

Supplementary Information for

Examining Clinical Similarities between Myalgic Encephalomyelitis/Chronic Fatigue

Syndrome and D-Lactic Acidosis: A Systematic Review

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Supplementary Table 2. Episodes that reported *matching* or *ambiguous/other* D-lactic acidosis (D-la) symptoms as a function of age and sex.

Other supplementary material for this manuscript includes the following:

Additional file 1: Supplementary Table 1 (excel file) Demographic and clinical data summary of D-lactic acidosis episodes ($n = 59$) included in the qualitative synthesis.

Legend: All episodes simultaneously reported at least one high D-lactate level (from blood or urine analysis) and documented neurological symptoms. Episodes were screened for information about patient demographics, neurological symptoms, non-neurological symptoms, D-lactate levels, L-lactate levels, anion gap, pH levels, microbial composition, proposed triggers, medical history/comorbid conditions and treatment. Numbers in brackets (1) and (2) indicate separate episodes for the same patient. The letters *a* and *b* identify different patient cases reported in the same reference. Episodes from non-SBS patients are marked with an asterisk (*).

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Supplementary Table 2. Episodes that reported *matching* or *ambiguous/other* D-lactic acidosis (D-la) symptoms as a function of age and sex.

ME/CFS ICC	D-la _b symptom overlap with ME/CFS	Episode Case Numbers					
		Paediatric (≤ 17 years)			Adult (≥ 18 years)		
		Male	Female	NI	Male	Female	
A. Postexertional neuroimmune exhaustion	Matching	14, 48, 50, 66,	40	-	36, 42	12b, 13, 58b,	
		53, 8, 47a, 47b				39	
	Ambiguous/other	-	-	-	-	-	
B. Neurological impairments	Matching	48, 47a, 30,	37, 17, 51b,	11	12a, 10, 29,	6, 16 ₁ , 16 ₂ , 65,	
		19, 52, 56, 66,	4, 40, 35		44, 58a, 25,	20, 34 ₂ , 57,	
		43, 53, 14, 50,			24, 49, 64, 59,	28, 41, 58b,	
		8			25, 42, 62, 9,	60, 21, 12b,	
					15, 23, 27, 33	13, 34 ₁	
		Ambiguous/other	43, 56, 66, 14,	35, 37	11	9, 29, 44, 58a,	57, 21, 13, 31,
			18, 51a, 8, 19,			42, 64, 15, 24,	39, 58b, 43 ₂ ,
			48, 53, 30			25, 49, 55, 23,	6, 16 ₁ , 16 ₂ , 65,
						26, 10, 62, 27,	20, 34 ₁ , 12b
						33	
	Ambiguous/other B1-B4	43, 56, 66	35	11	9, 29, 44, 58a,	57, 21, 13, 31,	
					42, 64, 15, 24,	39, 58b	
					25, 49, 33		
	Speech/Language	14, 48, 66, 53,	37, 35	11	10, 15, 44, 62,	6, 16 ₁ , 16 ₂ , 65,	
		8, 30, 19			64, 9, 23, 25,	20, 34 ₁ , 57,	
					58a, 27, 33	21, 12b	
	Consciousness	14, 18, 51a, 8,	-	-	55, 9, 23, 26,	34 ₂ , 57, 21	
		19			49		
C. Immune, gastrointestinal and genitourinary	Matching	50, 47b, 66	-	-	10, 62, 42, 44,	16 ₂ , 60, 16 ₁ ,	
					36, 15	65	
	Ambiguous/other	-	-	-	-	-	

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D. Energy	Matching	56, 66, 48,	51b, 17, 35	-	55, 42, 25, 49,	60, 39
production/ transportation impairments		51a, 52, 8, 43, 19			9, 29, 62	
	Ambiguous/other	-	-	-	42, 23	-
Mood /	Matching	52, 53	40	-	9, 10	60, 31, 16 ₂
Behavior	Ambiguous/other	50, 56, 30, 66, 53, 19	-	-	44, 58a, 62, 33	31, 58b, 20
Uncategorized D-la Symptoms						
	Metabolic	14, 18, 48, 50,	35, 51b, 37,	11	10, 15, 29, 44,	6, 12b, 16 ₁ ,
	acidosis	51a, 52, 66, 53, 56, 8, 43, 47a, 47b, 30, 19	40, 4, 17		55, 62, 64, 9, 12a, 23, 24, 25, 26, 49, 58a, 59, 27, 36, 42, 33	16 ₂ , 60, 65, 13, 20, 31, 34 ₁ , 34 ₂ , 57, 58b, 21, 28, 41
	Other	66, 43, 47a,	-	-	9, 2, 15, 26,	16 ₁ , 58b
	abnormalities	47b			59, 33	

ambiguous/other: symptoms that were not clearly identified as consistent with ME/CFS presentation (see Table 2 for detailed symptom delineation); D-la: D-lactic acidosis; ICC: International Consensus Criteria; *matching*: mapped overlap between ME/CFS and D-la symptoms; ME/CFS: myalgic encephalomyelitis/chronic fatigue syndrome; NI: sex not identified.

Legend. Subscript numbers (1 and 2) indicate separate episodes for the same patient. The letters *a* and *b* identify different patient cases reported in the same reference. ME/CFS broad category B. Neurological impairments are highlighted as the primary focus of this review and to show three subcategories of delineation under *ambiguous/other* symptoms (i.e., in accordance with specific ICC criteria (B1 – B4), speech/language symptoms, and level of consciousness; see Table 2). Therefore, the same episode code number can be shown several times to represent multiple symptoms during each D-la episode (see Table 1 for references). Descriptions of drunkenness were referred to in several studies. Adult males self-reported *feeling* “drunk” (12a, 27) whereas females were described as *appearing* “drunk” (16₁, 16₂, 20,

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58b). Rather than using this ambiguous term, the specific symptoms that were also referred to in each of these studies were categorized in the table. See Table 3 for a summary of symptom frequencies.