

Table I

Basic information of included studies

Author, Year	Study design	Intervention	Control	Outcome				
				Open field test (OFT)	Tail suspension test (TST)	Serotonin	Dopamine	Noradrenaline
Dudau et al., 2023	Animal study, rat, 21 days	Group I: Control, vehicle per oral	Group IV: Reserpine (1.5 mg/kg) ip	Reserpinised rat: 0.6 ± 1.2 Control: 4.6 ± 4.6, p=0.004	NI	NI	NI	NI
		Group II: CUR (150 mg/kg) ip	Group V: Reserpine and CUR (150 mg/kg) ip					
		Group III: Escitalopram (20 mg/kg) ip	Group VI: Reserpine (1.5 mg/kg)and escitalopram (20 mg/kg) ip					
Chang et al., 2016	Animal study, rat (n=30), 45 days	Group I: Sham control Group II: Olfactory bulbectomy (lesion) Group II: CUR (10 mg/kg) Group III: CUR (20 mg/kg) Group IV: CUR (40 mg/kg)	Group I: Sham control Group II: Olfactory bulbectomy (lesion) Group II: CUR (10 mg/kg) Group III: CUR (20 mg/kg) Group IV: CUR (40 mg/kg)	CUR treatment reduced the number peeping and rearing 27.1, 24.1 and 18.6 at 10, 20 and 40 mg/kg respectively Sham: 15.1 Olfactory bulbectomy: 30.3	NI	CUR treatment significantly increased the serotonin level 162.3, 210.3 and 257.7 ng/g at 10, 20 and 40 mg/kg respectively Sham: 279.2 Olfactory bulbectomy: 141.4 ng/g	CUR treatment significantly increased the dopamine level 247.4, 265.8 and 290.6 ng/g at 10, 20 and 40 mg/kg respectively Sham: 315 Olfactory bulbectomy: 215.5 ng/g	CUR treatment significantly increased the noradrenaline level 273, 299.5 and 335 ng/g at 10, 20 and 40 mg/kg respectively Sham: 351 Olfactory bulbectomy: 250.5 ng/g olfactory
CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (<i>Cochlospermum religiosum</i>); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of <i>Aegle marmelos</i> leaves); EEGS (Ethanol extract from <i>G. procumbens</i> stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (<i>Rosmarinus officinalis</i> extract); SNE (<i>Synedrella nodiflora</i> extract)								

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Gazal et al., 2013	Animal study, female rat (n=30), 15 days	Group I: Control, vehicle per oral	Group I: Control, vehicle per oral	CUR pretreatment: [F(2,26)=4.11, p<0.05]	NI	NI
		Group II: Ketamine (25 mg/kg)	Group II: Ketamine (25 mg/kg)	Ketamine treatment: [F(1,26)=16.93, p<0.05]		
		Group III: CUR 20 mg/kg	Group III: CUR 20 mg/kg	Interaction: [F(2,26)=7.99, p<0.05]		
		Group IV: Ketamine (25 mg/kg)/CUR (50 mg/kg)	Group IV: Ketamine (25 mg/kg)/CUR (50 mg/kg)	Lithium chloride pretreatment: [F(1,23)= 8.4, p<0.01]		
		Group V: Saline 45 mg/kg twice a day	Group V: Saline 45 mg/kg twice a day	Ketamine treatment: [F(1,23)=23.08, p<0.01]		
		Group VI: Lithium chloride (45 mg/kg) twice a day	Group VI: Lithium chloride (45 mg/kg) twice a day	Interaction: [F(1,23)=13.99, p<0.01]		
CBD (Cannabidiol), CCI (Chronic constriction injury), CSR (Cochlospermum religiosum); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of <i>Agave marmelos</i> leaves); EEGS (Ethanol extract from <i>G. procumbens</i> stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (Rosmarinus officinalis extract); SNE (Synedrella nodiflora extract)		From the 8th to the 14th day the animals also received saline or ketamine (25 mg/kg) once a day, total 4 experiments	From the 8th to the 14th day the animals also received saline or ketamine (25 mg/kg) once a day, total four experimental	No changes in the anxiety behavior were observed for % central crossings after ketamine treatment, (CUR pretreatment: [F(2,36)=0.60, p=0.55] Ketamine treatment: [F(1,36)=0.28, p=0.59] Interaction: [F(2,36)=0.25, p=0.77]) and (Lithium chloride pretreatment: [F(1,20)=0.005, p=0.93] Ketamine treatment: [F(1,20)=0.60, p=0.44]		

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Pan et al., 2021	Animal study, male mice (n=30), 5 days	J147 as derivative CUR dissolved in DMSO and diluted with polyethylene glycol 660 hydroxy stearate solution as the working solution or vehicle Imipramine (10 mg/kg) J147 2.5 mg/kg J147 5 mg/kg J147 10 mg/kg	J147 as derivative CUR dissolved in DMSO and diluted with polyethylene glycol 660 hydroxy stearate solution as the working solution or vehicle Imipramine (10 mg/kg) J147 2.5 mg/kg J147 5 mg/kg J147 10 mg/kg	NI	NI	Control: 363.3 ± 15.4 J147 2.5 mg/kg: 398.8 ± 21.4 J147 5 mg/kg: 444.1 ± 23.6 J147 10 mg/kg: 461.8 ± 22.2 Imipramine 10 mg/kg: 466.3 ± 14.0	Control: 61.7 ± 2.9 J147 2.5 mg/kg: 63.7 ± 7.2 J147 5 mg/kg: 63.8 ± 5.9 J147 10 mg/kg: 66.5 ± 6.6 Imipramine 10 mg/kg: 62.3 ± 6.9	Control: 238.2 ± 13.6 J147 2.5 mg/kg: 262.1 ± 11.5 J147 5 mg/kg: 297.3 ± 18.1 J147 10 mg/kg: 311.5 ± 20.1 Imipramine 10 mg/kg: 300.6 ± 16.7
Arora et al., 2011	Animal study, rat (n=48), 5 days	Group I: Control, vehicle per oral Group II: Reserpine (1 mg/kg) sc for 3 consecutive days Group III, IV, V: Reserpinised rats receiving CUR (100, 200 and 300 mg/kg) ip for 2 days after reserpine (i.e. day 4 and 5) Group VI: CUR (300 mg/kg) ip	Group I: Control, vehicle per oral Group II: Reserpine (1 mg/kg) sc for 3 consecutive days Group III, IV, V: Reserpinised rats receiving CUR (100, 200 and 300 mg/kg) ip for two day after reserpine (i.e. day 4 and 5) Group VI: CUR (300 mg/kg) ip	NI	NI	Reserpine: 8.2 ± 0.04 Reserpine + CUR 10: 1.6 ± 0.0 Reserpine + CUR 20: 2.5 ± 0.0 Reserpine + CUR 30: 4.3 ± 0.0 CUR 30: 8.3 ± 0.0	Reserpine: 2.1 ± 0.01 Reserpine + CUR 10: 0.8 ± 0.0 Reserpine + CUR 20: 1.0 ± 0.0 Reserpine + CUR 30: 1.1 ± 0.0 CUR 30: 2.1 ± 0.0	NI

CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (*Cochlospermum religiosum*); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of *Aegle marmelos* leaves); EEGS (Ethanol extract from *G. procumbens* stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (*Rosmarinus officinalis* extract); SNE (*Synedrella nodiflora* extract)

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				Open field test (OFT)	Tail suspension test (TST)	Serotonin	Dopamine	Noradrenaline
Zhao et al., 2014	Animal study, mice (n=49)	Sham mice Vehicle CUR (15 mg/kg) CUR (45 mg/kg) CCI → peripheral injury mice Vehicle CUR (5 mg/kg) CUR (15 mg/kg) CUR (45 mg/kg)	Sham mice Vehicle CUR (15 mg/kg) CUR (45 mg/kg) CCI → peripheral injury mice Vehicle CUR (5 mg/kg) CUR (15 mg/kg) CUR (45 mg/kg)	NI	NI	Vehicle + vehicle 538.5 ± 22.4 Vehicle + DSP-4 524.3 ±30.6 Vehicle + PCPA 192.9±17.7 CUR 45 mg/kg + vehicle 572.0 ± 31.5 CUR 45 mg/kg + DSP -4 549.3 ± 27.4 CUR 45 mg/kg + PCPA 208.1 ± 23.8		Vehicle + vehicle 427.8 ± 27.2 Vehicle + DSP-4 143.7 ± 15.4 Vehicle + PCPA 405.0 ± 12.9 CUR 45 mg/kg + vehicle 440.3 ± 22.1 CUR 45 mg/kg + DSP-4 128.7 ± 26.4 Cur 45 mg/kg + PCPA 418.6 ± 17.8
Moorkoth, et al., 2021	Animal study, mice (n=30), 1, 3 and 6 hours before FST	Group I: Control, vehicle per oral Group II: Escitalopram (10 mg/kg) per oral Group III: Dehydrozingerone 100 mg/kg per oral, 1 hour before FST Group IV: Dehydrozingerone 100 mg/kg per oral, 3 hour before FST Group V: Dehydrozingerone 100 mg/kg per oral, 6 hour before FST	Group I: Control, vehicle per oral Group II: Escitalopram (10 mg/kg) per oral Group III: Dehydrozingerone 100 mg/kg per oral, 1 hour before FST Group IV: Dehydrozingerone 100 mg/kg per oral, 3 hour before FST Group V: Dehydrozingerone 100 mg/kg per oral, 6 hour before FST	Number of crossing line Baseline 1st day Group I: 6.8 ± 1.3 Group II: (F4, 12 = 14.4, p = 0.0023) Group III: (F4,12 = 14.4, p = 0.9187) Group IV: (F4,12 = 14.4, p = 0.8140) Group V: (F4,12 = 14.4, p = 0.1320) 7th day Group I: 4.3 ± 0.5 Group II: (F4,13 = 17.1, p = 0.0004)	Immobility time, sec Baseline 1st day Group I: 3.3 ± 0.1 Group II: (F4,22 = 19.0 with p<0.0001) Group III: (F4,22 = 19.0, p<0.0001) Group IV: (F4,22 = 19.0, p = 0.0124) Group V: (F4,22 = 19.0, p = 0.9998) 7th day Group I: 3.3 ± 0.1 Group II: F4,17 = 18.7, p = 0.0055	7th day Group I: 4103.5 ± 43.43 ng/g of tissue Group II: Increase serotonin (p < 0.0001) Group III: Highest increase serotonin (p<0.0001) Group IV: Lower than group 3 (p <0.0001 for all three neurotransmitters) Group V: Lower than group 4, p <0.0001	7th day Group I: 5310.4 ± 31.48 Group II: Increase dopamine (p < 0.0001) Group III: Highest increase dopamine (p<0.0001), noradrenaline (p <0.0001) and serotonin (p<0.0001) Group IV: Lower than group III (p <0.0001 for all three neurotransmitters) Group V: Lower than group III (p<0.0001 for all three neurotransmitters)	7th day Group I: 817.3 ± 6.58 Group II: Increase noradrenaline (p = 0.0008) Group III: Highest increase noradrenaline (p <0.0001) Group IV: Lower than group III (p <0.0001 for all three neurotransmitters)
CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (<i>Cochlospermum religiosum</i>); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of <i>Aegle marmelos</i> leaves); EEGS (Ethanol extract from <i>G. procumbens</i> stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (<i>Rosmarinus officinalis</i> extract); SNE (<i>Synedrella nodiflora</i> extract)								

CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (*Cochlospermum religiosum*); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of *Aegle marmelos* leaves); EEGS (Ethanol extract from *G. procumbens* stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (*Rosmarinus officinalis* extract); SNE (*Synedrella nodiflora* extract)

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Author, Year	Study design	Intervention	Control	Outcome				
				Open field test (OFT)	Tail suspension test (TST)	Serotonin	Dopamine	Noradrenaline
Moorkoth, et al., 2021	Animal study, mice (n=30), 1, 3, and 6 hours before FST			Group III: (F4,13 = 17.1, p = 0.898) Group IV: (F4,13 = 17.1, p = 0.8613) Group V: (F4,13 = 17.1, p = 0.2929) Number of centre square entry Baseline 1st day Group I: 1.8 ± 0.2 Group II: (F4,18 = 16.6, p <0.0001) Group III: (F4,18 = 16.6, p = 0.0005) Group IV: (F4,18 = 16.6, p = 0.0141) Group V: (F4,18 = 16.6, p = 0.2098) 7th day Group I: 2.5 ± 0.3 Group II: (F4, 12 = 16.4, p = 0.0027) Group III: (F4, 12 = 16.4, p = 0.3723) Group IV: (F4, 12 = 16.4, p = 0.3723) Group V: (F4, 12 = 16.4, p = 0.1675)	Group III: (F4,22 = 19.0, p<0.0001) Group IV: (F4,22 = 19.0, p =0.0124) Group V: (F4,22 =19.0, p= 0.9998) 7th day Group I: 3.3 ± 0.1 Group II: F4,17 = 18.7, p= 0.0055 Group III: F4,17 = 18.7, p= 0.023) Group IV: (3.1 ± 0.2) Group V: (F4,17 = 18.7, p= 0.9529)		Group V: Lower than group 4, (p <0.0001)	Group V: Lower than group 4, p = 0.1214
CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (<i>Cochlospermum religiosum</i>); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of <i>Aegle marmelos</i> leaves); EFCS (Ethanol extract from <i>G. procumbens</i> stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LP5 (Lipopolysaccharide); ME (Microalgae extract); RE (<i>Rosmarinus officinalis</i> extract); SNE (<i>Synedrella nodiflora</i> extract)								

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Sartim et al., 2021	Animal study, mice (n=30)	Group I: Vehicle Group II: CBD 3 mg/kg Group III: CBD 10 mg/kg Group IV: CBD 30 mg/kg Group V: Imipramine 30 mg/kg	Group I: Vehicle Group II: Ketamine 2.5 mg/kg Group III: Ketamine 5 mg/kg Group IV: Ketamine 10 mg/kg Group V: Imipramine 30 mg/kg	None of the tested doses changed the distance travelled performed by animals in the OFT (n = 8–13, F4,55= 0.5656, p>0.05) S-Ketamine None of the tested doses changed the distance travelled performed by animals in the OFT (n=5–7, F4,29 = 1.147,	NI	NI	NI	NI
Zhai et al., 2024	Animal study, female mice, Chronic unpredictable mild stress (CUMS) 4 weeks followed by drugs administration 2 weeks	120 successful model of CUMS mice, divided into 8 groups (n=15) Group I: CUMS model Group II: EEGS low = EEGS-L (5 mg/kg/day) Group III: EEGS moderate = EEGS-M (25 mg/kg/day) Group IV: EEGS high = EEGS-H (125 mg/kg/day) Group V: FLU (10 mg/kg/day) Group VI: CA 10 mg/kg/day Group VII: t-p-CA 10 mg/kg/day 8. CA+t-p-CA 10 mg/kg/day + 10 mg/kg/day CA and t-p-CA mixed in a ratio of 1:1 respectively	Control group, which was not exposed to CUMS= received an equivalent volume of saline	1. Control (99.1 ± 8.7 sec for central time; 41.3 ± 3.4 times for central number) 2. Model group (47.1 ± 1.9 sec, p<0.001) and number (19.8 ± 0.8 times, p<0.001) to cross the central zone were shortened 3. EEGS-M = time in central zone 78.2 ± 5.23 sec, p<0.05) and number in central zone 34.3 ± 2.2 times, p<0.01) 4. CA + t-p-CA = increase the time (77.8 ± 33.7 sec, p<0.1) and number (31.7 ± 1.9 times, p<0.05) to cross the central zone	1. Control (96.9 ± 3.4 sec) 2. Model group (170.1 ± 2.8 sec, p<0.001) were significantly prolonged 3. EEGS-H in the mice, the immobility time 137.1 ± 5.9 sec, p<0.01) was shortened most significantly 4. EEGS-M = immobility time 124.6 ± 4.0 sec, p<0.001) 5. EEGS-M = time in central zone 78.2 ± 5.2 sec, p<0.05) and number in central zone 34.3 ± 2.2 times, p<0.01) 4. CA + t-p-CA = significantly shorten the TST of immobility time (140.7 ± 7.8 sec, p<0.05)	1. Control (20.1 ± 0.9 ng/mL for 5-HT) 2. CUMS (11.0 ± 0.6 ng/mL, p<0.001 for 5-HT) 3. EEGS-M was as effective as FLU compared with the model group 4. Up-regulation impact of 5-HT and NE by the combined effect of chlorogenic acid and trans-p-coumaric acid (p<0.05) was better than any monomer alone		1. Control group (10.15 ± 0.69 ng/mL for NE) 2. CUMS group NE (6.53 ± 0.61 ng/mL, P <0.01) 3. EEGS-M was as effective as FLU compared with the model group. 4. Up-regulation impact of 5-HT and NE by the combined effect of chlorogenic acid and trans-p-coumaric acid (p<0.05) was better than any monomer alone

CBD (Cannabidiol); CCI (Chronic constriction injury); CSR (*Cochlospermum religiosum*); CUMS (Chronic unpredictable mild stress); CUR (Curcumin); EAM (hydroethanolic extract of *Aegle marmelos* leaves); EEGS (Ethanol extract from *G. procumbens* stems); FLU (Fluoxetine); IMP (Imipramine); LBRD (Lily bulb and Rehmannia decoction); LPS (Lipopolysaccharide); ME (Microalgae extract); RE (*Rosmarinus officinalis* extract); SNE (*Synedrella nodiflora* extract)

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Zhai et al., 2024	Animal study, mice (n=36)	All treatments were administered via oral gavage for 14 consecutive days after dissolution in saline						
Bhatt, et al., 2022		Group I: Control, vehicle per oral Group II: FLU 20 mg/kg, ip Group III: IMP 20 mg/kg Group IV: CSR-50 (leaf extract at 50 mg/kg, per oral) Group V: FLU 10 mg/kg + CSR 25 mg/kg	NI	1. Control 136.7 ± 8.069 2. FLU 20 mg/kg, ip 98.2 ± 7.6 3. IMP 107.2 ± 5.9 4. CSR-50 = 103.8 ± 4.7 5. FLU 10 + CSR 25 = 100.0 ± 10.3 6. IMP 10 + CSR 25 = 113.3 ± 3.0	NI	NI	NI	
Chi et al., 2022		Group I: Control, vehicle per oral Group II: Model depression (LPS-induced) 0.3 mg/kg ip Group III: LPS (90 g/kg) + LBRD group gavage 2 h post LPS injection Group IV: LPS + FLU 2 mg/kg 2 hours post LPS injection	NI	NI	Control = 367.0 ± 14.1 Model group = 268.0 ± 7.5 LPS + LBRD = 304.0 ± 15.4	Control= 79.2 ± 2.1 Model= 62.7 ± 1.8 LPS + LBRD = 69.8 ± 2.7	NI	
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Sasaki, et al. 2022	Animal study, mice (n=28), 7 days	Group I: Control, vehicle per oral Group II: Bupropion 20 mg/kg Group III: ME 50 and 100 mg/kg Group IV: ME 100 mg/kg	Group I: Control, vehicle per oral Group II: Bupropion 20 mg/kg Group III: ME 50 and 100 mg/kg Group IV: ME 100 mg/kg	NI	1. Control = 64.6 ± 13.3 sec --> 81.0 ± 7.7 sec --> 103.8 ± 12.8 sec --> 139.4 ± 5.7 sec, --> 140.3 ± 8.0 sec --> 158.0 ± 12.2 sec, and 169.6 ± 9.9 sec 2. Positive control (bupropion 20 mg/kg) = 61.1 ± 8.3 sec --> 58.6 ± 6.0 sec --> 58.2 ± 6.5 sec --> 70.8 ± 2.5 sec --> 65.7 ± 10.0 sec --> 59.0 ± 9.9 sec, and 62.1 ± 10.5 sec 3. ME 50 mg/kg = 65.5 ± 6.1 sec --> 62.0 ± 9.8 sec --> 56.8 ± 13.3 sec --> 68.8 ± 8.3 sec --> 73.5 ± 4.6 sec --> 70.3 ± 11.9 sec and 71.7 ± 8.4 sec 4. ME 100 mg/kg = 65.7 ± 6.6 sec --> 96.0 ± 11.8 sec --> 119.0 ± 11.4 sec --> 123.0 ± 10.2 sec --> 124.0 ± 12.3 sec --> 133.6 ± 11.2 sec --> and 154.4 ± 8.9 sec	NI	NI	NI

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