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		10 20 30 40 50	
AKR1C2	1	MDSKYQCVKLNDGHFMPVLGFGTYAPAEVPKSKALEAVKLAIEAGFHHIDSAHVYNNEEQ MDSK+OCVKLNDGHFMPVLGFGTYAP EVP+SKALE KLAIEAGF HIDSAH+YNNEEO	60
AKR1C3	1	MDSKHQCVKLNDGHFMPVLGFGT <mark>Y</mark> APPEVPRSKALEVTKLAIEAGFRHIDSAHL <mark>Y</mark> NNEEQ	60
AKR1C2	61	VGLAIRSKIADGSVKREDIFYTSKLWSNSHRPELVRPALERSLKNLQLDYVDLYLIHFPV VGLAIRSKIADGSVKREDIFYTSKLWS HRPELVRPALE SLK QLDYVDLYLIH P+	120
AKR1C3	61	VGLAIRSKIADGSVKREDIFYTSKL <mark>W</mark> STFHRPELVRPALENSLKKAQLDYVDLYLI <mark>H</mark> SPM	120
AKR1C2	121	SVKPGEEV <mark>I</mark> PKDENGKILFDTVDLCATWEAMEKCKDAGLAKSIGVSNFNHRLLEMILNKP S+KPGEE+ P DENGK++FD VDLC TWEAMEKCKDAGLAKSIGVSNFN R LEMILNKP	180
AKR1C3	121	SLKPGEEL <mark>S</mark> PTDENGK <mark>V</mark> IFDIVDLCTTWEAMEKCKDAGLAKSIGVSNFNRRQLEMILNKP	180
AKR1C2	181	GLKYKPVCNQVECHPYFNQRKLLDFCKSKDIVLVAYSALGSHREEPWVDPNSPVLLEDPV GLKYKPVCNQVECHPYFN+ KLLDFCKSKDIVLVAYSALGS R++ WVDPNSPVLLEDPV	240
AKR1C3	181	GLKYKPVCNQVECHPYFNRSKLLDFCKSKDIVLVAYSALGSQRDKRWVDPNSPVLLEDPV	240
AKR1C2	241	LCALAKKHKRTPALIALRYQLQRGVVVLAKSYNEQRIRQNVQVFEFQLTSEEMKAIDGLN LCALAKKHKRTPALIALRYQLQRGVVVLAKSYNEQRIRQNVQVFEFQLT+E+MKAIDGL+	300
AKR1C3	241	LCALAKKHKRTPALIALRYQLQRGVVVLAKSYNEQRIRQNVQVFEFQLTAEDMKAIDGLD	300
AKR1C2	301	RNVRYLT <mark>l</mark> d <mark>i</mark> fagppnypfsdey 323 rn+ y d fa pnyp+sdey	
AKR1C3	301	RNLHY <mark>FNS</mark> D <mark>SF</mark> ASHPNYPYSDEY 323	

Supplemental. Sequence alignment of human AKR1C2 and AKR1C3 using the 'align two or more sequences' feature of protein BLAST (<u>https://blast.ncbi.nlm.nih.gov</u>). AKR1C3 shares 91% sequence similarity and 87% identity with AKR1C2. Identical residues are shown in the consensus sequence between the two compared sequences. Similar residues are marked with a '+'. Residues in contact with compound **2** in the structure of **AKR1C3·NADP+·2** are shown in bold and highlighted in yellow. Of these 11 residues, 3 residues that are different in AKR1C2 are shown in red and highlighted in yellow.