

Azole-resistant *Aspergillus fumigatus*, Iran

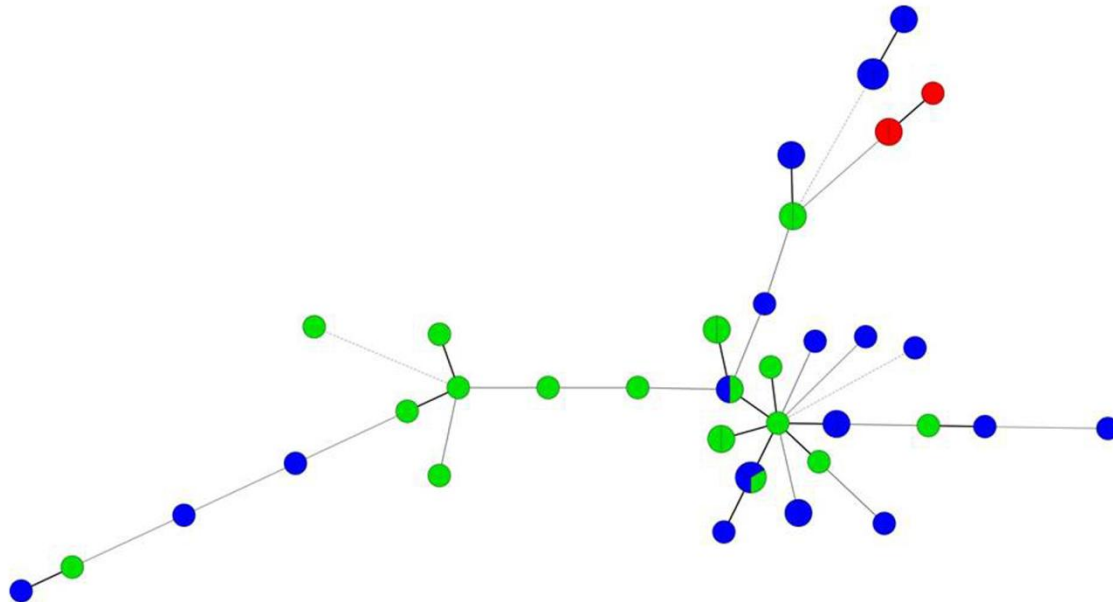
Technical Appendix

Technical Appendix Table 1. Distribution of azole-resistant and azole-susceptible *Aspergillus fumigatus* isolates, Iran, 2003–2009

Year of isolation	No. isolates with each phenotype and resistance mechanism		
	Wild type	Resistant, TR ₃₄ /L98H	Resistant, non-TR ₃₄ /L98H
2003	11	–	–
2004	18	–	–
2005	11	1	–
2006	17	–	–
2007	19	2	1
2008	20	–	–
2009	24	–	–
Total	120	3	1

Technical Appendix Table 2. First reports of multiple-triazole-resistant *Aspergillus fumigatus* isolate(s) carrying the TR₃₄/L98H mutations in the CYP51A gene, by country

Region/Country	First reported TR ₃₄ /L98H <i>A.fumigatus</i> isolate(s)	Reference	
Europe	Netherlands	1998	Snelders et al. 2008 (1)
	UK	1999	Howard et al. 2009 (2)
	Norway	2001	Snelders et al. 2008 (1)
	Spain	2002/2003	Mellado et al. 2012 (3)
	Denmark	2007	Mortensen et al. 2011 (4)
	Belgium	2008	Lagrou et al. 2008 (5)
	France	2010/2011	Burgel et al. 2012 (6), Morio et al. 2012 (7)
	Germany	2012	Rath et al. 2012 (8)
	Asia	Iran	2005
India		2008	Chowdhary et al. 2012 (9,10)
China		2008/2009	Lockhart et al. 2011 (11)



Technical Appendix Figure. Minimum spanning tree comparing genotypic relatedness of clinical azole-resistant *Aspergillus fumigatus* isolates carrying TR₃₄/L98H alteration in the *CYP51A* gene from Iran with those reported from European countries. Microsatellite typing of 6 STR loci demonstrated identical patterns for two of the three azole-resistant isolates from Iran, but the TR₃₄/L98H isolates from Iran did not cluster with those from the Netherlands and other European countries, indicating no close genetic relatedness. Each circle corresponds to a unique genotype, and each color indicates the origin of azole-resistant TR₃₄/L98H isolates: red, Iran (n = 3); green, the Netherlands (n = 20); blue, other European countries (n = 24). The size of the circle corresponds to the number of isolates with that genotype. Connecting lines correspond to the number of different microsatellite loci between the genotypes: solid thick and thin branches indicate 1 and 2 microsatellite marker differences, respectively; dashed branches indicate 3 microsatellite marker difference; dotted branches indicate ≥ 4 microsatellite marker differences between genotypes.

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