A null mutation of *C. elegans* vwa-8

Ming Zhu\(^1\), Andrew D Chisholm\(^1\) and Yishi Jin\(^1§\)

\(^1\)Section of Neurobiology, University of California San Diego, La Jolla, CA 92093, United States

\(^§\)To whom correspondence should be addressed: yijin@ucsd.edu

**Figure 1:** (A) shown is vwa-8 gene structure from WormBase (Version: WS276). Blue bar indicates vwa-8(ju1659) deletion. (B) vwa-8(ju1659) deletes 7989bp. The start and stop codons of the F11C1.5a.1 isoform are highlighted in yellow. Exonal sequences are shown in uppercase.

**Description**

VWA8 proteins, named for von Willebrand factor A (VWA) domain containing 8, are conserved from worm to mammals (Whittaker & Hynes, 2002). In human, two SNPs (rs9566845 and rs9566867) in vwa8 are found to be associated with bipolar disorder with comorbid migraine (Oedegaard et al., 2010). Another SNP (rs9532931) is tentatively associated with a specific sub-group of autism patients (Anney et al., 2010). The *C. elegans* VWA-8 long isoform shares 38% and 55% amino acid sequence identity and similarity, respectively, with human VWA8 long isoform. We showed that endogenous *C. elegans* vwa-8, we generated a null allele vwa-8(ju1659) by CRISPR-Cas9. vwa-8(ju1659) mutants are homozygous viable, and indistinguishable from wild type in gross phenotypes such as body size, brood size, growth rate and movement.

**Methods**

Request a detailed protocol

We generated vwa-8(ju1659) using two CRISPR RNA (crRNAs), 5’-AGTGAAACCCGTGTGATCAT-3’ and 5’-CTACAACGAGAGTTGCCTGT-3’ (Integrated DNA Technologies) targeting the start and stop codons of vwa-8, respectively. The crRNAs were injected into wild type hermaphrodites with purified Cas9 protein (MacroLabs, University of California Berkeley), trans-activating crRNA (tracrRNA) and dpy-10 crRNA, as described (Paix, Folkmann, Rasoloson, & Seydoux, 2015). The dumpy F1 progeny of the injected P0 wild type animals were singled to separate plates. F1 dumpy worms were then genotyped for the presence of potential vwa-8 deletions using the following primers: 5’-CCCTCAGGAGGCCCCCATATTTT-3’ and 5’-TGCTCTCGAACAACCTTGCTT-3’. Several independent vwa-8 deletions were identified. All deletion mutants behaved similarly. ju1659 is a 7989bp deletion of vwa-8 which removes nearly all the coding sequence of vwa-8, except the last 82bp of the last exon. CZ26606 vwa-8(ju1659) was generated after outcrossing with N2 for 3 times to remove the dumpy mutation.

**Reagents**

CZ26606 vwa-8(ju1659) will be available at the CGC.

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References

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