Primary/secondary outcome	Studies	Risk of	Inconsistency	Indirectness	Imprecision	Publication	Large	Overall quality of
	(N)	bias				bias	effect	evidence ^a
Prolactin level in all patients (ng/mL)	5 (385)	Serious ^b	Serious ^c	No	No	Serious ^d	Large ^e	+/+/-/-/; Low
Prolactin level in females (ng/mL)	2 (186)	Serious ^b	No	No	No	Serious ^d	Large ^e	+/+/+/-/; Moderate
Prolactin level in males (ng/mL)	2 (127)	Serious ^b	Serious ^c	No	No	Serious ^d	Large ^e	+/+/-/-/; Low
Total psychopathology	5 (385)	Serious ^b	No	No	No	Serious ^d	No	+/+/-/-/; Low
PANSS positive symptoms	3 (213)	Serious ^b	Serious ^c	No	No	Serious ^d	No	+/-/-/; Very low
PANSS negative symptoms	3 (213)	Serious ^b	No	No	No	Serious ^d	No	+/+/-/-/; Low
All caused discontinuation	2 (180)	No	No	No	No	Serious ^d	No	+/+/+/; Moderate
Any extrapyramidal symptoms	2 (172)	No	No	No	No	Serious ^d	No	+/+/+/; Moderate

Supplemental Table 1. Adjunctive aripiprazole for antipsychotic-related hyperprolactinemia: GRADE analyses

GRADE = grading of recommendations assessment, development, and evaluation; PANSS = the Positive and Negative Syndrome Scale.

^aGRADE Working Group grades of evidence: High quality=further research is very unlikely to change our confidence in the estimate of effect. Moderate quality=further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate. Low quality=further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate. Very low quality=we are very uncertain about the estimate.

^bMore than 50% studies reported as having a serious bias used an open-label method, only mentioned random allocation without describing the method and withdrawal from the study.

^cAll studies reported as having a serious inconsistency had $I^2 > 50\%$.

^d For continuous outcomes, N<400; For dichotomous outcomes, N< 300 following the methodology of a previous study^[1].

^eStudies with large effects provided increased quality of evidence. Large effects=standard mean differences<-0.8.

[1] Bo Q J , Wang Z M , Li X B , et al. Adjunctive metformin for antipsychotic-induced hyperprolactinemia: A systematic review[J]. Psychiatry Res, 2016, 237:257-263.

	Random sequence	generation (selection bias)	Allocation	concealment	(selection bias)	Blinding of	participants and	personnel	assessment	(Symptom reduction,	resnonse)	incomplete outcome data addressed	(attrition bias)	Selective reporting	(reporting bias)	Other sources of blas
Chen et al., 2009		+		?			+			?		+			÷	?
Chen et al., 2012		?		-			-			-		?			÷	?
Ren and Hu 2011		+		?			+			?		+			-	?
Sha et al., 2017		+		-			-			-		+			+	?
Zhou et al., 2014		?		-			-			-		+			+	?

Supplemental Figure 1: Risk of bias

+ : Low risk of bias, - : High risk of bias, ? : Unclear risk of bias, nd : not determined

Supplemental Figure 2: forest plot for all caused discontinuation and adverse drug reactions

		Risk Ratio	Risk Ratio
Secondary outcomes	Weight	M-H, Random, 95% CI	M-H, Random, 95% CI
2.1 Discontinuation due to	o any reason		
Chen et al., 2009	73.3%	0.67 [0.26, 1.70]	
Zhou et al., 2014	26.7%	1.00 [0.21, 4.72]	
Subtotal (95% CI)	100.0%	0.74 [0.33, 1.65]	•
Heterogeneity: Tau ² = 0.00;	Chi ² = 0.19, df	f = 1 (P = 0.66); l² = 0%	
Test for overall effect: Z = 0	.73 (P = 0.47)		
2.2 Any extrapyramidal sy	mptoms		
Ren and Hu 2011	12.0%	0.20 [0.01, 4.03]	
Zhou et al., 2014	88.0%	1.09 [0.71, 1.67]	
Subtotal (95% CI)	100.0%	0.89 [0.29, 2.70]	-
Heterogeneity: Tau ² = 0.32;	Chi ² = 1.26, df	f = 1 (P = 0.26); l ² = 21%	
Test for overall effect: Z = 0	.21 (P = 0.84)		
		0.01	0.1 1 10 10
		Favours [e	experimental] Favours [control

3

Supplementary material

4