

## Supporting Material

**Figure S1:** SDS-PAGE showing results of limited tryptic digest of  $\alpha$ CaN in the absence and presence of CaM. The  $\alpha$ CaN is protected from digestion by addition of CaM.

**Figure S2:** Deuterium uptake curves for RD-AID-CT peptides determined by HXMS. Different structured regions display distinct exchange kinetics. The horizontal dashed lines, where present, shows totally deuterated control exchange level, ^ denotes peptides used in the heat map, \* indicates uptake curves constructed from incomplete data sets,  $\delta$  shows ambiguous (isobaric) peptide assignment (see Table S1). Error bars were determined from two independent data sets.

**Table S1:** Ambiguous (isobaric) RD-AID-CT peptide assignments as indicated in Figure S6.

| <b>Residue Numbers</b> | <b>Sequence</b>  |
|------------------------|--|
| 377-412                | LGSEEDGFDGATAAARKEVIRNKIRAIGKMARVFSV   |
| 378-413                | GSEEDGFDGATAAARKEVIRNKIRAIGKMARVFSVL   |
| 375-413                | DELGSEEDGFDGATAAARKEVIRNKIRAIGKMARVFSVL  |
| 379-416                | SEEDGFDGATAAARKEVIRNKIRAIGKMARVFSVLREE   |
| 380-417                | EEDGFDGATAAARKEVIRNKIRAIGKMARVFSVLREES   |
| 381-418                | EDGFDGATAAARKEVIRNKIRAIGKMARVFSVLREESE   |
| 418-449                | ESVLTCLKGLTPTGMLPSGVLSGGKQTLQSATV  |
| 419-450                | SVLTCLKGLTPTGMLPSGVLSGGKQTLQSATVE  |
| 421-431                | LTLKGLTPTGM  |
| 422-432                | TLKGLTPTGML  |
| 421-495                | LTLKGLTPTGMLPSGVLSGGKQTLQSATVEAIEADEAIKGFSPQHKITSF<br>EEAKGLDRINERMPPRRDAMPDAN |
| 422-496                | TLKGLTPTGMLPSGVLSGGKQTLQSATVEAIEADEAIKGFSPQHKITSFE<br>EAKGLDRINERMPPRRDAMPDANL |
| 421-443                | LTLKGLTPTGMLPSGVLSGGKQT  |
| 422-444                | TLKGLTPTGMLPSGVLSGGKQTL  |
| 422-447                | TLKGLTPTGMLPSGVLSGGKQTLQSA   |
| 423-448                | LKGLTPTGMLPSGVLSGGKQTLQSAT   |

| <b>Residue Numbers</b> | <b>Sequence</b>   |
|------------------------|-------------------|
| 450-455                | EAIEAD            |
| 451-456                | AIEADE            |
| 452-457                | IEADEA            |
| 453-458                | EADEAI            |
| 454-470                | ADEAIKGFSPQHKITSF |
| 460-476                | GFSPQHKITSFEEAKGL |
| 456-470                | EAIKGFSPQHKITSF   |
| 457-471                | AIKGFSPQHKITSFE   |
| 456-471                | EAIKGFSPQHKITSFE  |
| 457-472                | AIKGFSPQHKITSFEE  |
| 458-473                | IKGFSPQHKITSFEEA  |
| 461-476                | FSPQHKITSFEEAKGL  |