評価対象論文リスト（要因：過剰飲酒（多量飲酒）、アウトカム：がん）

評価判定日：2024/1/25

①既存の系統的レビュー・メタ解析・統合解析

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■コホート研究

Reference			Study subjects						Category	Number among cases	Relative risk (95%CI or p)	P for trend	Confounding variables considered	**Magnitude of association**
Author	Title	Year	Study period	Number of subjects	Source of subjects	Event followed	Number of incident cases or deaths	Participant's race						
Manami Inoue et al	Impact of alcohol drinking on total cancer risk: data from a large-scale population-based cohort study in Japan	2005	1990-2001	73,281 men; 35,007 women; 38,274	JPHC Study; Population-based prospective study	**Cancer incidence** **Cancer deaths**	3403 1208	Japanese	**Men;** Nondrinkers Occasional drinkers Weekly etanol intake (g per w) 1 -149 150-299 300-449 ≧450 **Women;** Nondrinkers Occasional drinkers Weekly etanol intake (g per w) 1 -149 150-299 300-449 ≧450	**Total cancer incidence;** 360 138 353 359 339 355 **Total cancer death;** 161 59 138 119 133 148 **Total cancer incidence;** 1170 178 118 20 6 7 **Total cancer death;** 368 43 28 6 3 2	1.10 (0.90 –1.34) 1.00 (reference) 1.18 (0.96 – 1.44) 1.17 (0.96 –1.44) **1.43 (1.17 – 1.75)** **1.61 (1.32 – 1.97)** 1.10 (0.81 –1.49) 1.00 (reference) 1.06 (0.77 – 1.44) 0.92 (0.67 –1.26) 1.33 (0.97 – 1.83) **1.58 (1.16 – 2.15)** 0.94 (0.80 –1.11) 1.00 (reference) 0.80 (0.63 – 1.01) 0.68 (0.42 –1.11) 0.73 (0.32 – 1.66) 0.68 (0.32 – 1.46) 1.08 (0.79 –1.49) 1.00 (reference) 0.79 (0.49 – 1.27) 0.54 (0.19 –1.52) 1.27 (0.39 – 4.15) 0.68 (0.16 – 2.86)	**< 0.001** **< 0.001** 0.659 0.896	age at baseline (continuous), study area (9 PHC area), pack-years of smoking (0, 1 – 19, 20 – 29, 30 – 39, X40), green vegetable intake (p3 – 4 times per week, almost everyday), and leisure-time physical activity (p1 – 3 times per month, X1 – 2 times per week).	↑ ↑ ↑ ↑ - -
Ono, A. et al.	Impact of alcohol drinking on cancer risk with consideration of flushing response: The Japan Public Health Center-based Prospective Study Cohort (JPHC study)	2020	1993-2013	56,890 men; 26,305 women; 30,585	JPHC Study; Population-based prospective study	**Cancer incidence** **Alcohol-related cancer incidence**	men; 8,486 women; 5,134 total; 3,352 colon cancer; 4,386 gastric cancer; 1,524 breast cancer; 1,516 500	Japanese	**All cancers incidence** Men; Flusher Past drinker Non-drinker (< 1 day/month) Occasional drinker (1-3 days/month) Regular drinker (g/week) 0-149 g 150-299 g ≥ 150 g for women 300-449 g ≥ 450g Non flusher Past drinker Non-drinker (< 1 day/month) Occasional drinker (1-3 days/month) Regular drinker (g/week) 0-149 g	8,486	1.23(1.00–1.51) 1.04 (0.92–1.18) 0.98 (0.84–1.16) 1.11 (0.98–1.25) **1.18 (1.05–1.34)** **1.37 (1.19–1.57)** 1.10 (0.94–1.30) 0.85 (0.67–1.07) 1.00 (reference) 1.14 (0.93–1.40) 1.11 (0.99–1.26)		age, public health center area, smoking status (never, former,<20 pack-years, 20–29 pack-years, 30–39 pack-years, ≥ 40 pack-years), flushing response, intake of green vegetables, leisure-time sports or physical exercise, past	↑

	150-299 g ≥ 150 g for women	1.03 (0.92–1.16)	history of diabetes mellitus, past history of hypertension, pasthistory of hyperlipidemia and family history of cancers	
	300-449 g	1.09 (0.96–1.24)		
	≥ 450g	**1.22 (1.07–1.40)**		↑
Women;				
Flusher				
	Past drinker	1.36 (0.81–2.29)		
	Non-drinker (< 1 day/month)	1.07 (0.99–1.17)		
	Occasional drinker (1-3 days/month)	1.00 (0.82–1.21)		-
	Regular drinker (g/week)			
	0-149 g	0.93 (0.78–1.11)		
	150-299 g ≥ 150 g for women	1.16 (0.80–1.69)		
Non flusher				
	Past drinker	1.04 (0.66–1.63)		
	Non-drinker (< 1 day/month)	1.00 (reference)		
	Occasional drinker (1-3 days/month)	1.12 (0.96–1.30)		-
	Regular drinker (g/week)			
	0-149 g	1.00 (0.87–1.14)		
	150-299 g ≥ 150 g for women	1.14 (0.88–1.46)		
Alcohol-related cancer incidence		4,386		
Men;				
Flusher				
	Past drinker	1.37 (1.03–1.84)		
	Non-drinker (< 1 day/month)	0.96 (0.79–1.15)		
	Occasional drinker (1-3 days/month)	1.08 (0.86–1.36)		
	Regular drinker (g/week)			
	0-149 g	1.13 (0.96–1.34)		
	150-299 g ≥ 150 g for women	**1.47 (1.25–1.73)**		↑
	300-449 g	**1.69 (1.40–2.04)**		↑ ↑
	≥ 450g	**1.32 (1.06–1.65)**		↑
Non flusher				
	Past drinker	0.94 (0.68–1.31)		
	Non-drinker (< 1 day/month)	1.00 (reference)		
	Occasional drinker (1-3 days/month)	1.19 (0.89–1.60)		
	Regular drinker (g/week)			
	0-149 g	**1.31 (1.11–1.55)**		↑
	150-299 g ≥ 150 g for women	**1.24 (1.06–1.46)**		↑
	300-449 g	**1.34 (1.12–1.59)**		↑
	≥ 450g	**1.59 (1.32–1.91)**		↑ ↑
Women;				
Flusher				
	Past drinker	**2.34 (1.35–4.06)**		↑ ↑ ↑
	Non-drinker (< 1 day/month)	**1.15 (1.02–1.29)**		↑
	Occasional drinker (1-3 days/month)	1.16 (0.90–1.49)		
	Regular drinker (g/week)			
	0-149 g	1.04 (0.83–1.31)		
	150-299 g ≥ 150 g for women	1.53 (0.98–2.39)		
Non flusher				
	Past drinker	1.44 (0.84–2.45)		
	Non-drinker (< 1 day/month)	1.00 (reference)		
	Occasional drinker (1-3 days/month)	1.15 (0.94–1.42)		
	Regular drinker (g/week)			
	0-149 g	1.08 (0.91–1.29)		
	150-299 g ≥ 150 g for women	**1.49 (1.09–2.04)**		↑

■統合解析

Reference			Study subjects						Category	Number among cases	Relative risk (95%CI or p)	P for trend	Confounding variables considered	**Magnitude of association**
Author	Title	Year	Study period	Number of subjects	Source of subjects	Event followed	Number of incident cases or deaths	Participant's race						
Mizoue T et al	Alcohol Drinking and Colorectal Cancer in Japanese: A Pooled Analysis of Results from Five Cohort Studies	2008	1988-2004	209,763 men; 98,265 women; 111,498	Japan cohort consortium; 5 cohorts;	**Cororectal cancer incidence**	men; 1724 women; 1078	Japanese	**Men;**				area (Japan Public Health Center-based Prospective Study (I and II) and Japan Collaborative Cohort Study), age (years; continuous), smoking (never smoker, past smoker, current smoker of 1–19 cigarettes/day, or current smoker of ≧20 cigarettes/day), body mass index (weight (kg)/height (m)2; <22, 22–24.9, 25–27.9, or ≧28), and intakes of energy (continuous), red meat (quartiles), calcium (quartiles), fiber (quartiles), and folate (quartiles)	↑ ↑ ↑
									Nondrinkers	311	1.00 (reference)			
									Occasional drinkers(<once/week)	87	1.00 (0.79, 1.28)			
									Current drinkers(≧once/week)					
									0.1–22.9 g/day	295	1.22 (0.92, 1.61)			
									23–45.9 g/day	363	**1.42 (1.21, 1.66)**			
									46–68.9 g/day	374	**1.95 (1.53, 2.49)**			
									69–91.9 g/day	182	**2.15 (1.74, 2.64)**			
									≧92 g/day	112	**2.96 (2.27, 3.86)**			
									Alcohol intake as a continuous variable (per 15 g/day)	-	**1.11 (1.09, 1.14)**	**<0.001**		
									Women;					
									Nondrinkers	839	1.00 (reference)			
									Occasional drinkers(<once/week)	100	0.96 (0.70, 1.32)			
									Current drinkers(≧once/week)					
									0.1-22.9 g/day	97	0.93 (0.70, 1.23)			
									≧23 g/day	42	**1.57 (1.11, 2.21)**			
									Alcohol intake as a continuous variable (per 15 g/day)	-	**1.13 (1.06, 1.20)**	**<0.001**		
Manami Inoue et al	Impact of alcohol intake on total mortality and mortality from major causes in Japan: a pooled analysis of six large-scale cohort studies	2012	1988-2006	309,082 men; 144,012 women; 165,070	Japan cohort consortium; 6 cohorts;	**Total cancer death**	men; 8,584 women; 4,690	Japanese	**Men;**				age, area, smoking, body mass index, history of hypertension, history of diabetes, leisure-time sports or physical exercise	↑
									Non-drinkders	2648	1.00 (Reference)			
									Occasional drinkers(<once/week)	446	**0.75 (0.68 to 0.84)**			
									Current drinkers(≧once/week)					
									<23 g/day	1712	**0.86 (0.79 to 0.92)**			
									23 to 46 g/day	1911	**0.91 (0.84 to 0.98)**			
									46 to < 69 g/day	1099	0.95 (0.88 to 1.03)			
									69 to < 92 g/day	511	1.12 (0.94 to 1.33)			
									≧92 g/day	257	**1.24 (1.08 to 1.42)**			
									per 15 g increase		1.01 (0.999 to 1.01)	0.084		
									Women;					
									Non-drinkders	3787	1.00 (Reference)			
									Occasional drinkers(<once/week)	376	0.94 (0.81 to 1.09)			
									Current drinkers(≧once/week)					
									<23 g/day	402	0.88 (0.73 to 1.05)			-
									23 to 46 g/day	93	1.02 (0.81 to 1.29)			
									≧46 g/day	32	1.05 (0.73 to 1.50)			
									per 15 g increase		0.995 (0.98 to 1.01)	0.545		
Shimazu T; Sasazuki S; Wakai K; Tamakoshi A; Tsuji I; S z ugawara Y; Matsuo K; Nagata C; Mizoue T; Tanaka K; Inoue M; Tsugane	Alcohol drinking and primary liver cancer: a pooled analysis of four Japanese cohort studies.	2012	1988-2004	174,719 men; 89,863 women; 84,856	Japan cohort consortium; 4 cohorts;	**Liver cancer incidence**	men; 605 women; 199	Japanese	**Men;**				geopraphic area, age, history of diabetes mellitus, smoking, coffee intake	↑ ↑
									Nondrinkers	228	**1.70 (1.15–2.53)**			
									Occasional drinkers(<once/week)	29	1.00 (Reference)			
									Current drinkers, alcohol intake(g/day)					
									0.1-22.9	82	0.88 (0.57–1.36)			
									23.0-45.9	107	1.06 (0.70–1.62)			
									46.0-68.9	76	1.07 (0.69–1.66)			
									69.0-91.9	54	**1.76 (1.08–2.87)**			
									≧92.0 g/day	29	1.66 (0.98–2.82)			
									Alcohol intake as a continuous variable (per 10 g/day)		**1.02 (1.004-1.04)**	0.015		
									Women;					
									Nondrinkers	175	1.50 (0.69–3.25)			
									Occasional drinkers(<once/week)	7	1.00 (Reference)			
									Current drinkers, alcohol intake(g/day)					

										0.1-22.9	8	0.86 (0.26–2.88)			
										≧23.0 g/day	9	**3.60 (1.22–10.66)**			↑ ↑ ↑
										Alcohol intake as a continuous variable (per 10 g/day)		1.11 (0.96–1.29)	0.165		
Oze I; Charvat H; Matsuo K; Ito H; Tamakoshi A; Nagata C; Wada K; Sugawara Y; Sawada N; Yamaji T; Naito M; Tanaka K; Shimazu T; Mizoue T; Tsugane S; Inoue M;	Revisit of an unanswered question by pooled analysis of eight cohort studies in Japan: Does cigarette smoking and alcohol drinking have interaction for the risk of esophageal cancer?	2019	1984-2009	men; 162,826	Japan cohort consortium; **esophageal cancer inciden** 8 cohorts;	men;	954	Japanese	**Men;**				N/A;		
									Cigarette smoking status / Alcohol drinking status						
									Never / Never		1.00 (Reference)		age, area, body mass index, vegetables and fruit intake. Esophageal cancer arising within 2 years of the start of follow-up was excluded.		
									Ever / Never		**2.92 (1.59-5.36)**				
									Never / Ever		**2.73 (1.78-4.18)**				
									Ever / Ever		**8.86 (4.82-16.30)**				↑ ↑ ↑
									Pack-years / Amount of alcohol drinking (g/day)						
									0 / <23		1.00 (Reference)				
									0 / ≥23, <46		**3.90 (1.84-8.27)**				
									0 /≥23, <46		**5.62 (2.94-10.76)**				
									≦40 / <23		**2.77 (1.53-5.01)**				
									≦40 / ≥23, <46		**8.08 (4.61-14.15)**				
									≦40 / ≧46		**10.53 (6.04-18.38)**				
									>40 / <23		**5.16 (3.14-8.50)**				
									>40 / ≥23, <46		**13.65 (6.66-27.99)**				
									>40 / ≧46		**17.15 (11.04-27.77)**				↑ ↑ ↑
Masaoka, H. et al.	Alcohol Drinking and Bladder Cancer Risk From a Pooled Analysis of Ten Cohort Studies in Japan	2020		340497	Japan cohort consortium; **bladder cancer incidence** 10 cohorts;	men;	936	Japanese	**Men;**				N/A;		
									Non-drinker	247	1.00 (Reference)				
									Occasional drinker (<1/week)	80	0.89 (0.68–1.17)				
									Regular drinker (≥1 /week)						
									0.1-22.9	156	0.95 (0.77–1.17)				
									23.0-45.9	199	1.07 (0.88–1.30)		age (continuous), public health center area, smoking status		-
									46.0-68.9	160	1.13 (0.91–1.39)				
									≥69.0	94	1.02 (0.79–1.33)				
									Unit HR (per 10 g of ethanol among current drinkers)		1.01 (0.99–1.03)				
									Women;				N/A;		
									Non-drinker	246	1.00 (Reference)				
									Occasional drinker (<1/week)	41	1.31 (0.91–1.88)				-
									Regular drinker (≥1 /week)	38	0.96 (0.66–1.40)				
Iwase, M. et al.	Alcohol consumption and breast cancer risk in Japan: A pooled analysis of eight population-based cohort studies	2021	1984-2013	158164	Japan cohort consortium; **brest cancer incidence** 8 cohorts;		2208	Japanse	**Women;**						
									Premenopausal at baseline						
									Drinking frequency						
									Non	597	1.00 (Reference)				
									Occasional	121	1.01 (0.71-1.43)				
									Moderate (1-4 days/week)	130	1.19 (0.93-1.52)				
									Regular (≥5 days/week)	62	**1.37 (1.04-1.81)**				↑
									Trend (per category)		**1.09 (1.02-1.18)**	**0.017**			
									Amount analysis						
									Drinking amount (g/d)						
									0	635	1.00 (Reference)				
									0 to <11.5	121	1.21 (0.96-1.53)				
									11.5 to <23	27	1.28 (0.85-1.92)		age, area, smoking status, body mass index, age at menarche,		
									≥23	41	**1.74 (1.25-2.43)**				↑ ↑
									trend (per 10 g)		**1.05 (1.00-1.10)**	**0.041**			
									Postmenopausal at baseline						

									Drinking frequency				party		
									Non	972	1.00 (Reference)				
									Occasional	114	1.16 (0.95-1.41)				
									Moderate (1-4 days/week)	100	0.94 (0.70-1.27)				-
									Regular (≥5 days/week)	60	1.11 (0.80-1.55)				
									Trend (per category)		1.01 (0.92-1.11)	0.882			
									Amount analysis						
									Drinking amount (g/d)						
									0	945	1.00 (Reference)				
									0 to <11.5	99	1.05 (0.82-1.33)				
									11.5 to <23	21	0.96 (0.62-1.49)				-
									≥23	24	1.18 (0.66-2.11)				
									trend (per 10 g)		0.99 (0.93-1.06)	0.824			
Tamura, T. et al.	Alcohol intake and stomach cancer risk in Japan: A pooled analysis of six cohort studies	2022	1988-2013	256,478 men; 119,951 women; 136,527	Japan cohort consortium; 6 cohorts;	**stomach cancer incidence**	men; 6051 women; 2535	8586	Japanese	**Men;**					
										Site of cancer: total;					
										Nondrinkers	1465	1.00 (Reference)			
										Occasional drinkers (<once/wk)	343	1.00 (0.85-1.18)		age, area, smoking status, medical history of diabetes mellitus. total energy intake (quartiles), vegetable intake (quartiles), fruit intake (quartiles), salt intake (quartiles), and green tea consumption	
										Regular drinkers (≥once/wk)					
										<23 g/d	1001	0.98 (0.89-1.07)			
										23 to <46 g/d	1264	1.07 (0.99-1.16)			
										46 to <69 g/d	1206	**1.15 (1.05-1.25)**			↑
										69 to <92 g/d	531	**1.19 (1.05-1.36)**			↑
										≥92 g/d	241	**1.23 (1.05-1.43)**			↑
										Alcohol intake as a continuous variable (per 10 g/d)		**1.019 (1.007-1.032)**	<.002		
										Women;					
										Site of cancer: total;					
										Nondrinkers	2051	1.00 (Reference)			
										Occasional drinkers (<once/wk)	187	1.04 (0.88-1.22)			
										Regular drinkers (≥once/wk)					
										<23 g/d	210	0.93 (0.75-1.15)			
										≥23 g/d	87	**1.38 (1.10-1.74)**			↑
										Alcohol intake as a continuous variable (per 10 g/d)		1.031 (0.992-1.071)	0.12		

■症例対照研究

Reference			Study subjects					Category	Number among cases	Relative risk (95%CI or p)	P for trend	Confounding variables considered	**Magnitude of association**
Author	Title	Year	Study period	Type and source	Definition	Number of cases	Number of controls						
Matsuo K, Hamajima N, Shinoda M, Hatooka S, Inoue M, Takezaki T, Tajima K	Gene-environment interaction between an aldehyde dehydrogenase-2 (ALDH2) polymorphism and alcohol consumption for the risk of esophageal cancer	2001	Recruited; 1999-2000 Cases were first 1984-2000	Aichi Cancer Center	Cases; histologically confirmed esophageal cancer	male; 86	male; 118	**ALDH2 genotype, alcohol**					
								Glu/Glu			N/A		
					Controls; non-cancer outp	female; 16	female; 123	Other		1.00 (Reference)		age, sex	
								Heavy drinker		**7.84 (2.77-22.2)**			↑ ↑ ↑
								Glu/Lys & Lys/Lys					
								Other		1.00 (Reference)			
								Heavy drinker		**49.6 (14.5-169.4)**			↑ ↑ ↑
								All					
								Other		1.00 (Reference)			
								Heavy drinker		**13.5 (6.94-26.5)**			↑ ↑ ↑