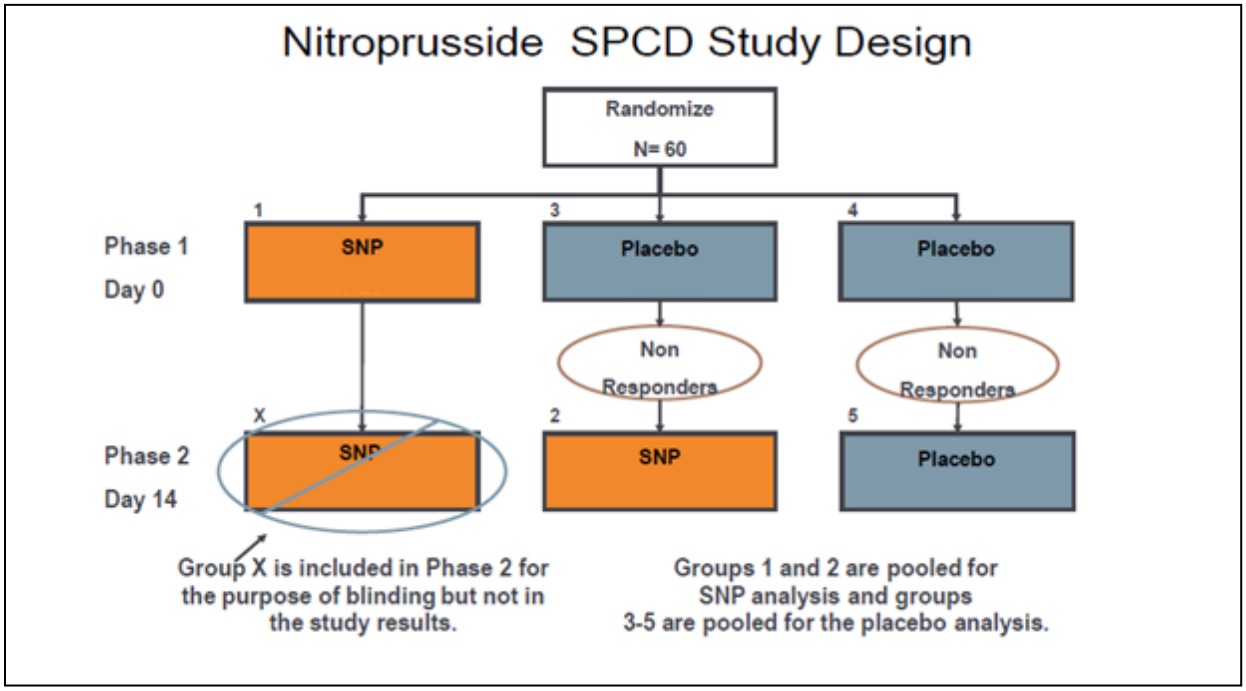
Supplementary Online Content

Brown HE, Freudenreich O, Fan X, et al. Efficacy and tolerability of adjunctive intravenous sodium nitroprusside treatment for outpatients with schizophrenia: a randomized clinical trial. *JAMA Psychiatry.* Published online March 27, 2019. doi:10.1001/jamapsychiatry.2019.0151

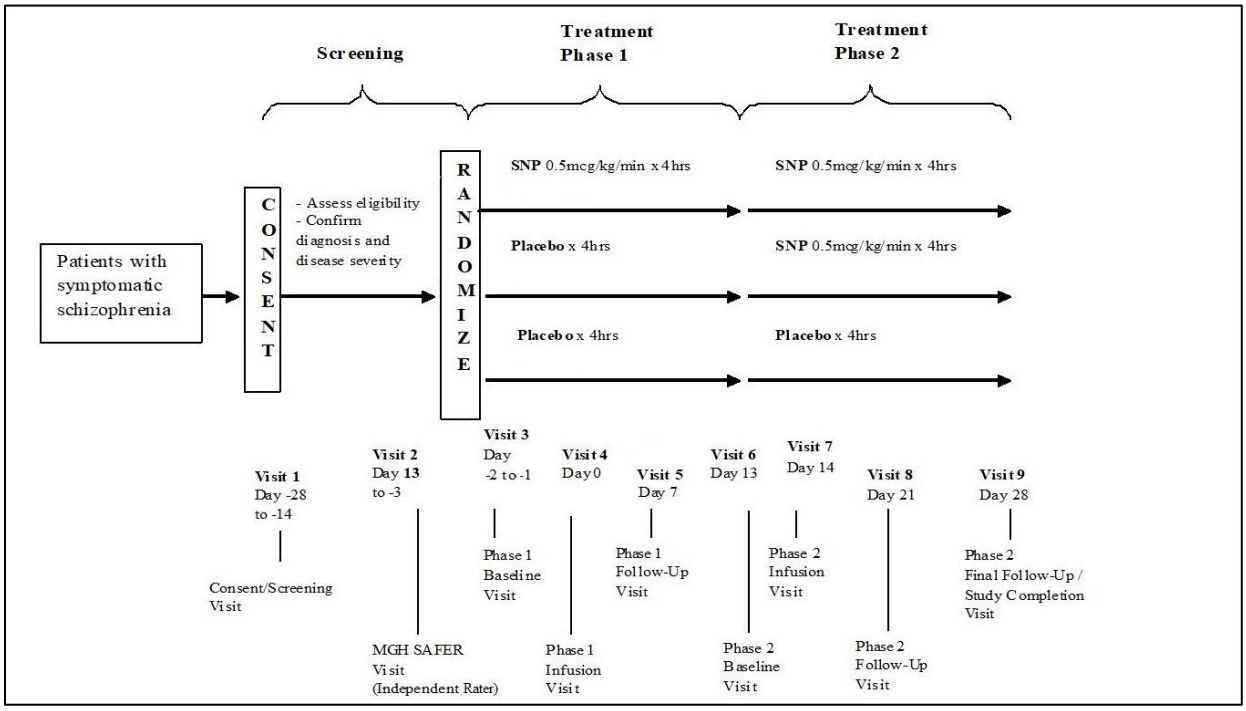
This supplementary material has been provided by the authors to give readers additional information about their work.

eFigure 1. Sequential Parallel Comparison Design (SPCD)



SPCD=Sequential Parallel Comparison Design; SNP=sodium nitroprusside

eFigure 2. Study Schema



SNP=sodium nitroprusside; MGH=Massachusetts General Hospital

eTable 1. Positive and Negative Syndrome Scale (PANSS) Scores Stratified by Treatment Status

		Clozapine		No Clozapine	
		n=19		*n=33*	
		p-value	**estimate**	**p-value**	**estimate**
Primary Outcome Measure					
	PANSS total (SPCD analysis)	0.41	-1.85	0.57	-1.50
Secondary Outcome Measure					
	PANSS - positive (SPCD analysis)	0.41	-0.83	0.55	-0.60
	PANSS - negative (SPCD analysis)	0.79	0.29	0.79	-0.22
	PANSS - general (SPCD analysis)	0.55	-1.08	0.97	0.06

SPCD=Sequential Parallel Comparison Design; PANSS=Positive and Negative Syndrome Scale

eTable 2. Positive and Negative Syndrome Scale (PANSS) Scores Stratified by PANSS-Negative Subscores

		High PANSS-N *n=27*			**Low PANSS-N** *n=25*	
		p-value	estimate		p-value	estimate
Primary Outcome Measure						
	PANSS total (SPCD analysis)	0.23	-2.95		0.66	1.17
Secondary Outcome Measure						
	PANSS - positive (SPCD analysis)	0.24	-1.06		0.86	0.15
	PANSS - negative (SPCD analysis)	0.71	-0.35		0.94	-0.08
	PANSS - general (SPCD analysis)	0.25	-1.59		0.41	1.30

SPCD=Sequential Parallel Comparison Design; PANSS=Positive and Negative Syndrome Scale; PANSS-N= Positive and Negative Syndrome Scale, negative symptoms subscore

eTable 3. Percentage of Participants Reporting Experiencing SAFTEE Symptoms at "Moderate" or "Severe" Level for the 10 Most Frequently Reported Symptoms Post Randomization

	SNP SNP			Placebo SNP			Placebo Placebo			Total
SAFTEE Symptoms	*n=18*			*n=16*			*n=18*			*n=52*
	BL	**FU**		**BL**	**FU**		**BL**	**FU**		**FU**
Hearing or seeing things	22%	33%		25.0%	31%		22%	17%		27%
Feeling drowsy or sleepy	17%	22.2%		0.0%	25%		22%	22%		23%
Trouble concentrating	17%	11.1%		6%	19%		22%	28%		19%
Poor memory	11%	5.6%		6%	25%		28%	22%		17%
Trouble Sleeping	17%	11.1%		6%	13%		28%	22%		15%
Feeling nervous or hyper	6%	5.6%		19%	19%		33%	22%		15%
Dry mouth	6%	5.6%		13%	25%		11%	17%		15%
Frequent need to urinate	0%	5.6%		6%	19%		11%	17%		14%
Stuffy nose	6%	5.6%		6%	19%		17%	11%		12%
Hot flashes	0%	11.1%		6%	6%		6%	17%		12%

BL = reported at baseline; FU = reported at any assessment thereafter (“follow-up”), including 7 hours after infusions 1 and 2, Baseline 2 (Day 14), and final follow-up, including early termination visits; SNP = sodium nitroprusside; SAFTEE = Systematic Assessment for Treatment Emergent Effects