

Table S3. Quality Assessment Checklist (1 point per criterion for fully satisfied, 0.5 for partially satisfied, 0 for otherwise)

Category 1: Subjects	Score (0/0.5/1)
1. Patients were evaluated prospectively, specific diagnostic criteria were applied, and demographic data were reported.	
2. Healthy subjects were evaluated prospectively, and psychiatric and medical illnesses were excluded.	
3. Important variables (such as age, gender, illness duration, onset time, medication status, comorbidity, and severity of illness) were checked, either by stratification or statistically.	
4. Sample size per group > 10.	
Category 2: Methods for image acquisition and analysis	
5. Magnet strength $\geq 1.5T$.	
6. MRI slice thickness ≤ 2 mm.	
7. The whole-brain analysis was automatically calculated with no prior regional selection.	
8. Coordinates were reported in a standard space.	
9. The imaging technique processing was described clearly enough to be reproducible.	
10. Measurements were described clearly enough to be reproducible.	
Category 3: Results and conclusions	
11. Statistical parameters were provided.	
12. Conclusions were consistent with the results obtained and the limitations were discussed.	
	TOTAL /12

Table S2. Meta-regression analysis of the correlation between GM alterations and clinical variables in migraine patients using the AES-SDM method.

	MNI coordinate			SDM z-score ^a	P value ^b	Number of voxels ^c	Cluster breakdown (number of voxels)
	X	Y	Z				
Duration of migraine							
Bilateral cerebellum and R dorsal medulla	0	-44	-46	1.932	<0.001	1422	R cerebellum, hemispheric lobule IX (335) L cerebellum, hemispheric lobule IX (170) R dorsal medulla (144)
L insula	-36	-12	14	1.398	0.001	465	L insula (255) L rolandic operculum (65)
Frequency of migraine attacks							
L rolandic operculum	-62	0	4	2.987	<0.001	273	L rolandic operculum (150) L superior temporal gyrus (84)
Age							
R dorsal medulla	6	-34	-42	1.377	0.003	77	
L amygdala	-26	-8	-24	-1.664	<0.001	413	L amygdala (96) L parahippocampus (66) L hippocampus (21)
L parahippocampus	-28	-32	-18	-1.336	<0.001	169	L parahippocampus (32)
R parahippocampus	20	2	-24	-1.234	0.001	33	R parahippocampus (28)

^a Peak height threshold: $z > 1$.

^b Voxel probability threshold: $P < 0.005$.

^c Cluster extent threshold: regions with < 20 voxels are not reported in the cluster breakdown.

Abbreviations: GM, gray matter; MNI, Montreal Neurological Institute; L, left; R, right; AES-SDM, anisotropic effect size-signed differential mapping.

Table S3. Subgroup analysis of GM alterations in migraine patients using the AES-SDM method.

	MNI coordinate			SDM z-score ^a	P value ^b	Number of voxels ^c	Cluster breakdown (number of voxels)
	X	Y	Z				
MwoA > HS							
L amygdala	-28	-4	-22	2.007	<0.001	739	L amygdala, BA28, BA34 (141) L temporal pole, superior temporal gyrus, BA28, BA38 (99) L parahippocampus, BA28 (43) L hippocampus, BA28 (20)
R parahippocampus	20	0	-24	1.980	<0.001	577	R parahippocampus, BA28, BA34 (150) R amygdala, BA34 (127)
L parahippocampus	-26	-34	-18	2.026	<0.001	428	L fusiform gyrus, BA30, BA37 (170) L cerebellum, hemispheric lobule IV / V, BA30, BA37 (63) L parahippocampus, BA30 (51) L hippocampus, BA20 (31)
R hippocampus	32	-24	-14	1.392	0.001	90	R hippocampus, BA20 (75)
L lingual gyrus	-18	-96	-20	1.400	0.001	38	L lingual gyrus, BA 18 (23)
MwoA < HS							
Bilateral dorsal medulla and cerebellum	0	-40	-52	-1.504	<0.001	746	R dorsal medulla (138) R cerebellum, hemispheric lobule IX (49) L dorsal medulla (45) L cerebellum, hemispheric lobule IX (34)
R inferior frontal gyrus	42	26	-14	-1.460	<0.001	227	R inferior frontal gyrus, orbital part, BA 38, BA 47 (175) R insula, BA 47 (25)
R middle frontal gyrus	32	44	30	-1.361	0.001	48	R middle frontal gyrus, BA46 (45)
R supplementary motor area	8	22	62	-1.292	0.002	32	R supplementary motor area, BA8 (20)
MwA > HS							
R inferior occipital gyrus	32	-92	-6	1.013	<0.001	364	R inferior occipital gyrus, BA18, BA19 (221) R middle occipital gyrus, BA18 (51)
L middle temporal gyrus	-46	-56	16	1.001	<0.001	244	L middle temporal gyrus, BA21, BA39 (185)
MwA < HS							
L supplementary motor area	2	20	44	-1.022	0.001	715	L superior frontal gyrus, medial, BA8, BA32 (150) L supplementary motor area BA8, BA32 (148)

							R median cingulate/paracingulate gyrus, BA24, BA32 (132)
							L median cingulate/paracingulate gyrus, BA24 (90)
							R superior frontal gyrus, medial, BA8, BA32 (75)
							R supplementary motor area, BA8, BA32 (73)
R cerebellum	2	-64	-56	-1.094	<0.001	79	R cerebellum, hemispheric lobule IX (24)
R superior frontal gyrus, dorsolateral	26	4	56	-1.012	0.002	35	
L temporal pole, superior temporal gyrus	-58	6	-2	-1.003	0.003	26	
EM > HS							
L temporal pole, superior temporal gyrus	-28	8	-24	1.777	<0.001	1125	L temporal pole, superior temporal gyrus, BA28, BA38 (280)
							L amygdala, BA28, BA34 (91)
							L temporal pole, middle temporal gyrus, BA20 (38)
							L insula, BA48 (34)
							L parahippocampus, BA28 (29)
L middle temporal gyrus	-46	-56	16	2.204	<0.001	822	L middle temporal gyrus, BA21, BA22, BA37, BA39 (582)
							L angular gyrus, BA39 (87)
							L superior temporal gyrus, BA22 (25)
R superior frontal gyrus	10	46	38	1.624	<0.001	375	R superior frontal gyrus, BA9, BA10, BA32 (232)
R superior temporal gyrus	44	-34	2	1.349	0.001	316	R superior temporal gyrus, BA21, BA22, BA42 (103)
							R middle temporal gyrus, BA21, BA22 (95)
R amygdala	30	-4	-18	1.272	0.002	160	R amygdala, BA34, BA36 (76)
EM < HS							
Bilateral dorsal medulla and cerebellum	-8	-42	-46	-1.863	<0.001	720	R dorsal medulla (130)
							L dorsal medulla (47)
							L cerebellum, hemispheric lobule IX (21)
L anterior thalamic projection	-4	-16	-6	-1.281	0.002	42	
CM > HS							
R middle frontal gyrus	42	36	22	1.489	<0.001	457	R middle frontal gyrus, BA45, BA46 (307)
							R inferior frontal gyrus, triangular part, BA45 (99)
R caudate nucleus	8	16	10	1.332	<0.001	349	R caudate nucleus, BA25 (148)
							R anterior thalamic projection (87)
L rolandic operculum	-62	0	6	1.231	0.002	53	L rolandic operculum, BA48 (42)
L superior occipital gyrus	-16	-94	32	1.275	0.002	29	

CM < HS							
L inferior temporal gyrus	-34	-10	-44	-1.151	0.004	972	L temporal pole, middle temporal gyrus, BA20, BA36, BA38 (233) L inferior temporal gyrus, BA20, BA36 (197) L fusiform gyrus, BA20, BA36 (101) L temporal pole, superior temporal gyrus, BA38 (89)
R inferior frontal gyrus, orbital part	40	24	-12	-1.156	0.003	72	R inferior frontal gyrus, orbital part, BA47 (27)
R inferior temporal gyrus	40	2	-44	-1.157	0.003	41	R inferior temporal gyrus, BA20 (22)
L superior frontal gyrus, dorsolateral	-28	66	6	-1.240	0.002	30	
R superior frontal gyrus, orbital part	16	70	-8	-1.220	0.003	26	
L precentral gyrus	-34	-20	64	-1.155	0.003	26	L precentral gyrus, BA6 (25)
VM > HS							
L superior occipital gyrus	-20	-84	26	1.021	<0.001	1443	L superior occipital gyrus, BA18, BA19 (388) L middle occipital gyrus, BA18, BA19 (340) L cuneus cortex, BA18, BA19 (220)
VM < HS							
L superior temporal gyrus	-52	-6	2	-2.594	<0.001	3396	L superior temporal gyrus, BA22, BA41, BA42, BA48 (789) L insula, BA48 (616) L rolandic operculum, BA48 (489) L supramarginal gyrus, BA2, BA48 (272) L lenticular nucleus, putamen, BA48 (173) L heschl gyrus, BA48 (168) L postcentral gyrus, BA48 (83)
R superior temporal gyrus	48	-24	12	-2.514	<0.001	2683	R insula, BA48 (672) R rolandic operculum, BA48 (598) R superior temporal gyrus, BA22, BA48 (306) R lenticular nucleus, putamen, BA48 (214) R heschl gyrus, BA48 (171)
R middle frontal gyrus	42	48	16	-2.536	<0.001	566	R middle frontal gyrus, BA10, BA45, BA46 (441) R superior frontal gyrus, dorsolateral, BA10 (49)
Migraine patients in interictal period > HS							
R amygdala	24	0	-20	1.677	<0.001	1055	R parahippocampus, BA28, BA34, BA36 (229) R amygdala, BA34, BA36 (158)

L amygdala	-26	-8	-22	1.387	<0.001	413	R temporal pole, superior temporal gyrus, BA 38 (42) L amygdala, BA 28, BA 34 (107) L hippocampus, BA 28 (20)
L lingual gyrus	-18	-96	-20	1.184	<0.001	136	L lingual gyrus, BA18 (73) L inferior occipital gyrus, BA18 (35)
L hippocampus	-28	-22	-14	1.143	<0.001	133	L hippocampus, BA20 (65)
R cerebellum, hemispheric lobule III	10	-34	-12	1.025	0.001	85	R cerebellum, hemispheric lobule III, BA30 (28)
R lingual gyrus	22	-94	-20	1.099	<0.001	79	R lingual gyrus, BA18 (52)
R hippocampus	32	-22	-16	1.012	0.001	58	R hippocampus, BA20 (44)
Migraine patients in interictal period < HS							
L rolandic operculum	-44	-6	6	-1.962	<0.001	1120	L insula, BA48 (432) L rolandic operculum, BA48 (261) L heschl gyrus, BA48 (117) L superior temporal gyrus, BA48(114)
R middle frontal gyrus	36	46	26	-1.792	<0.001	291	R middle frontal gyrus, BA46 (263)
R dorsal medulla	0	-42	-50	-1.502	0.002	247	R dorsal medulla (23) R cerebellum, hemispheric lobule IX (22)
L cerebellum, hemispheric lobule VI	-28	-66	-22	-1.494	0.002	195	L cerebellum, hemispheric lobule VI, BA19, BA37 (182)
R inferior parietal (excluding supramarginal and angular) gyrus	46	-46	44	-1.472	0.002	80	R inferior parietal (excluding supramarginal and angular) gyrus, BA40 (64)
R rolandic operculum	54	8	2	-1.456	0.002	62	R rolandic operculum, BA48 (42)

^a Peak height threshold: $z > 1$.

^b Voxel probability threshold: $P < 0.005$.

^c Cluster extent threshold: regions with < 20 voxels are not reported in the cluster breakdown.

Abbreviations: GM, gray matter; L, left; R, right; MwA, migraine with aura; MwoA, migraine without aura; HS, healthy subjects; MNI, Montreal Neurological Institute; AES-SDM, anisotropic effect size-signed differential mapping; EM, episodic migraine; CM, chronic migraine; VM, vestibular migraine.

Table S4. Subgroup analysis of GM alterations in migraine patients using the ALE method.

Cluster no.	Cluster size (mm ³)	Weighted center (x, y, z)			x	y	z	ALE value (×10 ⁻³)	Label (Nearest Gray Matter within 5mm)
MwoA > HS									
1	720	-21.5	-31.4	-13.8	-22	-32	-14	18.222	L parahippocampus, BA35
EM > HS									
1	600	-19.6	-30.9	-11.1	-20	-32	-12	16.102	L parahippocampus, BA28
VM > HS									
1	1728				-19	-84	28	9.246	L occipital gyrus, BA18
Migraine patients in interictal periods > HS									
1	760	-21.6	-31.3	-13.8	-22	-32	-14	18.222	L parahippocampus, BA35

Abbreviations: GM, gray matter; ALE, activation likelihood estimation; L, left; MwoA, migraine without aura; EM, episodic migraine; VM, vestibular migraine; HS, healthy subjects; R, right.

Table S5. Heterogeneity of altered GM regions between migraine patients and HS in VBM studies using the AES-SDM method.

Regions	MNI coordinate			SDM z-score ^(a)	p-value ^(b)	Number of voxels ^(c)
	x	y	z			
L insula	-38	-10	12	2.274	<0.001	822
R middle frontal gyrus	46	40	30	2.815	<0.001	117

^a Peak height threshold: $z > 1$;

^b Voxel probability threshold: $p < 0.005$;

^c Cluster threshold: Regions with less than 10 voxels are not reported.

Abbreviations: GM, gray matter; HS, healthy subjects; L, left; R, right; MNI, Montreal Neurological Institute; AES-SDM, anisotropic effect size-signed differential mapping; VBM, voxel-based morphometry.

Table S6. Sensitivity analysis of VBM meta-analysis using the AES-SDM method.

[illegible]

[illegible]

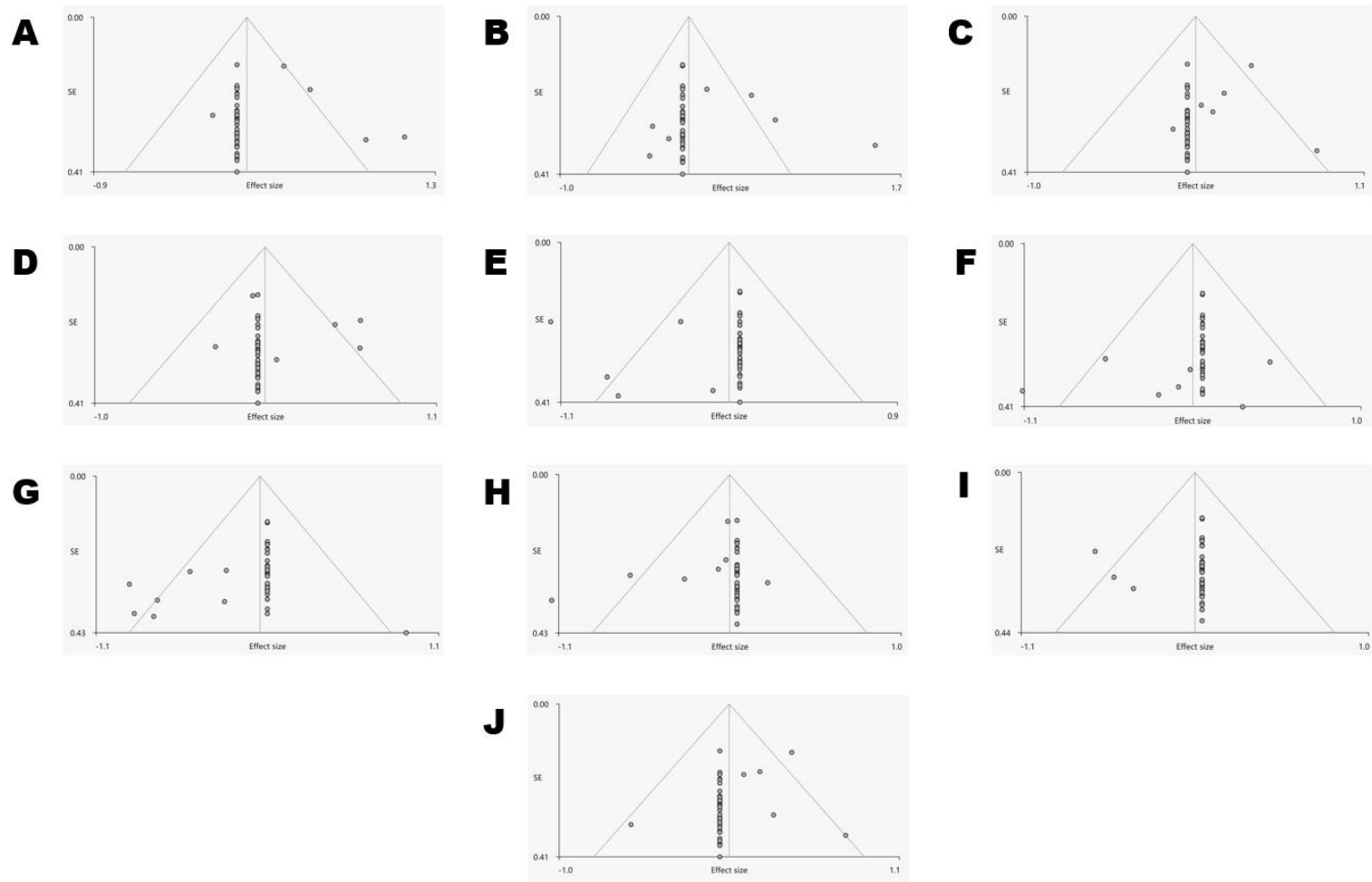
Bonanno L 2020[28] (MwoA)	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes
Bonanno L 2020[28] (MwA)	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Li ZJ 2020[29]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	Yes	Yes	Yes	Yes
Liu HY 2020[30]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No
Zhe X 2021[31]	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes	Yes	No	Yes	Yes	Yes
Yu Y 2021[32] (Episodic)	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No
Yu Y 2021[32] (Chronic)	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Chou KH 2021[33]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Masson R 2021[34]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Zhao L 2011[35]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Only left	Yes	Yes
Chen XY 2014[36] (Episodic)	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No
Chen XY 2014[36] (Chronic)	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Yao Q 2017[37]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Zhe X 2018[38]	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes	Yes	Only right	No	No	No
Li MQ 2020[39]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Wang JH 2021[40]	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Only left	Yes	Yes	Yes
	47/47	46/47	46/47	47/47	45/47	44/47	45/47	46/47	46/47	41/47	41/47	40/47	40/47

Abbreviations: VBM, voxel-based morphometry; AES-SDM, anisotropic effect size-signed differential mapping; L, left; R, right; GM, gray

matter.

Figure S1. Results of funnel plot analysis for the meta-analysis of all included VBM studies using the AES-SDM method.

The Egger's test and funnel plots revealed no significant publication bias in the (A) right amygdala ($Z=-0.31$, $t=-0.78$, $df=45$, $P=0.441$), (B) right superior frontal gyrus ($Z=0.10$, $t=0.22$, $df=45$, $P=0.826$), (C) left hippocampus ($Z=-0.53$, $t=-1.98$, $df=45$, $P=0.054$), (D) left middle temporal gyrus ($Z=-0.19$, $t=-0.57$, $df=45$, $P=0.570$), (E) bilateral cerebellum and right dorsal medulla ($Z=-0.06$, $t=-0.13$, $df=45$, $P=0.894$), (F) left insula ($Z=-1.06$, $t=-1.79$, $df=45$, $P=0.080$), (G) right rolandic operculum ($Z=-0.58$, $t=-1.46$, $df=45$, $P=0.151$), (H) right middle frontal gyrus ($Z=-0.64$, $t=-1.63$, $df=45$, $P=0.110$) and (I) right inferior parietal gyrus ($Z=-0.53$, $t=-1.38$, $df=45$, $P=0.175$). Significant publication bias was reported in the (J) left amygdala ($Z=-0.76$, $t=-2.44$, $df=45$, $P=0.019$) by the Egger's test and funnel plots.



Abbreviations: VBM, voxel-based morphometry; SE, standard error; AES-SDM, anisotropic effect size-signed differential mapping.

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