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FIG. S1. Schematic diagrams of TbrPDEB1 and TbrPDEB2 showing functional domains (grey boxes) and allelic polymorphisms in T. b. brucei strain Lister 427 and the Cpd A resistant R0.8 cell line. The ORFs are represented by the largest box (white and grey parts); flanking regions are represented by thick black horizontal lines. Allelic polymorphisms are represented by thin (single nucleotide) or thicker (clusters of multiple nucleotides) vertical lines.

FIG. S2. Western Blot analysis of CARP1 repression (1 µg/ml tetracycline for 24h, + Tet) in additional CARP1 RNAi cell lines with in situ tagged CARP1 either with an N-terminal 4xTy1 (upper panel) or a C-terminal 3xHA tag (lower panel). CARP1 protein levels were normalized to PFR-A/C detected by the monoclonal antibody L13D6 (36) and set to 100% in the absence of Tet. The relative scan gain in the 800 nm channel was set to 1 for the upper panel and to 3 for the lower panel. Relative expression levels are indicated as percentage of the non-induced cultures.

FIG. S3. CARP1-4 mRNA expression levels in T. b. brucei strain Lister 427 wild type (WT) and derived R0.8 cell line in the presence or absence of Cpd A, determined by quantitative real-time PCR. Relative expression normalized to TERT as reference gene (37) and the wild type line is given with standard error (SEM) of three biological replicates.
Allelic Polymorphisms of *TbrPDEB1*

- Allelic Polymorphism 1 (bp 738)
- Allelic Polymorphism 2 (bp 1362)
- Allelic Polymorphism 3 (bp 1602)

Allelic Polymorphisms Cluster 1 (bp 780-792)
Allelic Polymorphisms Cluster 2 (bp 882-855)
Allelic Polymorphisms Cluster 3 (bp 1002-1008)
Allelic Polymorphisms Cluster 4 (bp 1070-1077)

Allelic Polymorphisms of *TbrPDEB2*
CARP1 RNAi / p3077 CARP1 (in situ N-terminal 4xty1)

pool 1                  clone 1                   clone 2
- Tet       + Tet      - Tet      + Tet         - Tet        + Tet
100%       55%       100%      38%        100%        58%

4xty1-CARP1
PFR-A/C

CARP1 RNAi / pMOTag2H CARP1 (in situ C-terminal 3xHA)

pool 1                  pool 2                           clone 1
- Tet       + Tet       - Tet           + Tet           - Tet          + Tet
100%        55%          100%           56%            100%         60%

CARP1-3xHA
PFR-A/C
CARP1-4 mRNA expression levels

Fold Change

WT
WT + Cpd A
R0.8
R0.8 + Cpd A

CARP1
CARP2
CARP3
CARP4