Women's Mental Health in Midlife and Beyond: Research Gaps

Lauren M. Osborne, MD Assistant Director, Women's Mood Disorders Center Johns Hopkins University School of Medicine



- None
- Research support from NIMH, ABPN, Doris Duke Foundation, Brain & Behavior Foundation

Menopause & Beyond

- 4-6x risk of depressive sx for those with a history, 2-3x risk for those without
- Cognitive effects who is at risk, how to treat?
- Link with early life adversity
- Connection between mental health and physical symptoms – direction?
- Stress eating a significant mediator between depressive symptoms and weight in postmenopausal women
- Research has focused on estrogen gaps in other hormones, other biological systems; different populations





Salk et al., Psychol Bull 2017 14(8):783-8333; Lee et al., Social Science & Medicine 105 (2014) 122e130

Early Life Adversity

- Childhood sexual abuse: 1 in 5 girls, 1 in 20 boys
- All abuse associated with midlife metabolic syndrome; sexual for women only
- Childhood abuse associated with greater all-cause mortality at midlife and beyond for women only, even when sexual abuse excluded
- SES and life-course mediators alter relationship for men but not women
- Treatment? Resilience? Biological mediators? Other populations?



A, Plot of cumulative mortality hazard by years since study entry for women with and without reported severe childhood physical abuse. B, Cumulative mortality hazard for women with all types of reported childhood abuse (emotional, moderate physical, and severe physical), some types of reported abuse, and no reported abuse. Analyses control for age, race/ethnicity, education, history of heart disease, history of cancer, alcohol use, and smoking.

Chen et al., JAMA Psychiatry. 2016;73(9):920-927; Lee & Ryff, Social Science & Medicine 240 (2019) 112566; Lee et al., Social Science & Medicine 105 (2014) 122e130

Figure. Cumulative Hazard Plots by Abuse Group

Anxiety

- Aging Anxiety (health declines, loss of fertility, appearance)
- Late-life onset of generalized anxiety (25%)
- Only 1/3 of elderly women with anxiety are treated
- What are the biological pathways?
- Alternative treatments?



Estrogen and fear conditioning



Li and Graham, Lancet 2017



Figure 3: Hypothesised model of the influence of sex hormones on biological, behavioural, and cognitive pathways that promote or reduce anxiety disorder vulnerability and maintenance in women

The Connection between Mind and Body

https://nutrition.org/inflammation-what-is-it-and-how-can-my-diet-and-behavior-affect-it/mind-body-connection-1/

Overweight & Obesity

- Overall rates higher in men than in women, but obesity higher in women
- Weight gain in both associated with job demands, bills, but perceived constraints and family relationships only in women
- Higher BMI predicts poorer psychological well-being (women)
- Substantial racial/ethnic differences; stronger relationship between WC and inflammation in Black women
- What are the biological connections?



Block et al., Am J Epidemiol 2009;170:181–192; Bookwala & Boyar, *Psychology of Women Quarterly*, 32 (2008), 188–195; Stepanikova et al., Ethnicity & Health, 22:2, 169-183, DOI: 10.1080/13557858.2016.1235681

Prevalence of Self-Reported Obesity Among Non-Hispanic White Adults, by State and Territory, BRFSS, 2016-2018





Prevalence of Self-Reported Obesity Among Non-Hispanic Black Adults, by State and Territory, BRFSS, 2016-2018





*Sample size <50 or the relative standard error (dividing the standard error by the prevalence) \ge 30%.

Racial/ethnic differences



Prevalence of CES-D ≥16, frequent irritability, nervousness and psychological distress at baseline by race/ethnicity.

Bromberger Am J Pub Health 2001;91:1435-42; Freeman EW et al. AGP 2004;61:62-7

The Isms (Ag-, Rac-, Sex-)



- Subjective experiences of aging & positive affect
- Discrimination associated with physiological markers of stress and with poor psychological well-being
- Little research in sexual minority populations
- What are the biological pathways?

Barrett & Toothman, Journal of Women & Aging 2016; Brownlow et al., <u>Curr Psychiatry Rep.</u> 2019 Nov 4;21(11):112.

Outcome group	Outcome	r	Lower CI	Upper CI	z	p-value	k	Q-value	p-value Q
Negative mental health (NM)	DEP	-0.21	-0.23	-0.19	-18.40	<0.001	109	976.78	<0.001
	DIS	-0.22	-0.25	-0.19	-14.11	< 0.001	55	447.87	< 0.001
	STR	-0.27	-0.30	-0.23	-12.82	< 0.001	66	891.69	< 0.001
	ANX	-0.24	-0.29	-0.19	-9.50	< 0.001	40	249.34	< 0.001
	INT	-0.26	-0.34	-0.17	-5.65	< 0.001	9	39.11	< 0.001
	NA	-0.20	-0.24	-0.16	-10.00	< 0.001	23	69.73	< 0.001
	PTS/PTSD	-0.34	-0.40	-0.27	-8.96	< 0.001	16	68.48	< 0.001
	SOM	-0.23	-0.29	-0.17	-7.61	< 0.001	13	40.07	< 0.001
	SUI	-0.16	-0.19	-0.12	-8.57	< 0.001	10	3.76	0.927
	MHS	-0.21	-0.29	-0.12	-4.72	< 0.001	11	136.39	< 0.001
	GMH	-0.18	-0.24	-0.12	-5.55	< 0.001	12	48.84	< 0.001
	Overall NM	-0.23	-0.24	-0.21	-27.28	< 0.001	227	2278.70	< 0.001
Positive mental health (PM)	SE	-0.12	-0.15	-0.10	-9.28	< 0.001	78	284.86	< 0.001
	CON	-0.11	-0.14	-0.07	-5.92	< 0.001	18	40.56	0.001
	LS	-0.16	-0.22	-0.10	-5.35	< 0.001	29	295.24	< 0.001
	PA	0.00	-0.06	0.07	0.09	0.926	4	1.02	0.796
	WB	-0.19	-0.26	-0.12	-5.10	< 0.001	10	33.89	< 0.001
	Overall PM	-0.13	-0.16	-0.10	-9.36	< 0.001	113	945.00	< 0.001
Physical health (PH)	BP & HTN	0.00	-0.02	0.01	-0.24	0.814	24	25.78	0.312
	CHO	0.00	-0.02	0.02	-0.10	0.919	4	1.84	0.606
	DIA	-0.02	-0.09	0.04	-0.70	0.482	7	14.98	0.020
	HRT	0.00	-0.05	0.06	0.15	0.880	8	9.79	0.201
	OW	-0.08	-0.11	-0.05	-5.31	< 0.001	15	22.16	0.075
	Misc	-0.13	-0.18	-0.08	-5.15	< 0.001	20	251.00	< 0.001
	Overall PH	-0.09	-0.12	-0.06	-5.384	< 0.001	50	445.520	< 0.001
General health (GH)	GH	-0.13	-0.18	-0.09	-5.61	< 0.001	30	615.85	< 0.001

Table 3. Effect sizes for associations between racism and health outcomes.

DEP-Depression; DIS-Distress; STR-Stress; ANX-Anxiety; INT-Internalizing; NA-Negative affect; PTS/PTSD-Post-traumatic stress and post-traumatic stress disorder; SOM-Somatization; SUI-Suicidal ideation, planning, and attempts; MHS-Other mental health symptoms (e.g., paranoia, psychoticism); GMH-General mental health; Overall NM-Overall negative mental health; SE-Self-esteem; CON-Control/Mastery; LS-Life satisfaction; PA-Positive affect; WB-Wellbeing; Overall PM-Overall positive mental health; BP & HTN-Blood pressure and hypertension; CHO-cholesterol; DIA-Diabetes; HRT-Heart conditions/illnesses; OW-Overweight (BMI, WC, WHR, overweight, obesity); MISC-Miscellaneous physical health; Overall PH-Overall physical health; GH-General health (unspecified/ physical & mental)

doi:10.1371/journal.pone.0138511.t003

Paradies Y, Ben J, Denson N, Elias A, Priest N, et al. (2015) Racism as a Determinant of Health: A Systematic Review and Meta-Analysis. PLOS ONE 10(9): e0138511. https://doi.org/10.1371/journal.pone.0138511

https://journals.plos.org/plosone/article?id=10.1371/journal.pone.0138511

Inflammation/Immune

- Major depression associated with increased odds of allergy in midlife women but not men
- Greater negative affect reactivity associated with increased inflammation in women but not men
- Likely bidirectional but what are the causal pathways? How does this differ in different populations?
- What is the role of OTHER aspects of the immune system, and how do they intersect with other systems?



Goodwin et al., Psychosomatic Medicine 68:94–98 (2006)

Sleep

- Roughly half of the racial difference in cardiometabolic risk is due to sleep health
- Higher sleep latency associated with increased inflammation and insulin resistance in midlife, women only
- Sleep efficiency and total sleep time associated with higher BMI and waist circumference in women only
- What are the pathways connecting weight, inflammation, racism, sleep, and mental health?





Curtis et al., PNAS | August 15, 2017 | vol. 114 | no. 33 | 8889–8894; Kim et al., Sleep Medicine 27-28 (2016) 72e79; Mezick et al., Sleep Medicine 15 (2014) 64–70

Marriage, Mothering, Caregiving & Work

- Low levels of spousal support associated with higher inflammation among women but not men
- Intensive mothering associated with worse psychological well-being in midlife
- Mothers have similar work effort and intensity with other groups, but fathers and childless women report that home life relaxes and recharges them for work
- Women are much more likely to be caregivers
- For women only, having multiple roles is associated with HIGHER psychological well-being
- What elements can mitigate this?



Ahrens & Ryff, Sex Roles (2006) 55:801–815; Donoho et al., Journal of Marriage and Family 75 (February 2013): 127 – 141; Gunderson et al., Journal of Family Issues 2017, Vol. 38(7) 992– 1009; Kmec JA, Social Science Research 40 (2011) 444–459

Sexual Health and Function

- Sexual function closely correlated with relationship satisfaction and overall well-being
- 45% of midlife women report sexual problems
- Genitourinary changes, pelvic organ prolapse, medical problems and medications, partner issues
- Gaps
 - Sexual health education for providers
 - Screening tools
 - Protective factors
 - Changes in sexual minority groups
 - Treatment

Figure 2. Frequency of sexual activity among sexually active women in the Survey of Midlife Development in the United States (MIDUS II), by age (n = 1,345).



Thomas et al., ANNALS OF FAMILY MEDICINE, VOL. 13, NO. 4;2015; 337-342; Thomas et al., Obstet Gynecol Clin N Am 45 (2018) 709–722

Summary of Research Gaps

- Connection between psychosocial stressors and biological pathways
 - Stressors include early life adversity, discrimination, marital stress, caregiving, work
- Role of hormones other than estrogen
- Links with cognitive health
- Role of immune system (inflammation and beyond)
- Connections with sleep, weight
- Racial/ethnic differences
- Anxiety
- Sexual health and functioning