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2 lola1 Is an Evolutionarily New Epigenetic Regulator of dpp Transcription during Dorsal-Ventral Axis Formation.

Quijano JC, Wisotzkey RG, Tran NL, Huang Y, Stinchfield MJ, Haerry TE, Shimmi O, Newfeld SJ

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Mol Biol Evol. 2016 Oct; 33(10):2621-32

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Very Good

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NEW FINDING

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The fate of new genes, duplicates of existing genes, is often elimination by natural selection. In some cases, these genes provide a novel function or more efficiently execute an existing function, offering the organism a selective advantage and are maintained. Yet how a new gene with a novel function becomes integrated into an existing genetic network is not well known. The distribution of these events across the phylogenetic landscape is also unknown.

The same group of authors has produced a pair of comparative experimental studies of the genetic network underlying the formation of the dorsal-ventral body axis during development in *Drosophila* and vertebrates. First, they found that a new gene unique to vertebrates, a Ring class ubiquitin ligase called TRIM33, had replaced an older gene called Nedd4, a Hect class ubiquitin ligase, that was common to both phyla {1}. Those experiments revealed that the vertebrate dorsal-ventral pathway, until then considered universally conserved with *Drosophila*, was able to incorporate a new gene.

The current paper nicely complements that analysis by demonstrating that a new gene in the fly lineage, called *Lola1* has been incorporated into the conserved dorsal-ventral patterning network. *Lola1* is present only in insects and crustaceans, yet it functions as part of a multiprotein complex that maintains the expression of Dpp - a key signaling molecule in dorsal-ventral axis formation that is absolutely conserved as vertebrate BMP2/4. The authors predict that while *Lola1* itself is not present in vertebrates, its function in the maintenance of BMP2/4 expression almost certainly is. They suggest several candidates and await the outcome.

Overall, these two studies demonstrate the new genes are capable of incorporation into existing, highly conserved genetic networks across phyla. From a larger perspective, they reveal that the dorsal-ventral pattern formation pathway functioning in all Bilateria has both highly conserved components and species-specific components, an advance on the current model of monolithic conservation.

References

1. New gene evolution in the bonus-TIF1-γ/TRIM33 family impacted the architecture of the vertebrate dorsal-ventral patterning network.

Wisotzkey RG, Quijano JC, Stinchfield MJ, Newfeld SJ. Mol Biol Evol. 2014 Sep; 31(9):2309-21

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Disclosures

None declared

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Abstract:

ABSTRACT

Secreted ligands in the Dpp/BMP family drive dorsal-ventral (D/V) axis formation in all Bilaterian species. However, maternal factors regulating Dpp/BMP transcription in this process are largely unknown. We identified the BTB domain protein longitudinals lacking-like (*lola1*) as a modifier of decapentaplegic (*dpp*) mutations. We show that *Lola1* is evolutionarily related to the Trithorax group of chromatin regulators and that *lola1* interacts genetically with the epigenetic factor Trithorax-like during Dpp D/V signaling. Maternally driven *Lola1*(HA)... more » is found in oocytes and translocates to zygotic nuclei prior to the point at which *dpp* transcription begins. *lola1* maternal and zygotic mutant embryos display significant reductions in *dpp*, *pMad*, and *zerknüllt* expression, but they are never absent. The data suggest that *lola1* is required to maintain *dpp* transcription during D/V patterning. Phylogenetic data revealed that *lola1* is an evolutionarily new gene present only in insects and crustaceans. We conclude that *Lola1* is the first maternal protein identified with a role in *dpp* D/V transcriptional maintenance, that *Lola1* and the epigenetic protein Trithorax-like are essential for Dpp D/V signaling and that the architecture of the Dpp D/V pathway evolved in the arthropod lineage after the separation from vertebrates via the incorporation of new genes such as *lola1*.

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