- 1 Supplemental Tables and Figures.
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- 3 Supplemental Table ST1. Chemical composition of Anidulafungin loaded
- 4 liposomes (AFG-LLs).

	µg in each			Moles percent relative
	preparation of	μg / μmole	µmoles	to moles of liposomal
Compound	liposomes	Mol. Wgt.	total	lipid
DSPC (1,2-distearoyl-sn-				
glycero-3-phosphocholine)	6516.00	790.00	8.25	50.00
cholesterol	1591.00	387.00	4.11	45.00
mPEG2000-DSPE	2302.10	872.00	<u>2.64</u>	5.00
Total mmoles and moles				
percent liposomal lipid			15.00	100.00
Anidulafungin (AFG)	1030.00	1140.00	0.90	6.02
Lissamine Rhodamine B 1,2-				
Dihexadecanoyl-sn-Glycero-				
3-Phosphoethanolamine,				
Triethylammonium Salt	400.00	1333.81	0.30	2.00

7 Supplemental Fig. SF1. Treatment regimens used to assay liposome binding, fungal burden and mouse survival- immunosuppression, infection, liposome 8 9 treatments and endpoints. A neutropenic mouse model was used to ensure 10 reproducible infection by C. albicans yeast cells. A. The regimen used to assay binding of DectiSomes to infection centers in the kidneys as compared to AmB-LLs and the 11 12 impact of DectiSomes on fungal burden as compared to untargeted liposomes or control buffer. Liposome binding was assayed on Day 4 (D4) PI and fungal burden was 13 determined on D1 PI. B. Regimen used to assay mouse survival after treatment with 14 15 DEC1-AmB-LLs, DEC2-AmB-LLs, AmB-LLs or control buffer. The Day(s) (D) of A. Regimen for assaying liposome binding in the kidneys or fun 16 treatment are burden in the kidneys Intravenous Intravenous Fungal burden Immunoindicated before and 17 or liposome suppression C. albicans Liposomes binding in cells 18 after the day of CP + TCkidneys 19 infection (D0). CP, D-3 D0 or D0+D1 D1/D4 D0 ŧ ¥ ¥ ŧ 20 cyclophosphamide. Timeline in hrs & Days (D) *م* 21 TC, triamcinolone. 22 Survival was 23 monitored until D10. B. Regimen for assaying the survival of mice with candidiasis Intravenous Intravenous Immuno-Mouse 24 C. albicans Liposomes suppression survival CP + TC D-3 D3 D0, D1, D2 D0 D3 to D10 ł ŧ Ł + + + + Timeline in Days (D)

- 25 Supplemental Fig. SF2. DectiSomes targeted by Dectin-1 or Dectin-2 and
- 26 delivered intravenously are concentrated in *C. albicans* infection centers in the
- 27 mouse kidney. Replicate images for the experiment illustrated in Fig. 2 showing the
- superior binding of DEC1-AmB-LLs and DEC2-AmB-LLs to *C. albicans* colonies in the
- 29 kidneys relative to untargeted AmB-LLs.



Supplemental Fig. SF3. Dectin-2 coated anidulafungin loaded liposomes, DEC2AFG-LLs, delivering 0.6 mg/kg AFG were more effective at reducing the burden of *C. albicans* in the kidneys than untargeted AFG-LLs. Relative Quantity (RQ) of *C. albicans* rDNA was determined by qPCR analysis of DNA isolated from duplicate kidney
samples to those in Fig. 3E, serving as a confirmation of the CFU data.



37 Supplemental Fig. SF4. Experimental data used to guantify the moles percent of AFG loaded into AFG-LLs. Light scattering from liposomes prevents a direct 38 measurement of the UV absorbance of drugs loaded into liposomes. Hence, we 39 40 estimated the percent of AFG loaded into liposomes by the subtraction of that which 41 was not loaded into liposomes as we have done before to estimate the loading of AmB 42 (Ambati et al., 2019, mSphere 4:1-15). A. Standard curve plotting the absorbance of AFG at A340 in a dilution series in a 96 well microtiter plate vs the amount mg amount of 43 AFG dissolved in 50 uL of DMSO. When more than 0.12 mg of AFG were examined the 44 45 A340 absorbance values were too high to be read. B. When 1.7 mg of AFG was tumbled for 3 days in 273 uL of liposome buffer, was 81.3% of the total remained 46 47 insoluble. This was determined by taking the insoluble AFG precipitate and dissolving it in 50 uL DMSO and reading A340 for a dilution series. C. After incubating liposomes 48 and 1.7 mg of AFG together in 273 uL the insoluble AFG was spun down and assayed 49 by in a dilution series in 50 uL DMSO. This is the amount of AFG not taken up by 50 51 liposomes and not soluble in the buffer surrounding them. **D.** The amount of AFG that remained soluble in 273 uL of liposome buffer was estimated by subtracting the data in 52



56 Supplemental Fig. SF5. Assays of *C. albicans* microcolonies to determine fungal

57 burden in the kidneys of neutropenic mice with candidiasis than AmB-LLs (an

58 example experiment). Aliquots of homogenized kidney tissue were diluted into PBS,

59 plated on YPD agar, incubated 11 hr at 37°C, and microcolonies counted. **A.** A bar plot

60 compare the average number of CFUs of *C. albicans* for mice treated once with DEC2-

61 AmB-LLs or AmB-LLs delivering 0.2 mg/kg AmB or with liposome dilution buffer.

62 Standard errors are indicated by a line and whisker. **B**, **C**, **& D**. Examples of the images

63 used to make CFU estimates. Microcolonies ranging from 5 to 300 microns in diameter

64 were counted from the bottom of agar petri plates on an EVOS imaging system at 4X

65 magnification. The number of CFUs was corrected for the area of the entire plate

relative to each microscopic field, the amount of homogenized kidney tissue plated, and
the weight of each kidney pair. Six mice were in each treatment group in this example.



