

## Supplementary Material

## Supplementary Figures



**Supplementary Figure S1: SmHGPRTAses expression profiles in different cell clusters identified in** *Schistosoma mansoni* adult female, immature female, and male worms. UMAP projection graphs with scRNAseq expression profile in females, immature females (IM) and males of *S. mansoni* from SmHGPRTAses 1 (A), 2 (B), 3 (C), and 4 (D). Expression values were normalized and represented by colors (gray = not expressed, red = expressed).



## Transcript levels in sporocyst knocked-down for SmHGPRTases

**Supplementary Figure S2:** *SmHGPRTases* **transcript levels after sporocysts exposure to dsRNAs.** Bars representing the transcript levels of *SmHGPRTase 1, SmHGPRTase 2/4/5,* and *SmHGPRTase 3* in sporocysts after exposure to the specific dsRNAs separately or in combination (combined) relative to the untreated (grey) or unspecific control (black) after 2 and 7 days. Above the 2-day bars are represented the standard error of the mean of three replicates. The dashed line represents the normalized values in the controls. After verifying the normality using the Shapiro-Wilk test, significant differences compared to control conditions were analyzed by unpaired t-test (\*  $p \le 0.05$ ).



Transcript levels in schistosomula knocked-down for SmHGPRTases

**Supplementary Figure S3: SmHGPRTases transcript levels and area of schistosomula after exposure to dsRNAs.** Bars representing the transcript levels of *SmHGPRTase 1, SmHGPRTase 2/4/5,* and *SmHGPRTase 3* in schistosomula after exposure to the specific dsRNAs separately or in combination (combined) relative to the untreated (grey) or unspecific control (black) after 2 days.



**Supplementary Figure S4: Knockdown effects in adenosine levels after parasites exposure to dsRNAs.** Bars representing the adenosine levels in schistosomula (A) and in male adult worms (B) in the respective groups: unspecific control, parasites exposed to *SmHGPRTase 1*-dsRNA, *SmHGPRTase 2/4/5*-dsRNA, *SmHGPRTase 3*-dsRNA, and the combined group. Above the bars are represented the standard error of the mean of three replicates.



**Supplementary Figure S5: Representative images of schistosomula exposed to SmHGPRTasesdsRNA.** Untreated control (A) and schistosomula exposed to *SmHGPRTases*-dsRNA (B). Black arrows indicate changes in phenotype or dead schistosomula.

## Male adult worm motility after electroporation with SmHGPRTases-dsRNA



**Supplementary Figure S6: Adult male worm's motility after electroporation with SmHGPRTases-dsRNA.** Dots represent the average of the movement units of male adult worms that were electroporated with *SmHGPRTases*-dsRNAs. Blue: parasites exposed to *SmHGPRTase 1*-dsRNA. Green: parasites exposed to *SmHGPRTase 2/4/5*-dsRNA. Red: parasites exposed to SmHGPRTase 3-dsRNA. Black: parasites exposed to the combination of the three *SmHGPRTases*-dsRNAs. The results were normalized according to the movement units of adult worms from control groups (dotted line).