

Correction to Radiolabeled Absorption, Distribution, Metabolism, and Excretion Studies in Drug Development: Why, When, and How?

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Ref 93 should be as follows: (93) Prakash, C., Cui, D., Baxter, J. G., Bright, G. M., Miceli, J., and Wilner, K. (1998) Metabolism and excretion of a new antianxiety drug candidate, CP-93,393, in healthy male volunteers. *Drug Metab. Dispos.* 26, 448–456.

Table 3 was corrected.

Table 3. Comparison of Exposure of Human Metabolites of a Drug Candidate with Preclinical Species

analyte	mean % of circulating radioactivity			mean exposure (AUC) (ng equiv-h/mL)		
	human ^b	rat ^c	monkey ^d	human ^b	rat ^c	monkey ^d
parent	9.80	31.5	3.23	89.5	25500	388
M7	2.50	10.4		22.6	8410	0
M9	3.70	4.40	0.96	33.3	3560	115
M13	8.80	7.60	25.8	80.6	6150	3100
M15	54.2	24.3	43.1	494	19700	5180
M18 ^a	10.3		17.0	94.0	0	2040
M19	1.60			14.6	0	0

^aM18 was coeluted with M7 in monkeys. ^bPrakash, C., Cui, D., Baxter, J.G., Bright, G.M., Miceli, J., and Wilner, K. (1998) Metabolism and excretion of a new antianxiety drug candidate, CP-93,393, in healthy male volunteers. *Drug Metab. Dispos.* 26, 448–456. ^cPrakash, C., and Soliman, V. (1997) Metabolism and excretion of a new anxiolytic drug candidate, CP-93,393, in Long-Evans rat. *Drug Metab. Dispos.* 25, 1288–1297. ^dPrakash, C., and Cui, D. (1997) Metabolism and excretion of a new anxiolytic drug candidate, CP-93,393, in cynomolgus monkey: Identification of the novel pyrimidine ring cleaved metabolites. *Drug Metab. Dispos.* 25, 1395–1406.

Metabolites M15 and M18 were the major circulating metabolites accounting for 54.2% and 10.3% of the circulating radioactivity, respectively, whereas M7, M9, and M13 accounted for only 2.5, 3.7, and 8.8% of the total circulating radioactivity, respectively. M19 was present at 1.6% of the total radioactivity (Table 3). In the monkey plasma, M15 and M18+M7 constituted 43.1% and 17.0% of the radioactivity. The major circulating metabolite in rat plasma was M15 and represented 24.3% of the total circulating radioactivity, whereas other metabolites (M7, M9, M13, and M10) accounted for 4.4 to 10.4% of the circulating radioactivity. M18 was not detected in the rat (Table 3).

Using eq 3, the mean exposures (AUC) of M15 and M18 were 494 and 94 ng-equiv-h/mL, respectively, in humans, and the estimated corresponding exposures in the monkey were 5080 and 2040 (+M7) ng-equiv-h/mL, respectively. Metabolite M18 was not present in the rat; however, the estimated AUC of M15 in the rat was 19700 ng-equiv-h/mL.

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