

UPDATE ON THE MEDICAL MANAGEMENT OF PSC

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HIGH-DOSE URSO FOR PSC RESULTS



Primary Endpoints

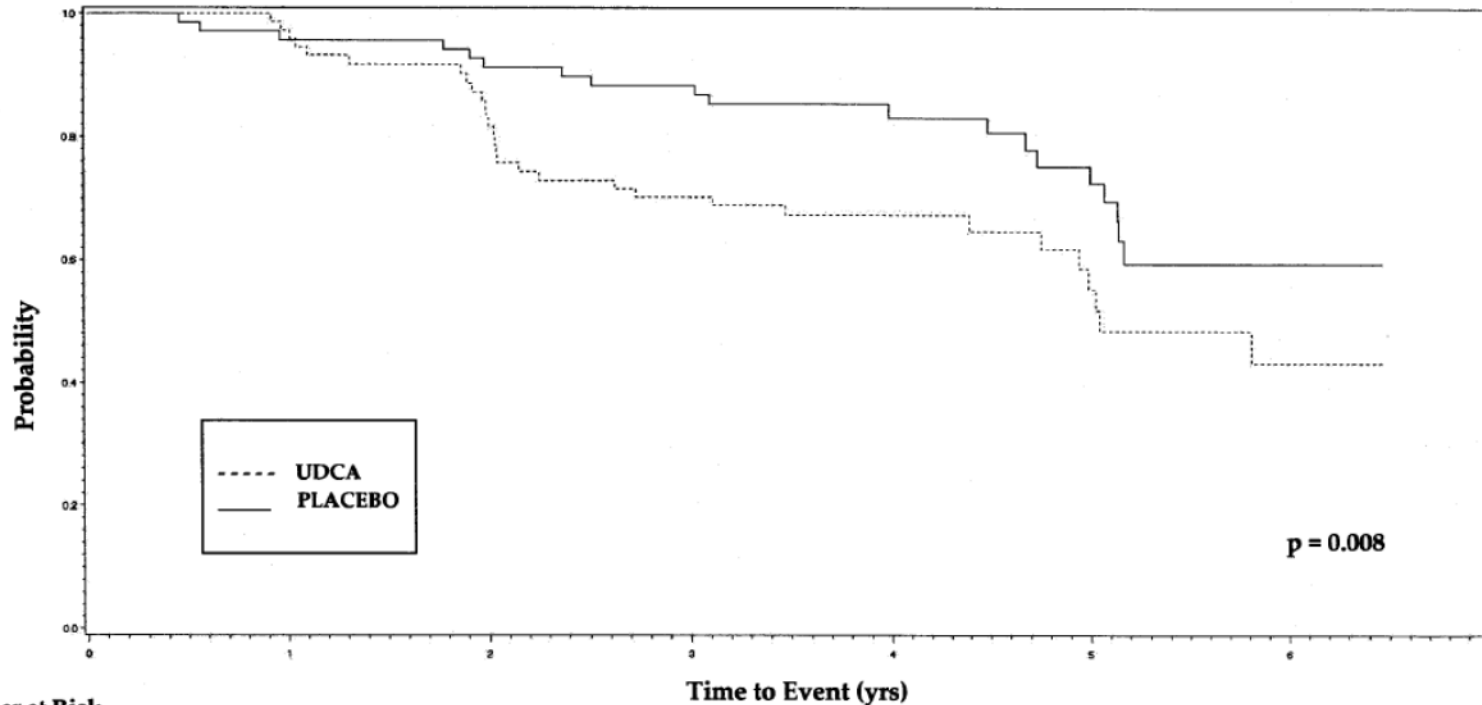
Primary Endpoints	UDCA	Placebo
Death	5	3
Liver Transplant	11	5
Minimal Listing Criteria for Liver Transplant	13	10
Development of Cirrhosis	6	4
Esophageal and/or Gastric Varices	15	5
Cholangiocarcinoma	2	2
Total Endpoints	52	29

Lindor KD, et al. Hepatology 2009;50(3):808-14

RESULTS



Model of All Primary Endpoints
Adjusted for Mayo Risk Score, Presence of Varices, and Stage

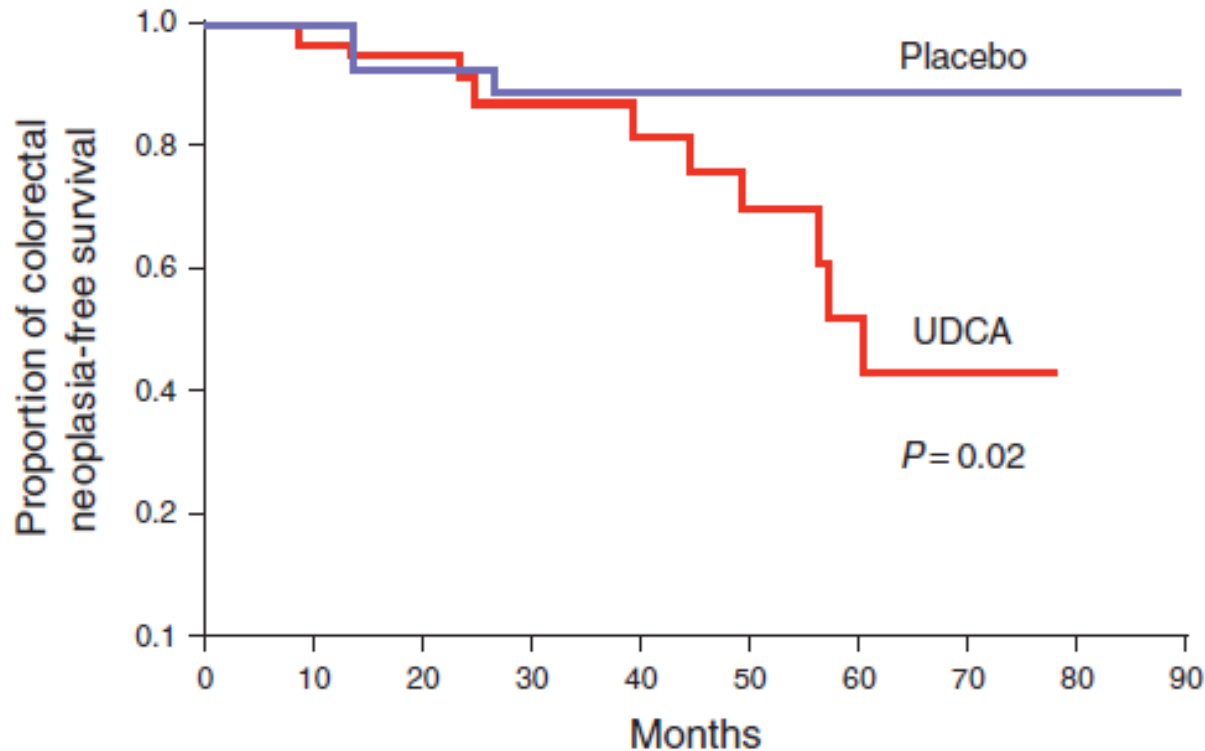


Number at Risk

UDCA	76	73	60	51	34	18	9	0
PLACEBO	74	65	60	58	41	24	7	0

Lindor KD, et al. Hepatology 2009;50(3):808-14

HIGH-DOSE URSO IN UC AND PSC PATIENTS



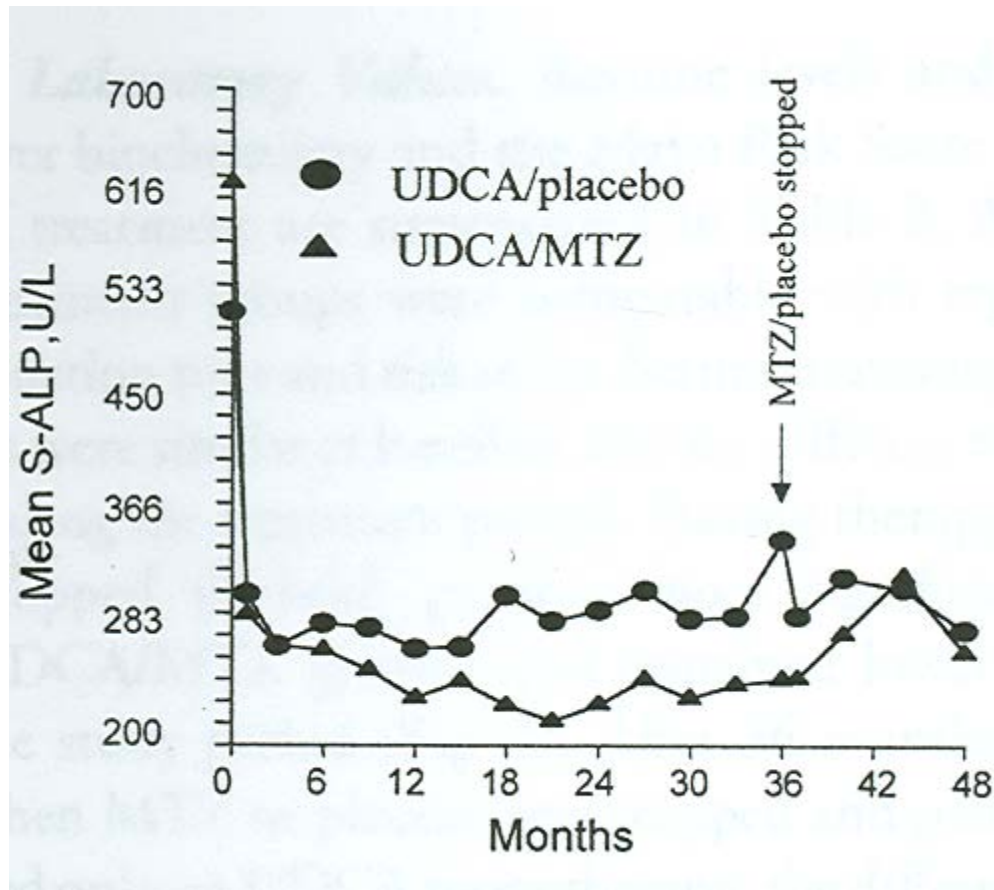
Eaton J, et al. Am J Gastroenterol 2011;106:1638-45

ANTIBACTERIAL AGENTS



- Metronidazole and UDCA for PSC
 - A randomized trial
 - 80 patients
 - UDCA (15 mg/kg/d)
 - UDCA and Metronidazole (600-800 mg/d)
 - 3 years

METRONIDAZOLE AND UDCA FOR PSC



Farkkila M, et al. Hepatology 2004;40:1379-86

OTHER ANTIBIOTICS IN PSC



Drug	# Months	% change from baseline		
		ALP	AST	ALT
Tetracycline (n=5)	1-10	-45	-60	-45
Vancomycin (short term) (n=3)	Mean 9	--	--	-89
Vancomycin (long term) (n=14)	54 ₊ 43	--	--	-78
Azithromycin (n=1)	5	-72	-31	-33

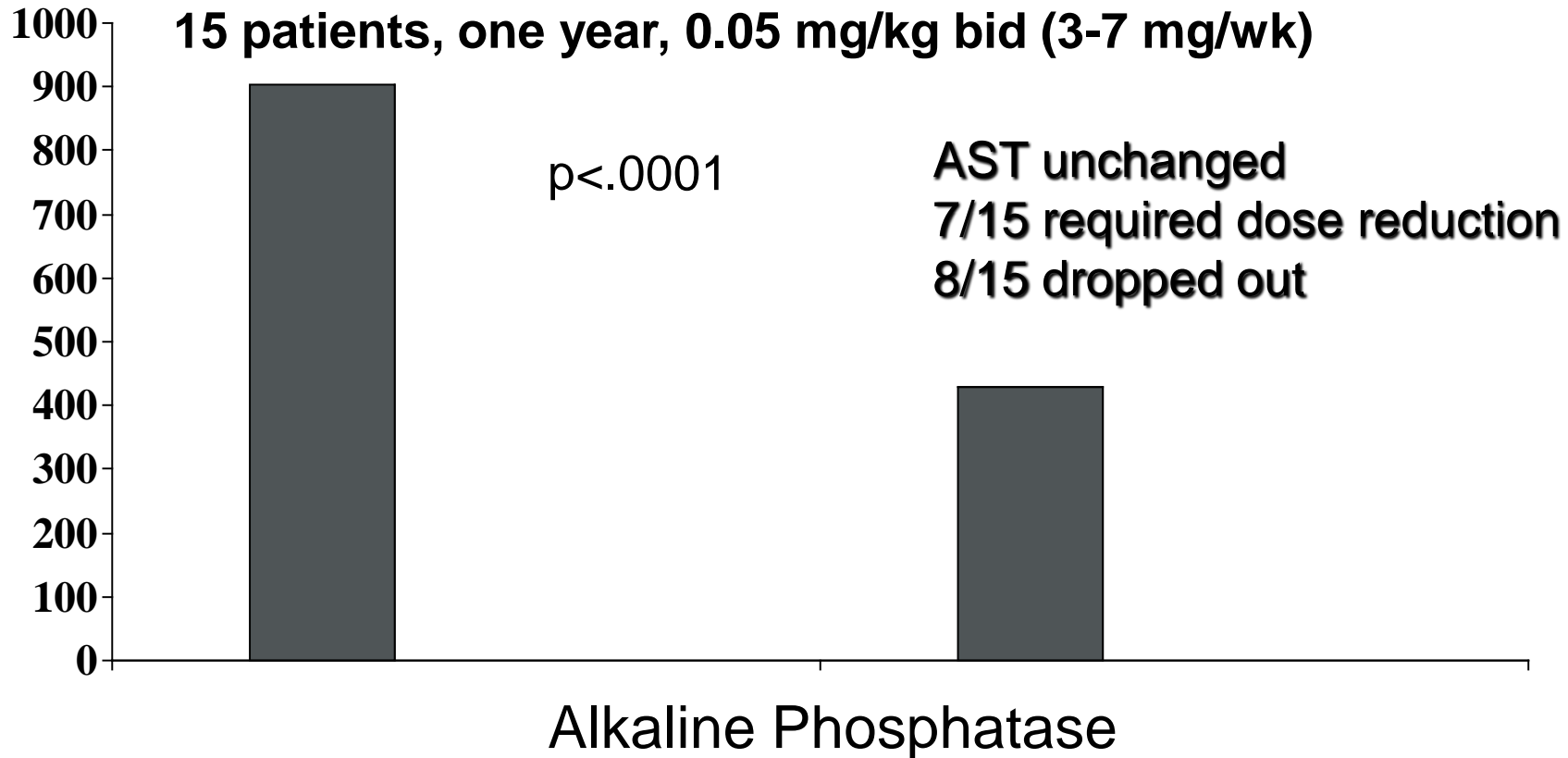
Elfaki D, Lindor KD. Am J Ther 2009 (in press)

IMMUNOLOGIC ABNORMALITIES



- Tacrolimus
- Budesonide
- IgG4

TACROLIMUS IN PSC

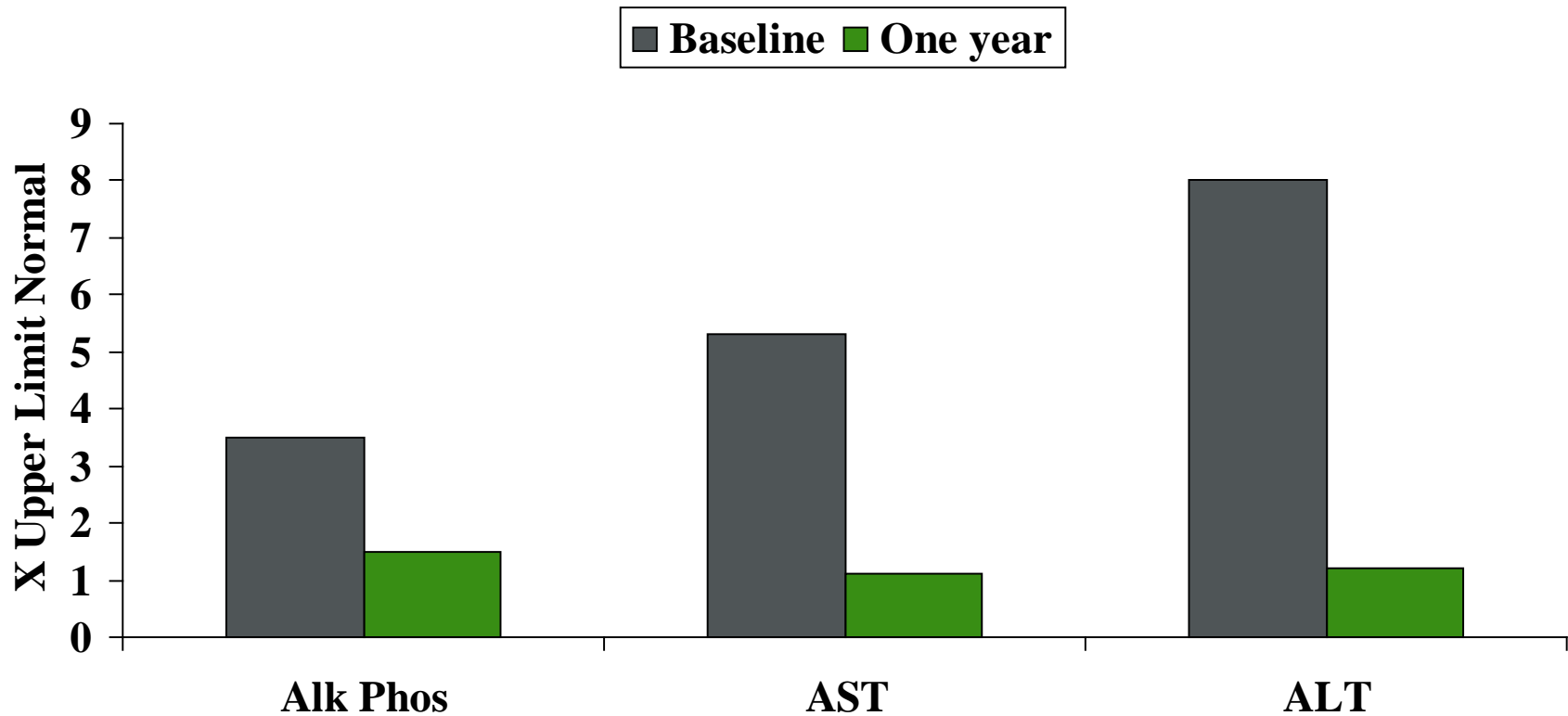


Talwalkar J. Gastroenterology 2005;128(4):A775

TACROLIMUS IN PSC



10 patients, 1 year, serum FK 506 level of 0.6 to 1.0 ng/ml



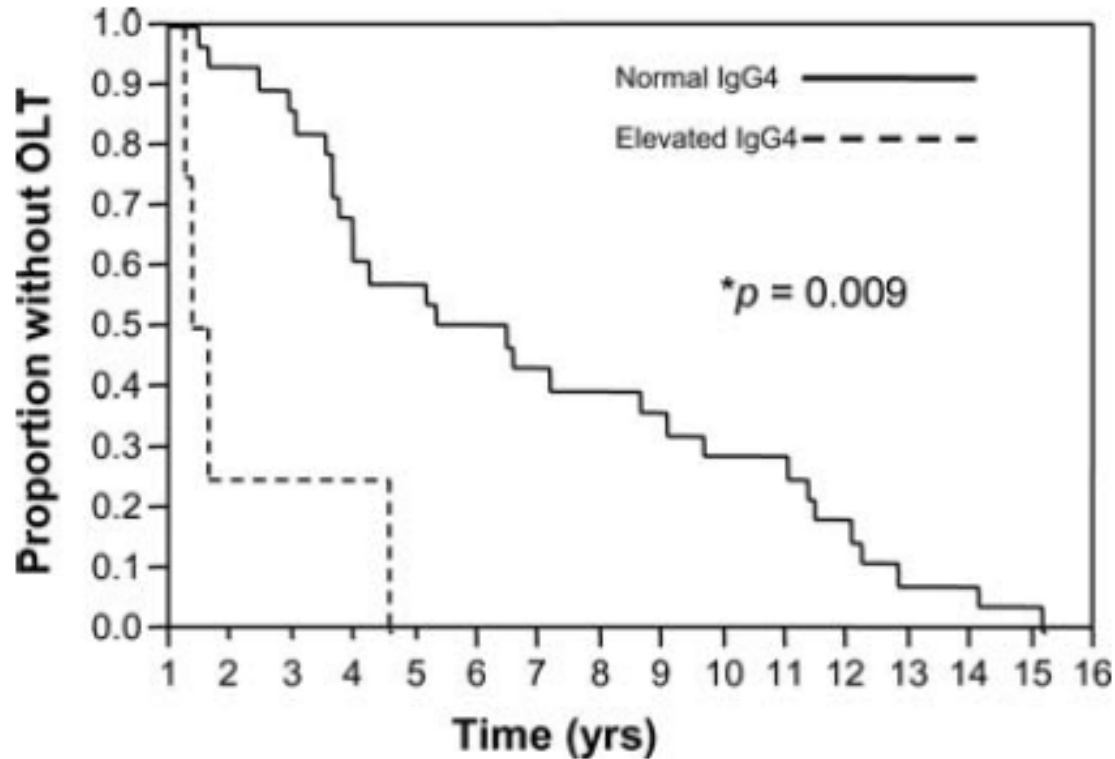
Van Thiel, et al. Am J Gastro 1005;90(3):455-459.

AUTOIMMUNE PANCREATITIS/ CHOLANGITIS IN PSC



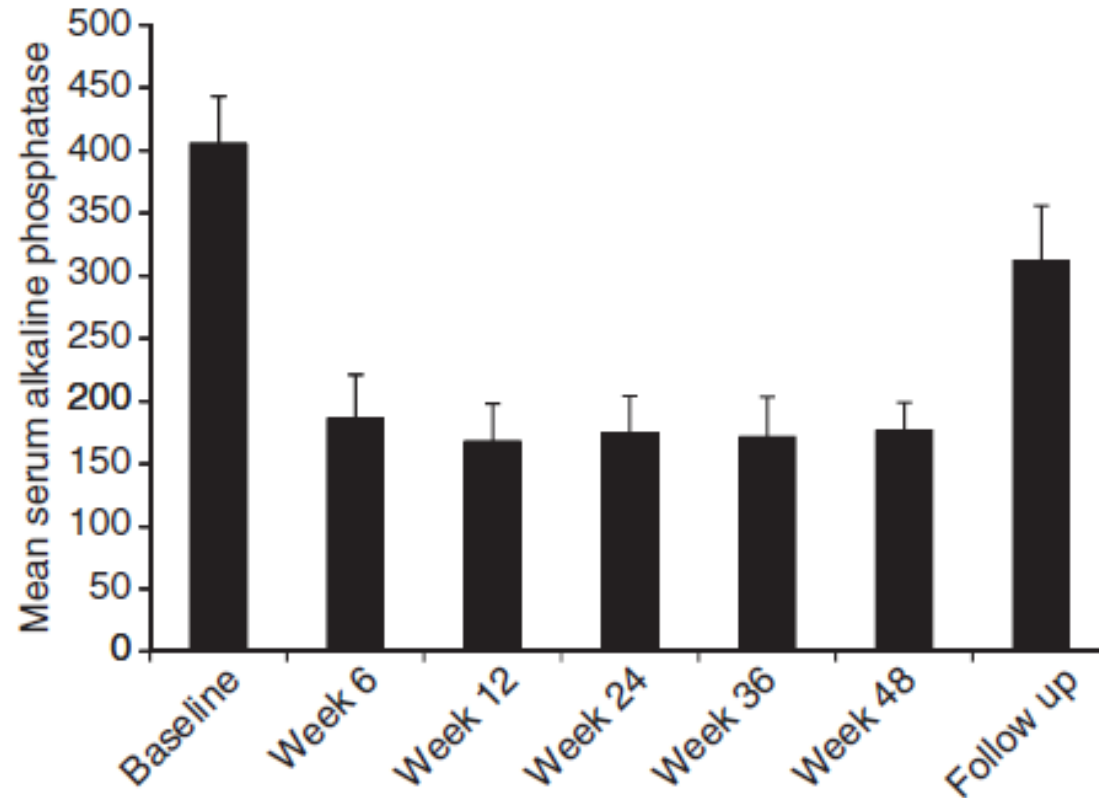
- IgG4 elevated in 9% PSC patients
- These patients may be more steroid responsive.

NATURAL HISTORY “PSC” & IgG4



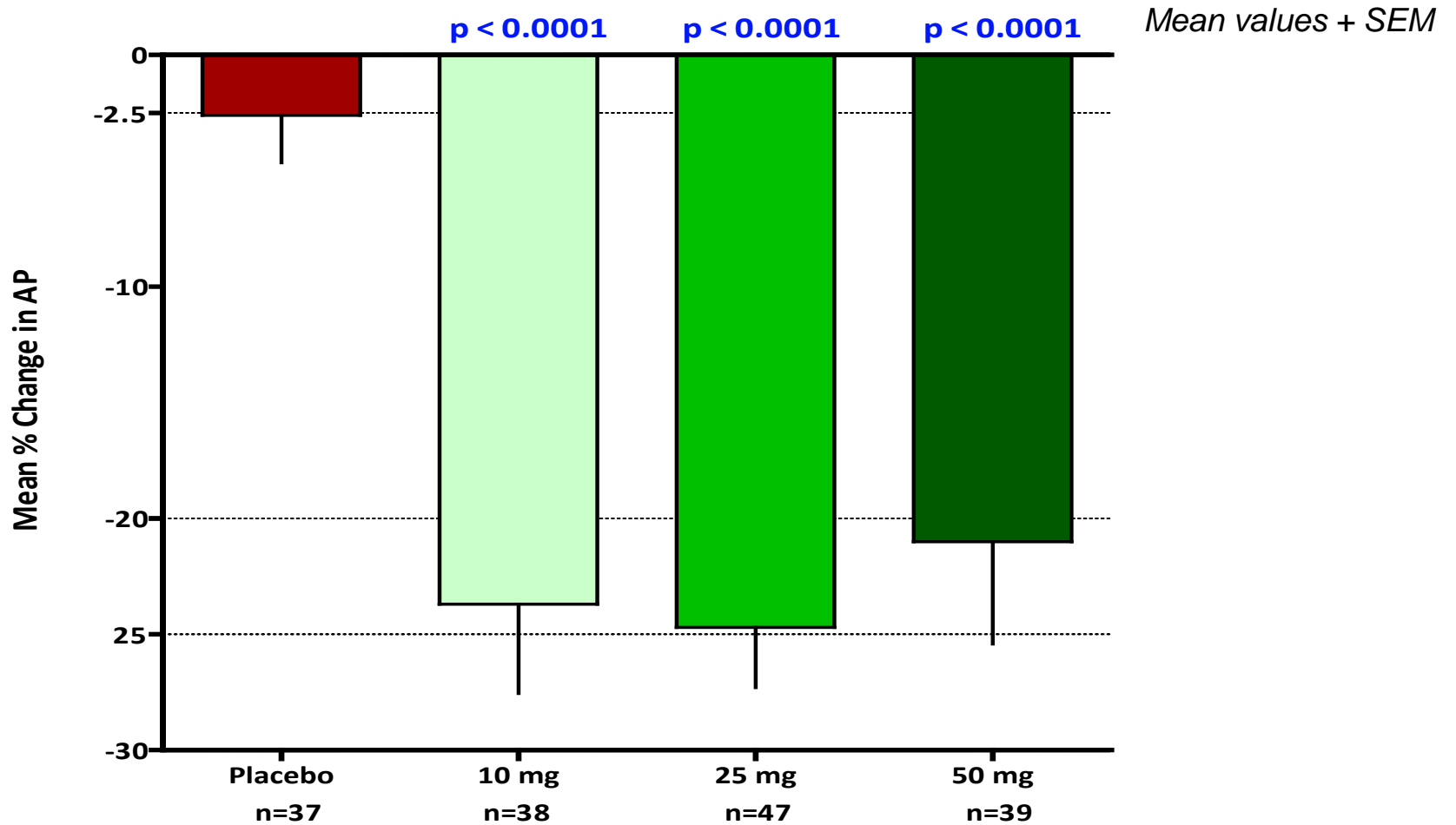
Mendes F, et al. Am J Gastroenterol 2006;101:2070-75

FENOFIBRATE FOR PBC



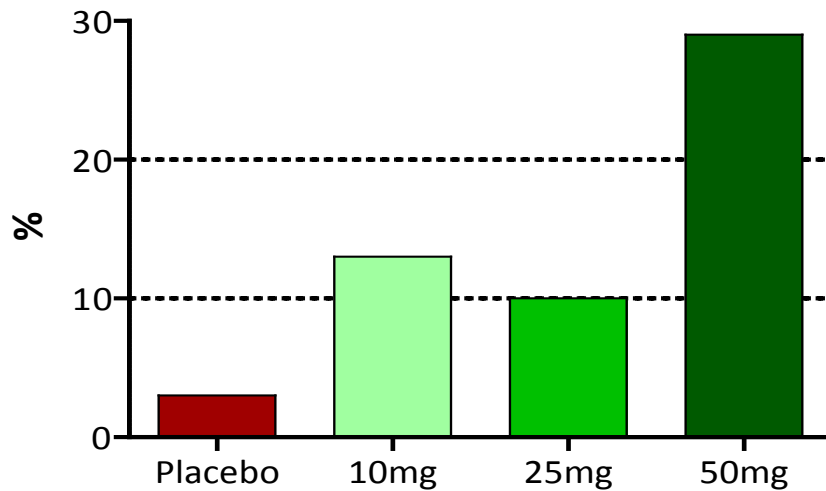
Levy C, et al. Aliment Pharmacol Ther 2011;33:235-42

% Δ – Alkaline Phosphatase

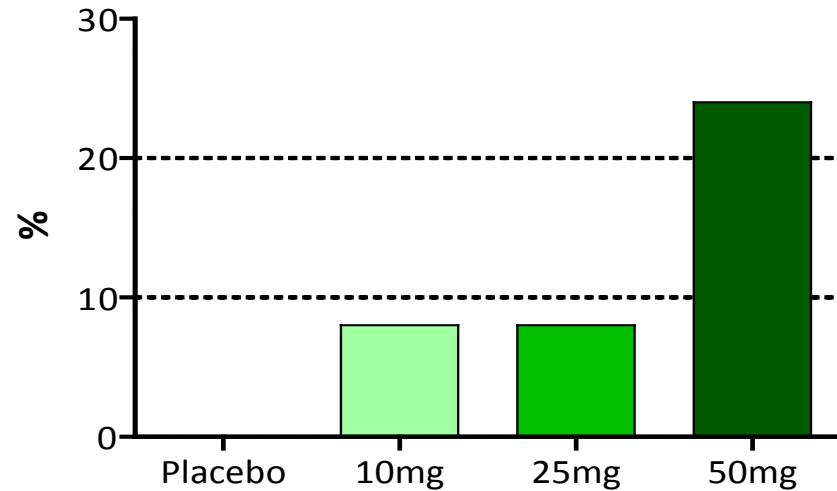


Discontinuations for AEs

Any AE



Pruritus



Conclusion



- No definitive therapy