## **1 SUPPLEMENTARY TABLE:**

NAME	ID*	Effect	% INH/ENH	IC <sub>50</sub>
				(µM)
C1	F1115-0016	INH	0.17 ±0.1	1.36 ±3.54
C2	F1041-0001	INH	0.19 ±0.1	$1.34\pm0.27$
C3	F3215-0002	INH	0.65 ±0.1	0.57 ±0.06
C4	F1744-0034	INH	1.57 ±0.5	3.46 ±0.08
C5	F0909-0123	INH	1.82 ±2.2	$1.12\pm0.18$
C6	F1092-0914	INH	10.9 ±15.0	$6.48\pm0.83$
C7	F5040-0271	INH	11.7 ±4.4	$8.70 \pm 1.03$
C8	F0406-1005	ENH	205.3 ±7.6	$0.14 \pm 0.02$
C9	F1874-0558	ENH	207.4 ±7.6	1.35 ±0.41
C10	F0779-0483	ENH	210.2 ±4.1	$1.24\pm0.68$
C11	F0745-0014	ENH	213.8 ±18.8	$0.21\pm0.05$
C12	F0862-0340	ENH	214.1 ±31.0	1.19 ±0.22
C13	F3220-0514	ENH	220.3 ±1.3	$0.38\pm0.26$
C14	F5570-0789	ENH	269.5 ±10.0	5.54±4.28

2 Suplementary Table 1: Summary of effects of 14 validated compounds identified in this study

3 \*Life Chemicals ID. Can be searched at http://www.emolecules.com

## 4 SUPPLEMENTARY FIGURES AND LEGENDS:



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7 viability. All validated compounds recorded absorbance levels similar to vehicle while

8 staurosporine (Stau\*) recorded low absorbance signal, an indication of toxic effects on the host

9 cells.

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Figure S2. Bioluminiscence signal data showing the effects of C1,C2, C3 and C5 on parasite growth in mice. C2-treated mice recorded low signal during days 1-6 compared to controls, which was significantly different on D4. B) Total flux (photons/s) data showing how each compound affected parasite growth in mice. C2 was effective in reducing parasite growth during days 1-6.



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Figure S3. Dorsal signal during days 15-25 after infection with strain 5A10 and treatment with either vehicle or C2. Despite the dramatically reduced parasite burden in C2-treated mice during the acute phase of infection (Fig. 9A-B), all mice eventually exhibited significant signal in the brain that was similar between vehicle and C2-treated mice.



Figure S4. Tanimoto based hierarchical clustering and heat map of the 14 identified compounds in our study and the 31 parasite invasion modulators identified in Carey et al (7). The compound dataset was clustered using the R package ChemmineR (1) using atom pair fingerprints and hclust (ward method). The cluster plot, heatmap, and heat map key were generated using the R package gplots (2). As shown in the heat map key histogram, the highest similarity value was 0.55 and 95% of the compounds had similarities less than 0.3.

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## 32 Supplementary References cited:

- Cao, Y., A. Charisi, L. C. Cheng, T. Jiang, and T. Girke. 2008. ChemmineR: a compound mining
  framework for R. Bioinformatics 24:1733-1734.
- Gregory R. Warnes et al. 2012. gplots: Various R programming tools for plotting data. R package
  2.11.0. http://CRAN.R-project.org/package=gplots

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