

Har-P weaponizes P-transposase to severely impact Drosophila gonad development.

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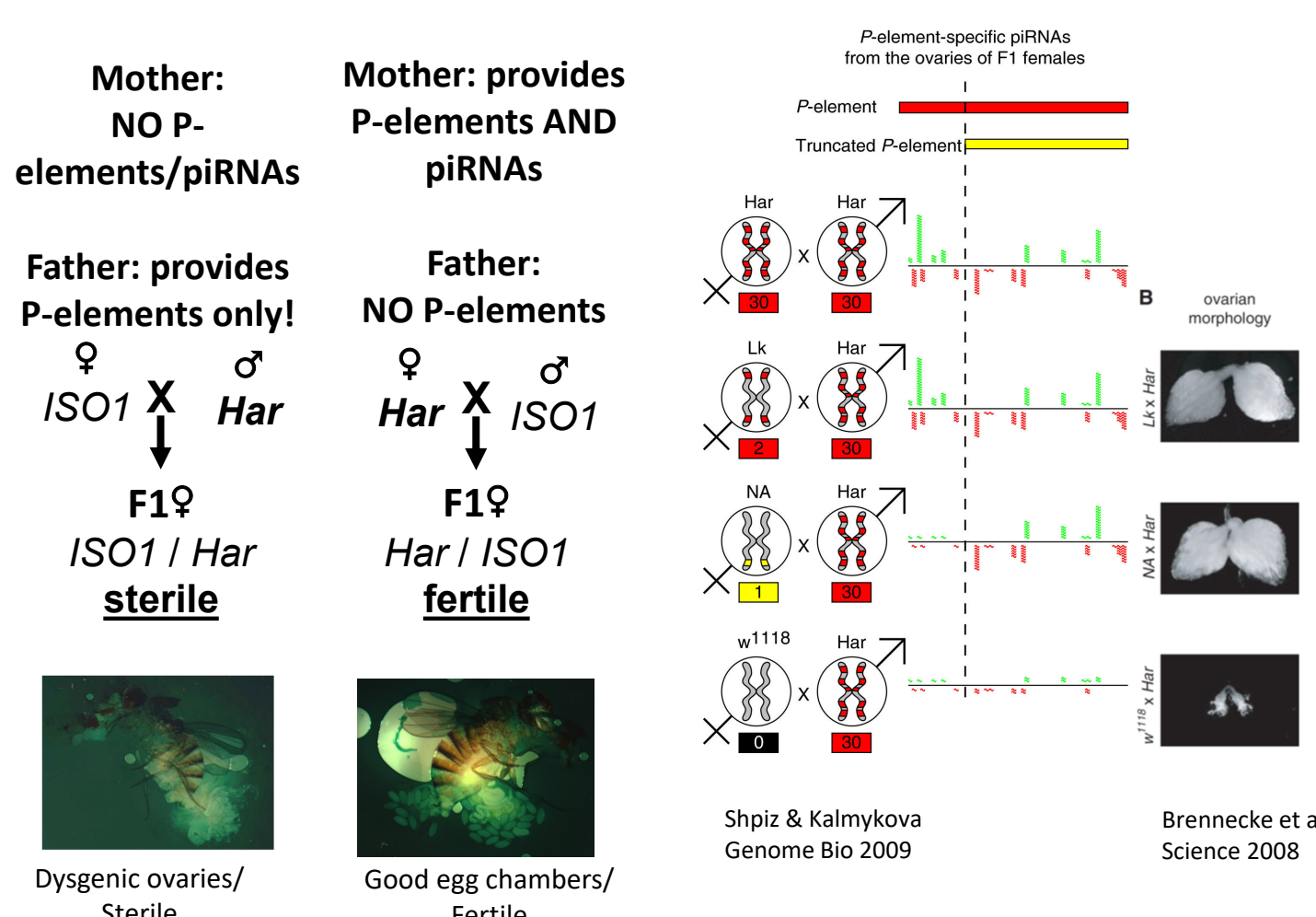
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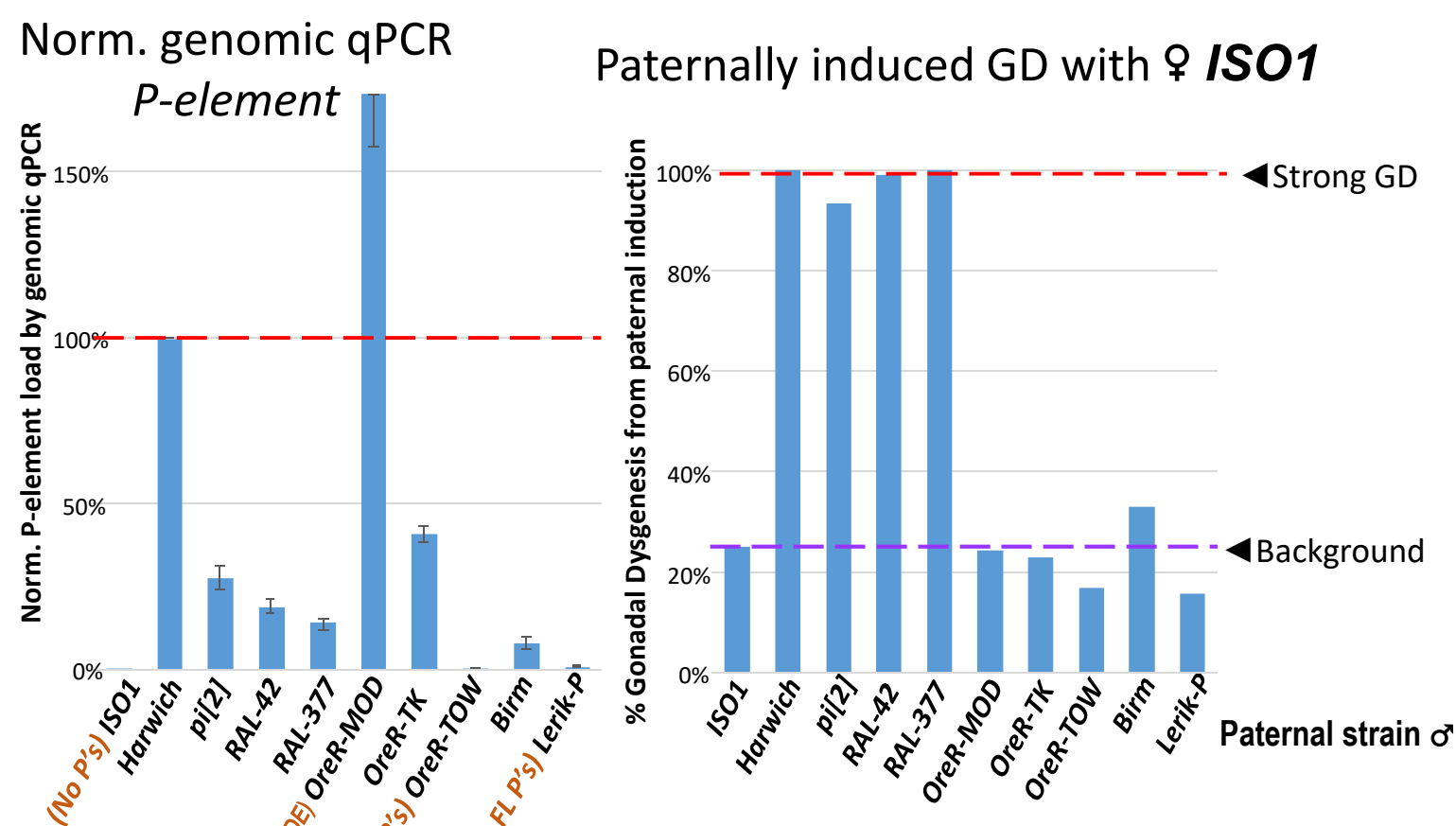
Abstract

When *Drosophila* mothers lacking *P*-element-piRNAs (i.e. *ISO1*) mate with certain fathers harboring active *P*-elements (i.e. *Harwich*, *Har*), hybrid daughters suffer from gonadal dysgenesis (GD), where ovaries collapse from uncontrolled *P*-element transposition. To discover the *P*-element driving GD, we selected hybrid *D. melanogaster* lines with portions of *Har* DNA within the *ISO1* genome. We generated *HISR-D* and *HISR-N* lines retaining or losing GD induction, respectively. Despite greatly-reduced numbers of *P*-elements, *HISR-D* lines produced as many *P*-element piRNAs as parental *Har*. In these lines, we discovered a highly-truncated *P*-element variant: “*Har-P*”, which mobilized *de novo* in all *HISR* lines. Crossing *P*-transposase with *Har-P*s in *HISR-N* lines restored GD and revealed a paternal *P*-element-piRNA-directed imprint on *Har-P*s to suppress lethal somatic transposition. *Har-P* may weaponize *P*-transposase during catastrophic transposition.

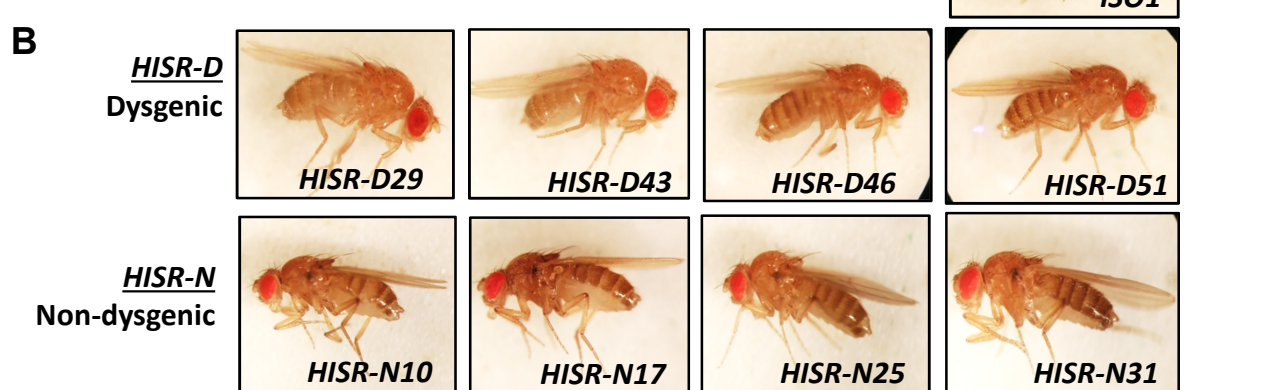
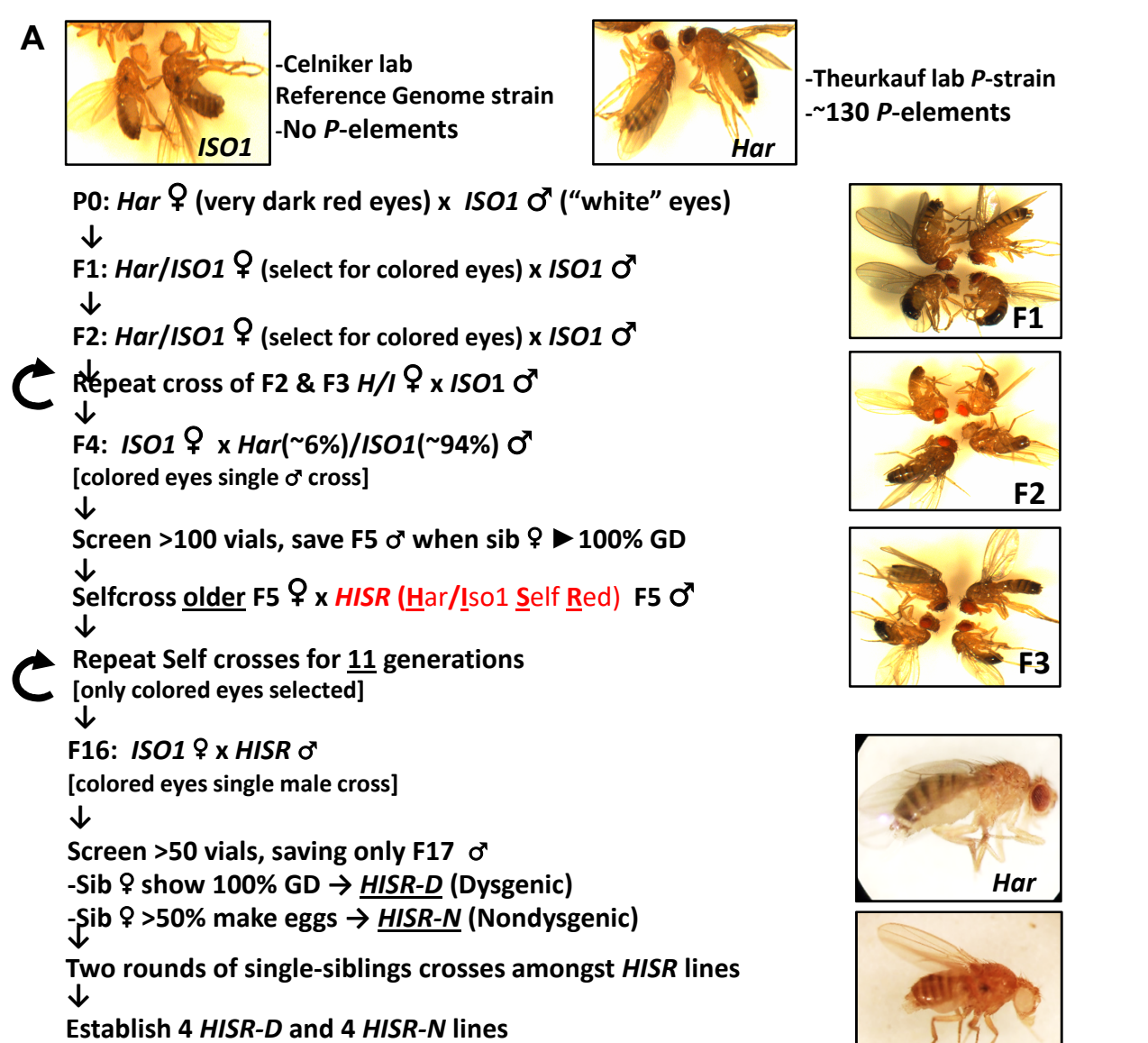
Hybrid dysgenesis is a fertility disease from missing P-elements & piRNAs



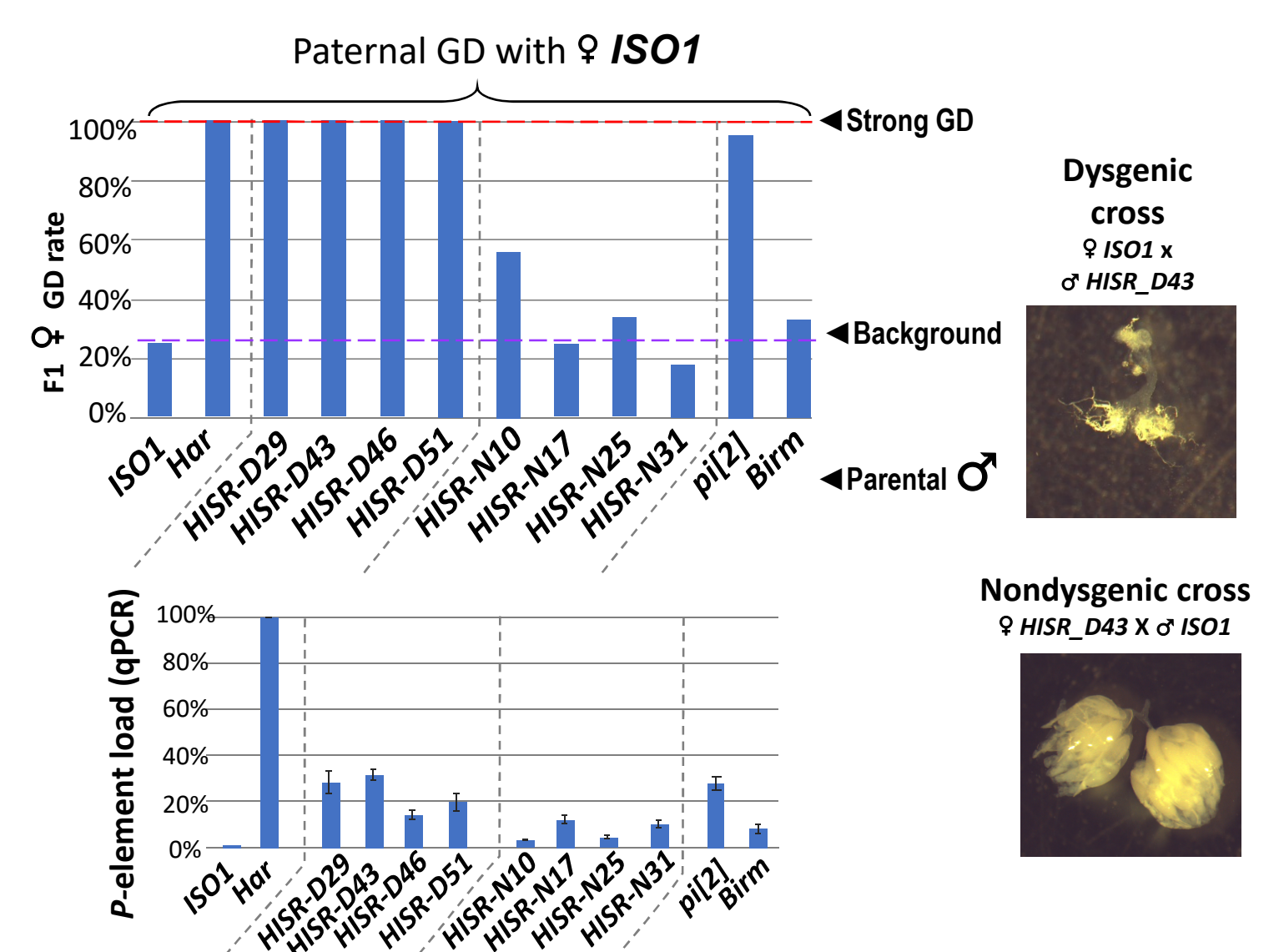
Diversity in P-element copy numbers and hybrid dysgenesis severity



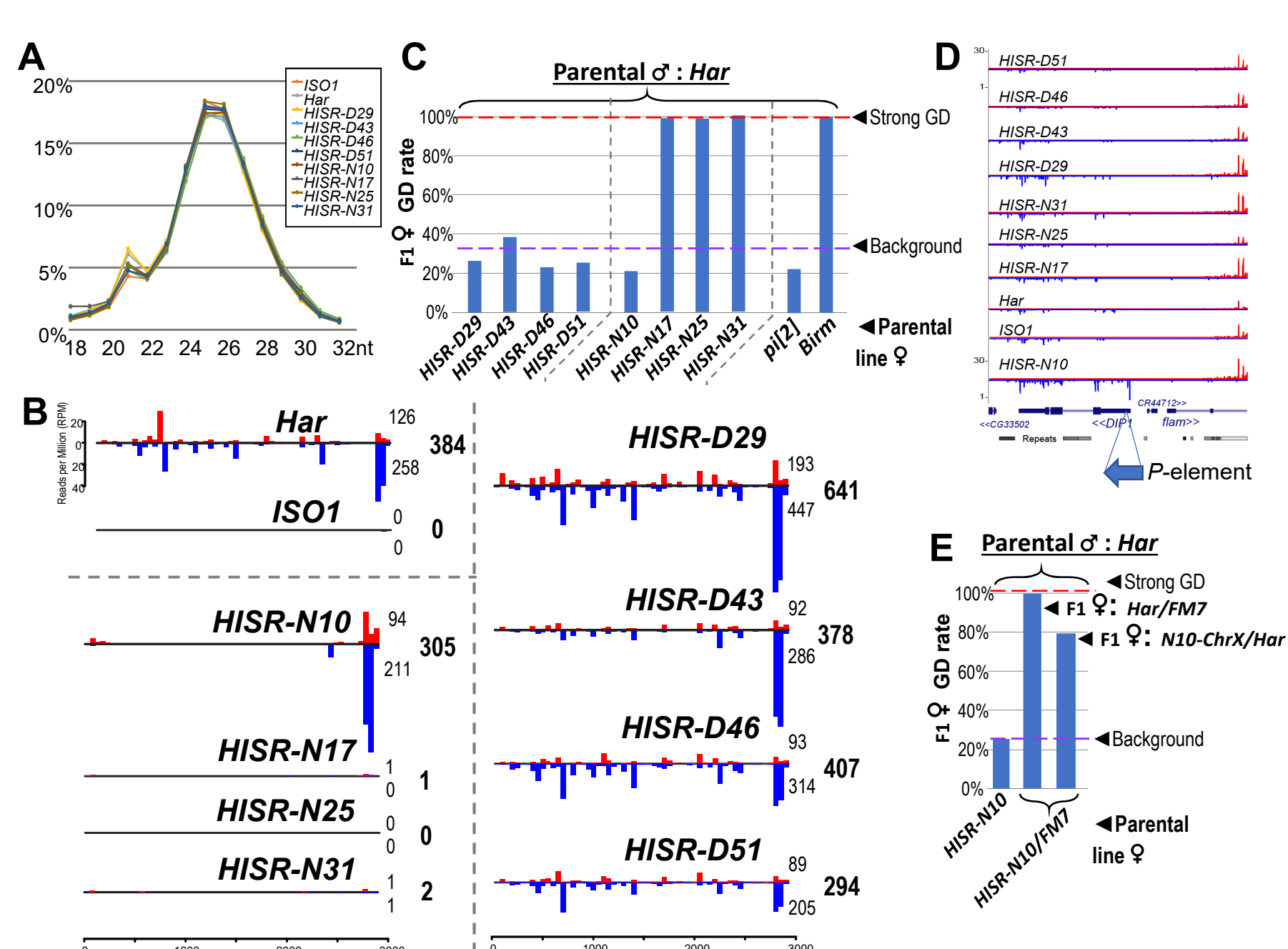
Can we genetically isolate a “Super” P-element from Harwich?



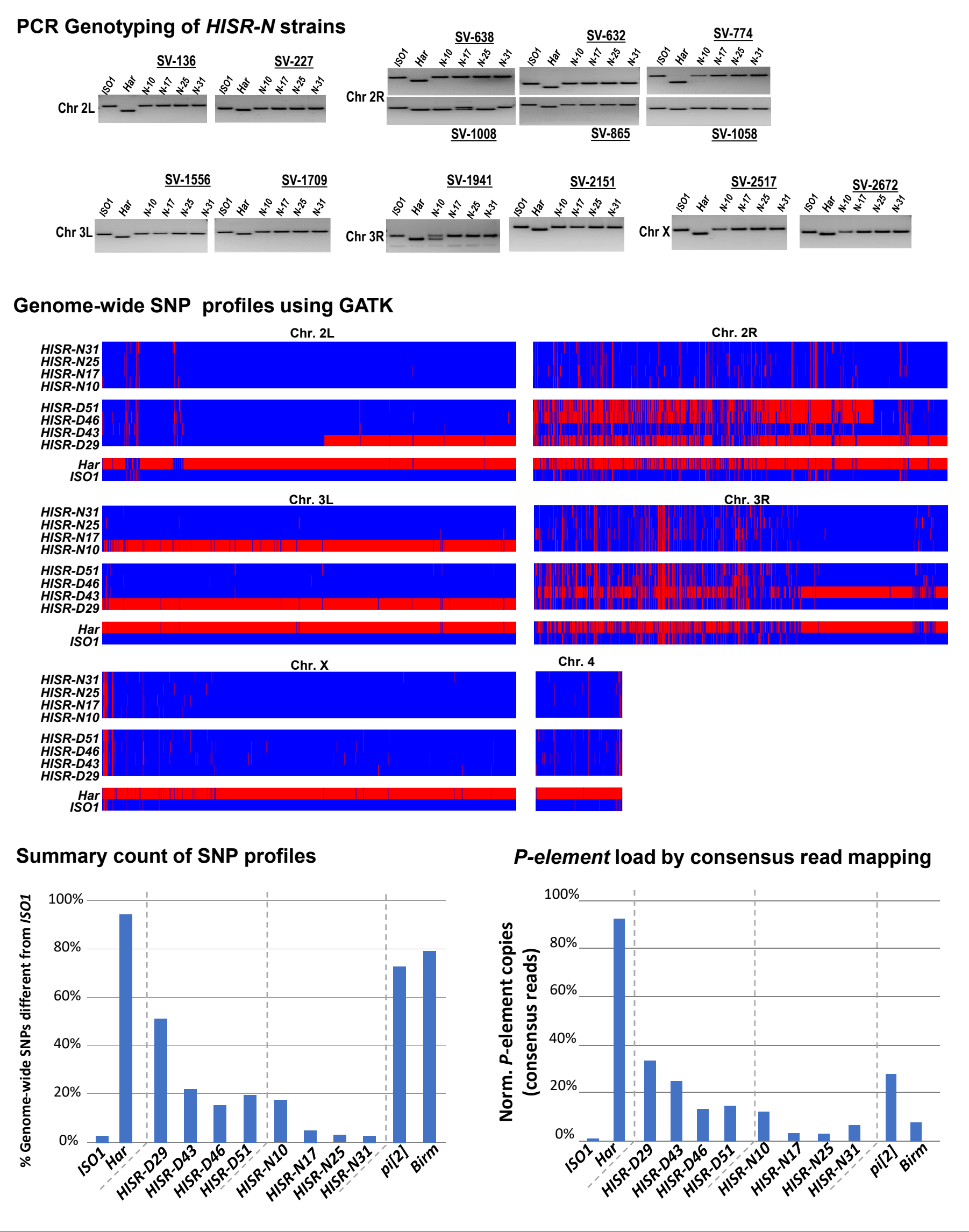
HISR hybrids with reduced P copies Strong GD (HISR-D) versus Low GD (HISR-N)



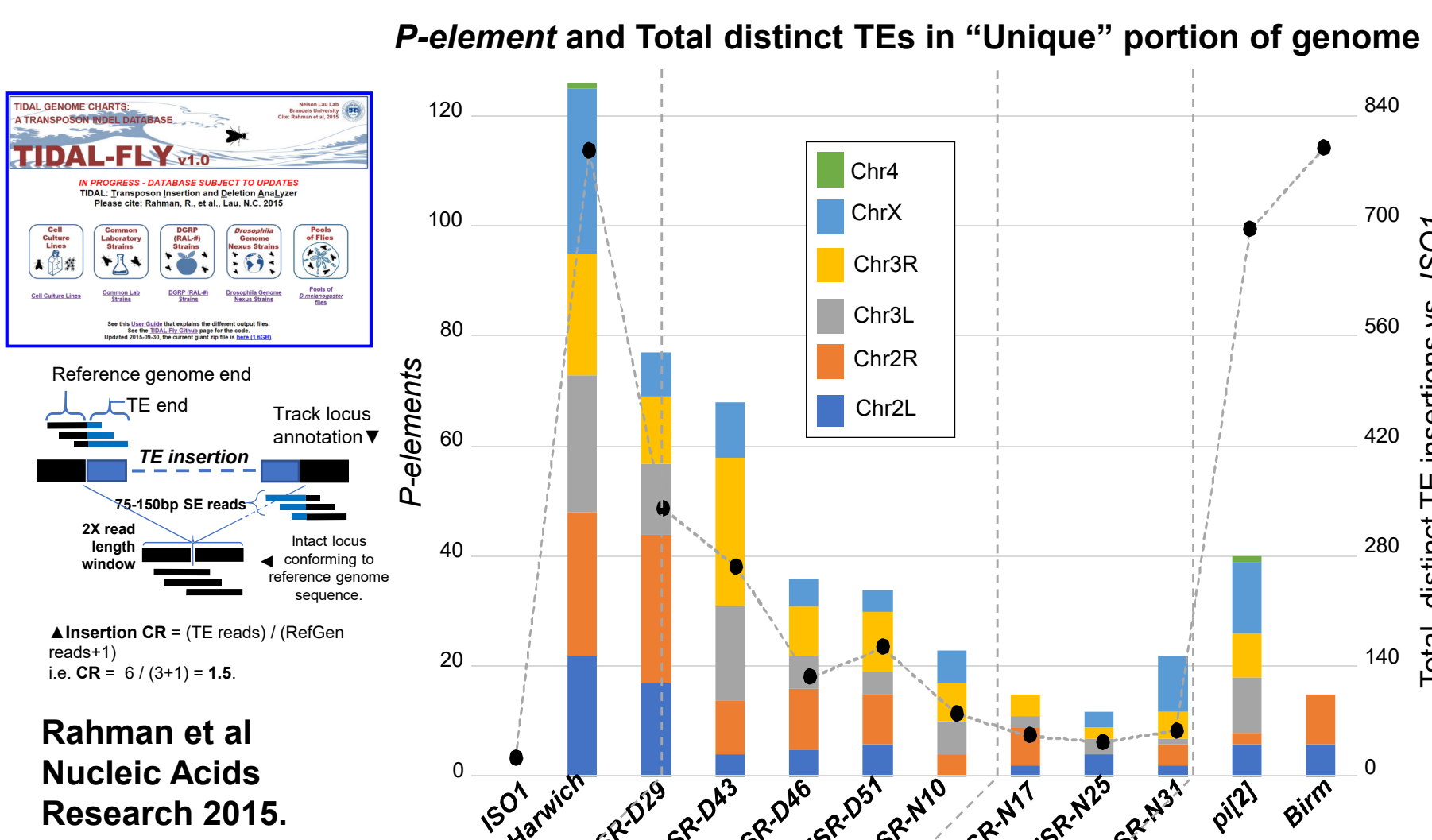
Altered P-element piRNA patterns in HISR hybrids



Whole genome sequencing of HISR hybrids



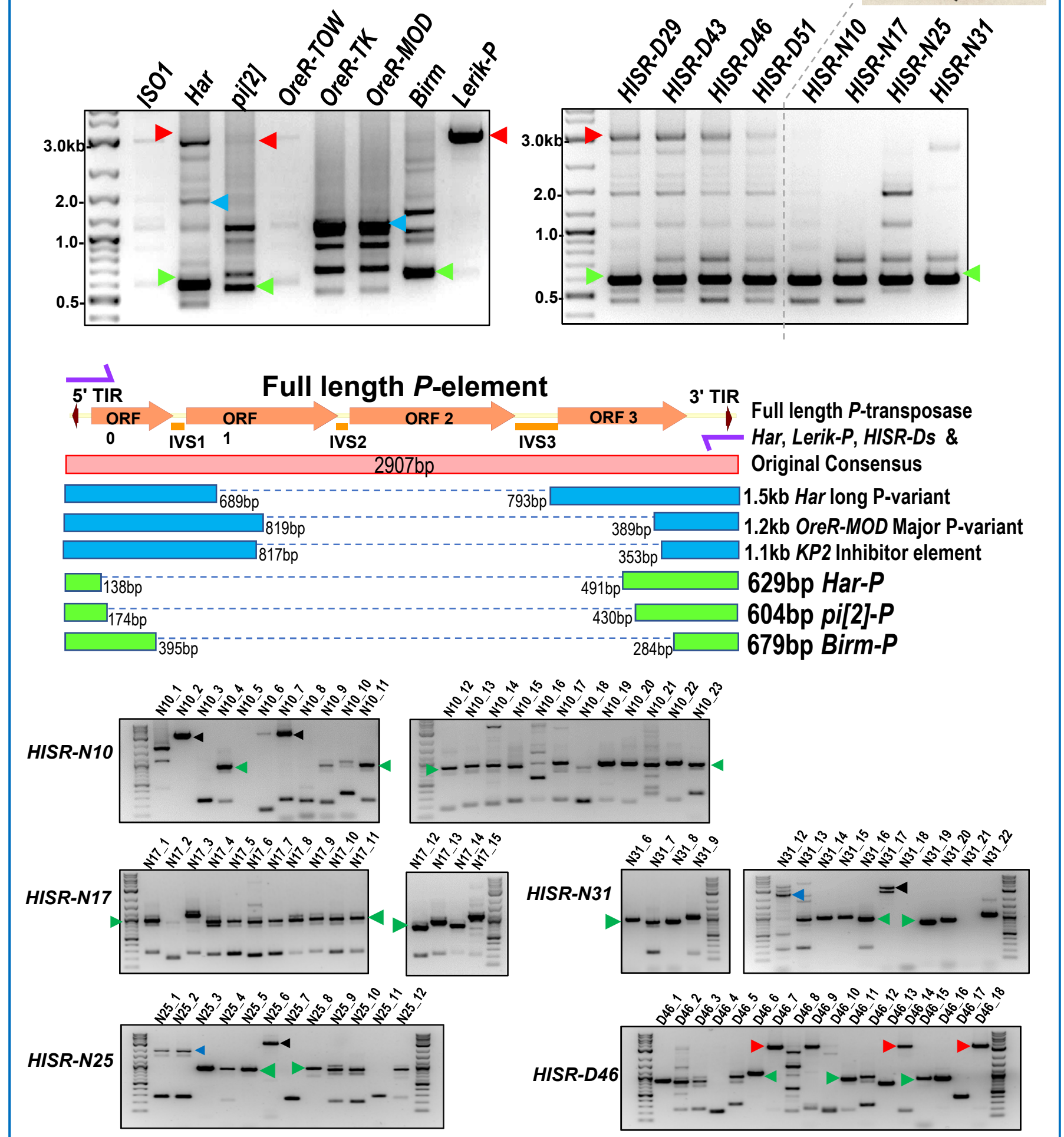
TIDAL-Fly analysis of TE landscape in HISR hybrids



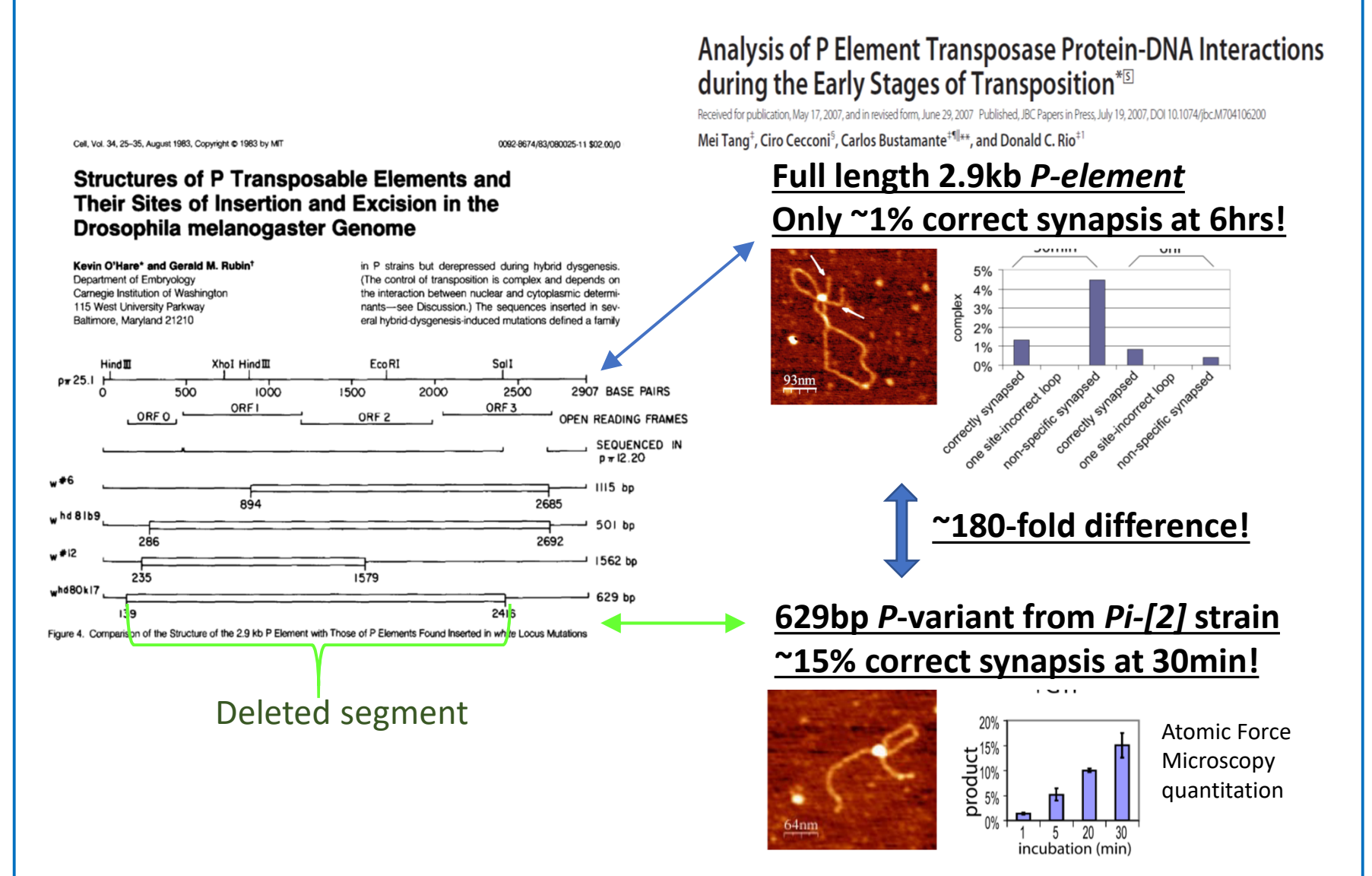
Low conservation of parental Harwich P's in HISRs - de-novo P-element mobilization

P-element copy #		Chr2L - 22	Chr2R - 26	Chr3L - 25	Chr3R - 23	ChrX - 30
127	Har					
77	HISR-D29	42				
68	HISR-D43	19	9			
36	HISR-D46	9	9	7		
34	HISR-D51	11	11	7	27	
23	HISR-N10	5	3	4	1	1
15	HISR-N17	0	1	1	0	0
12	HISR-N25	1	1	0	0	0
22	HISR-N31	2	3	1	0	1

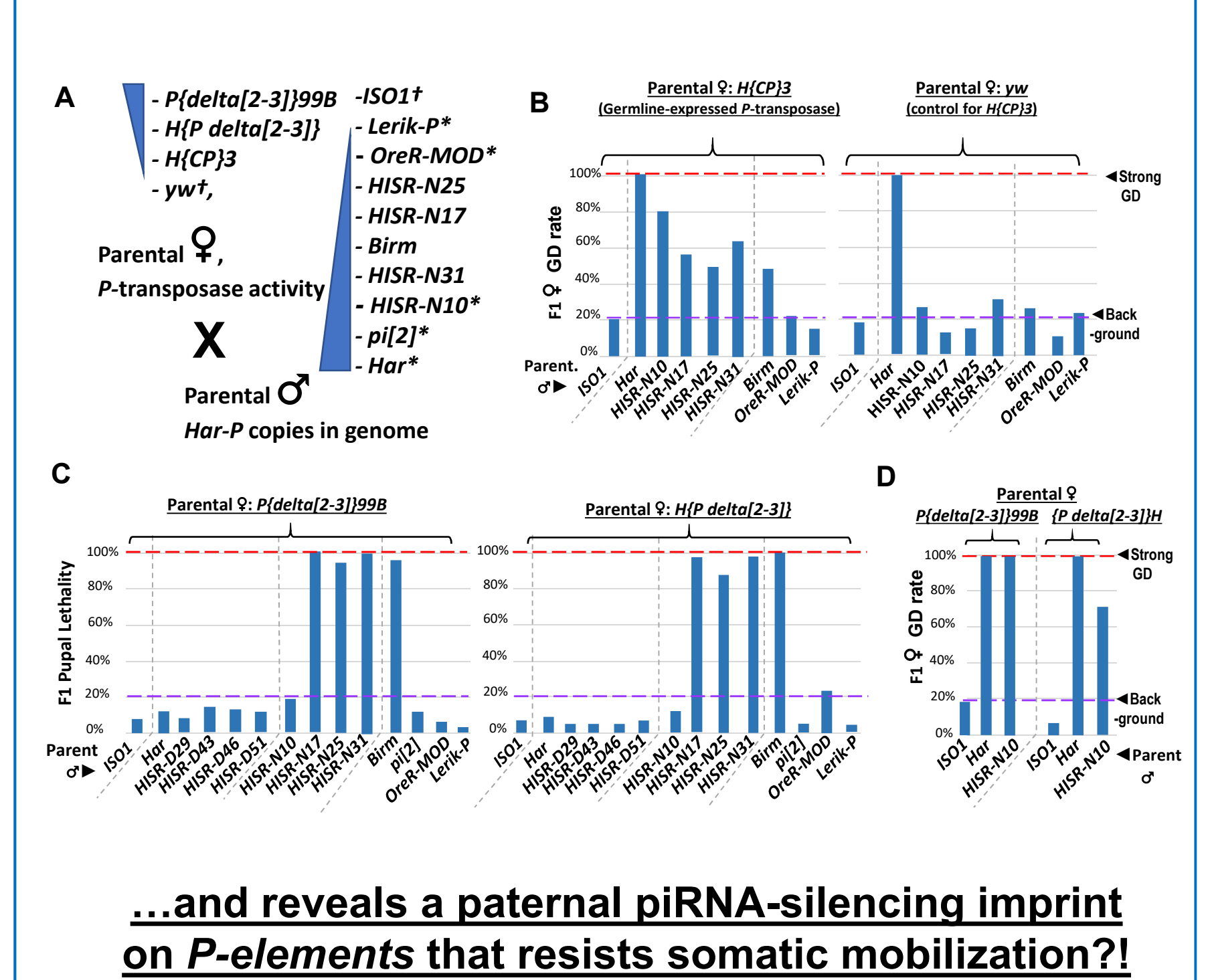
PCR and cloning in HISRs uncovers Har-P as a highly mobile short P-element variant



Earlier studies of naturally short P-element variants



Crossing back a Full-length P-transposase into HISR-N strains “restores” strong GD...



Full Length P-transposase is the bow; Har-P's are arrows that drive strong GD

