession #?            | Accession #?        | Accession #?    | Accession #?      |
| 5. localization       | ICC                 | ICC                     | Tag NADH,           | In situ, immuno | In situ, immuno   |
|                       |                     |                         | imaging to localize |                 |                   |
| 6. function           | Pharmacol block,    | Pharmacol block,        | Transgenic with     | RNA knockdown   |                   |
|                       | predict             | predict                 | conditional KO of   | in culture      |                   |
|                       | hyperexcitability   | hyperexcitability:ephys | transporter—        |                 |                   |
|                       |                     | measure of Vm           | specificity will be |                 |                   |
|                       |                     |                         | critical: in vitro  |                 |                   |
|                       |                     |                         | model unclear       |                 |                   |
| 7. caveats            |                     |                         | General             |                 |                   |
|                       |                     |                         | metabolism          |                 |                   |
| 8.pitfalls            |                     |                         |                     |                 |                   |
| 9. design             |                     |                         |                     |                 |                   |
| comparative           |                     |                         |                     |                 |                   |
| analysis              |                     |                         |                     |                 |                   |
| 10. cost              | 31K                 | 31K                     | \$31K               | \$32K           | \$32K             |

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| Level of analysis     | Team 4                         | Team 4b                               | Team 5a                   |
|-----------------------|--------------------------------|---------------------------------------|---------------------------|
| 1. define molecular   | Pyruvate transporter           | Pyruvate transporter                  | Monocarboxylate           |
| exchange              | (astrocytic export)            | Neuronal import                       | transporters              |
|                       |                                |                                       | Lactate                   |
| 2. ID key gene        | ?                              | ?                                     | ID transporter            |
| products              |                                |                                       |                           |
| 3. Assess specificity | ?                              | ?                                     | ?                         |
|                       |                                |                                       |                           |
| 4. genomic analysis   | Accession #?                   | Accession #?                          | Accession #?              |
| 5. localization       | ICC                            | ICC                                   | ICC (LM,EM)               |
| 6. function           | Pharmacol                      | Pharmacol                             | In vivo transgenic        |
|                       | inhibition, ephys              | inhibition, ephys                     | with tissue-specific      |
|                       | measures                       | measures                              | promoter (mean            |
|                       | Pulse chase                    | Pulse chase                           | cell-type specific?)      |
|                       | metabolic labeling of pyruvate | metabolic labeling of pyruvate        | Behavioral analysis of KO |
|                       | 15                             | I I I I I I I I I I I I I I I I I I I | Tissue culture            |
|                       |                                |                                       | (apoptosis, slice         |
|                       |                                |                                       | ephys + rescue)           |
| 7. caveats            |                                |                                       | ,                         |
| 8.pitfalls            |                                |                                       |                           |
| 9. design             |                                |                                       |                           |
| comparative analysis  |                                |                                       |                           |
| 10. cost              | 31K                            | 31K                                   | \$\$75K (per gene?)       |