

**Order: 9191 Doxycycline HCl 50uM + GSK2126458 10uM**

**Experiment type: Sponsor a combinatorial intervention**

**Thank you for purchasing a lifespan analysis study on your chosen interventions!**

Below you will find the data for the treatment you sponsored. Please remember that this data is conducted in a model system and is not intended to inform personal health decisions. Please consult your physician before making changes to your personal health regimen.

### **Advice on data interpretation**

At Ora Biomedical Inc. we conduct rigorous science with the goal of producing the highest quality, reproducible data. While a single promising result is exciting (and a negative result may be disheartening), please keep in mind that this data represents a single experiment performed at a single dose. Treatments were compared to matched solvent-only controls conducted at the same time. Insights gleaned from this study only reflect the specific dose and testing regime used. For context, highly promising interventions produce lifespan extensions above 30%, depending on drug dosage and other environmental factors. If you have questions regarding your treatment, feel free to reach out and we will do our best to contextualize the data for you. Please note, we pool data from all control populations tested along with your treatment and their number will often greatly exceed that of treated animals.

Lastly, do not get discouraged by a negative result! As we explore chemical space, understanding which treatments do not effect aging is as important as finding the ones that do! Keep in mind that the number of treatments with no or negative effect on aging will be greater than those that increase lifespan. Often these results are not published in academic research which can result in wasted resources and effort as others try and re-try an experiment.

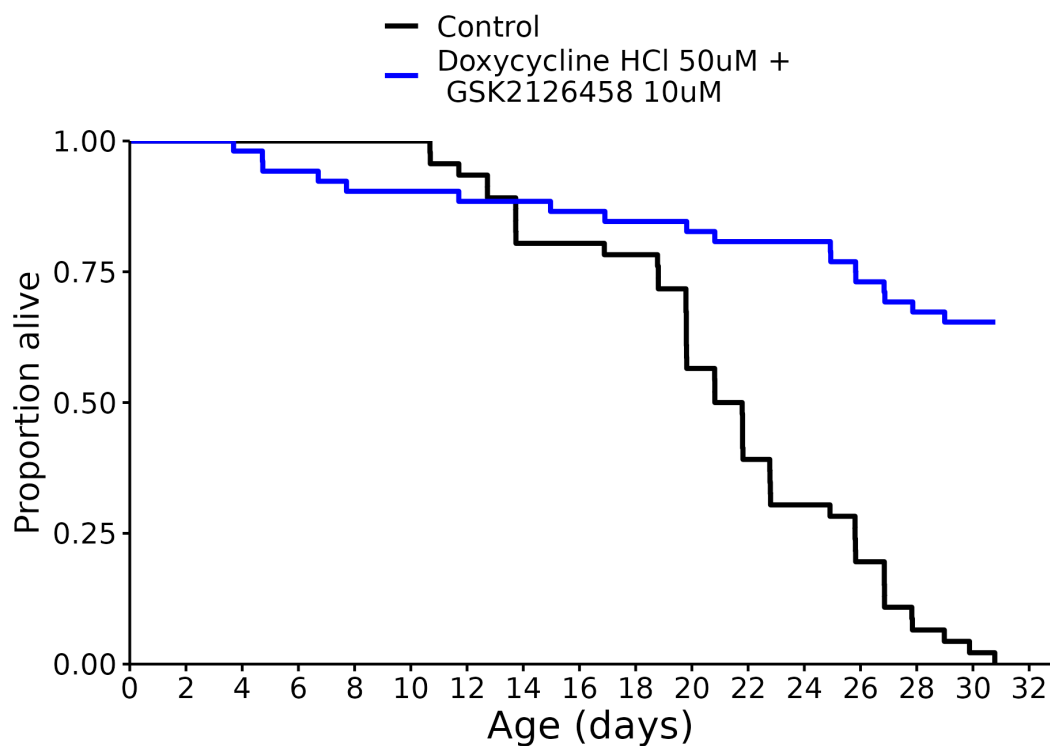
### **Methods**

Experiments were conducted at 20 degrees Celsius using adult N2 Bristol *Caenorhabditis elegans* from the *Caenorhabditis* Genetics Center. Animals were grown from egg to enrollment in the study at 20C from age-synchronized populations on nematode-growth-media (NGM) agar and fed with laboratory grown *E. coli* bacteria. At late L4 larval stage, 20-25 animals were transferred to NGM agar supplemented with either the compound(s) of interest or relevant solvent controls. Floxuridine was added to prevent progeny production. Observations were made every day using the WormBot automated capture platform and images were analyzed via the WormBot-AI analysis pipeline. For your experiment, we tested compound(s) at the concentration specified below in at least two replicate populations. Data was pooled across these replicate populations and compared to pooled control data. To assess whether the tested treatment extended organismal lifespan, we compared treatment to solvent control using a standard logrank statistical test. See page two for drug information, Kaplan-Meier survival curve, and summary data table. Thank you from all of us at Ora Biomedical!

## Drug information from chemical library

Compound Name	Formula	Pathway	Target
Doxycycline HCl	C <sub>22</sub> H <sub>24</sub> N <sub>2</sub> O <sub>8</sub> .HCl	Proteases	MMP
GSK2126458	C <sub>25</sub> H <sub>17</sub> F <sub>2</sub> N <sub>5</sub> O <sub>3</sub> S	PI3K/Akt/mTOR Signaling	mTOR

## Survival curve



## Summary table

Treatment	Median lifespan	# animals	# replicates	95% LCL	95% UCL	Log-rank p-value
Control	21.3	46	5	19.8	22.8	-
Doxycycline HCl 50uM + GSK2126458 10uM	NA	52	5	NA	NA	< .001

Questions? Comments? Want to design an additional experiment using a disease model or examine how the treatments shown here may protect against physiological stress?

Please contact us at [mitchell@orabiomedical.com](mailto:mitchell@orabiomedical.com).

## Terms and Conditions

*Effective Date: Jan 1, 2024*

*Revised: Feb 12, 2024*

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