

# Diabetic ketoacidosis associated with severe hypertriglyceridemia and acute pancreatitis in type 1 diabetes: 4 pediatric cases

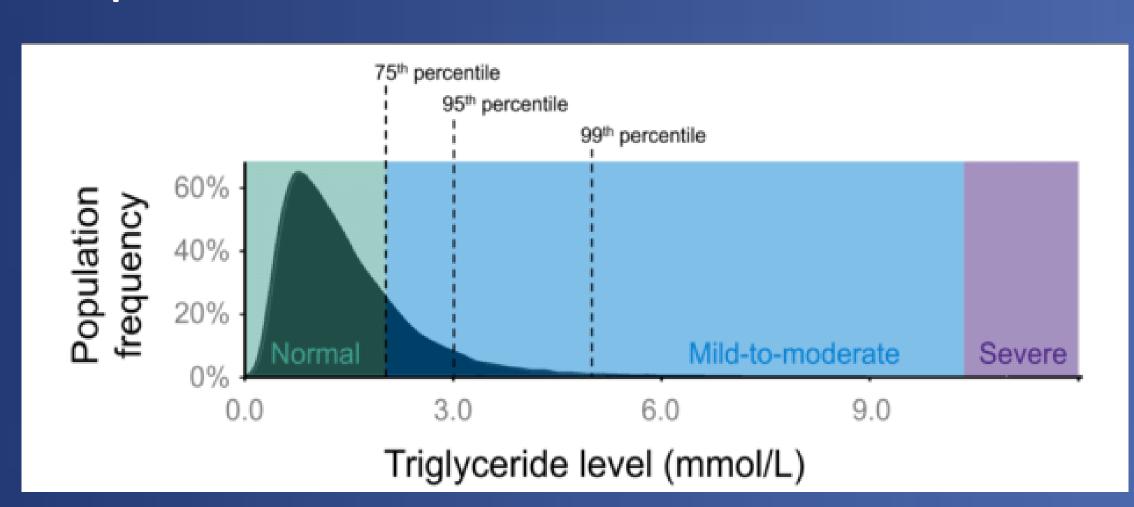


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## Background<sup>1</sup>

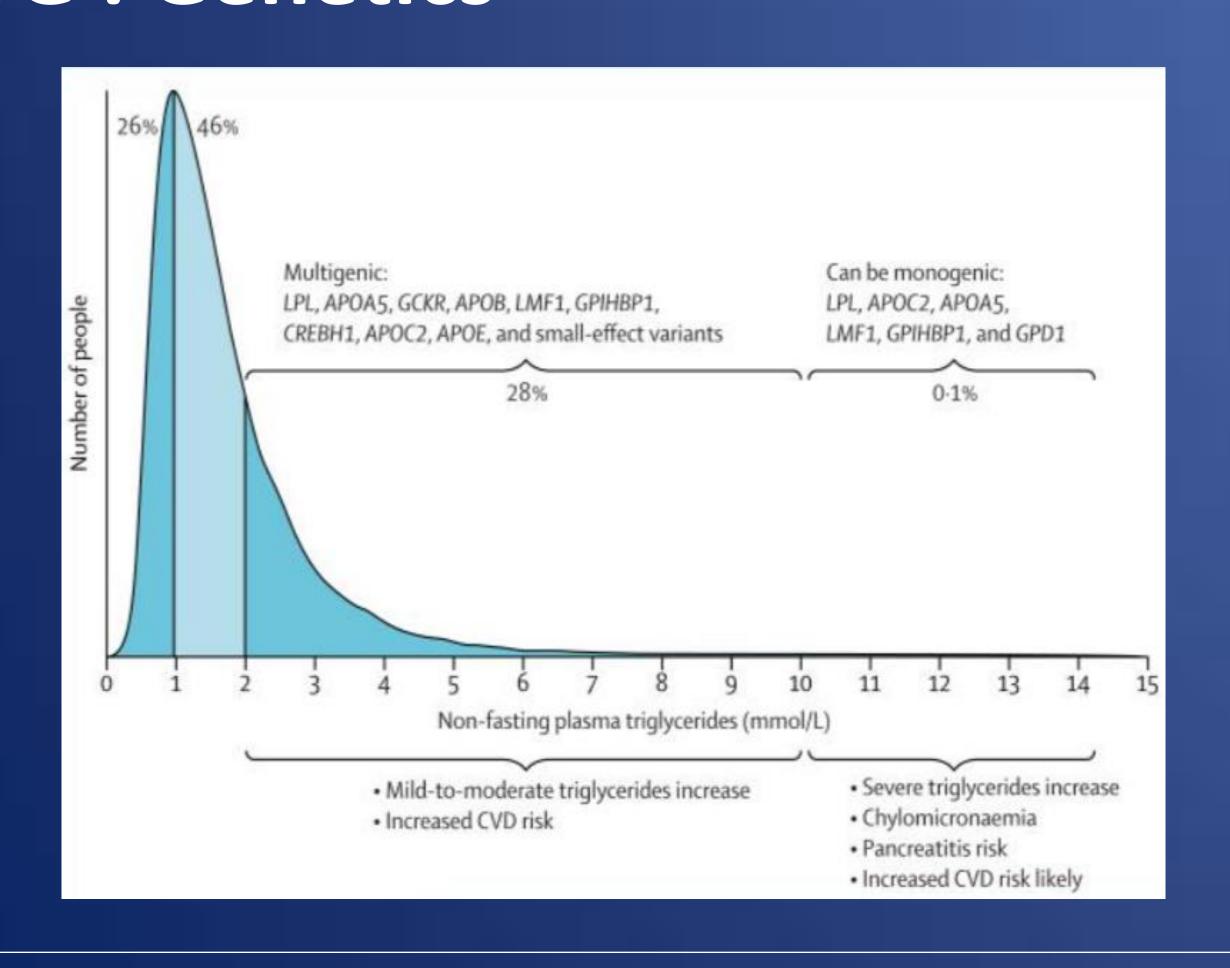
- Definition of severe hypertriglyceridemia (HTG): TG > 10 mmol/L, very severe HTG  $> 20 \text{mmol/L}^1$ .
- Most cases due to monogenic causes combined with secondary factors, such as obesity or medications.
- HTG due to diabetic ketoacidosis is rare, especially in pediatric patients.





Lipemic aspect of blood in hyperTG

# HTG: Genetics<sup>1</sup>



### Clinical cases

		Patient 1 : 17-year-old girl	Patient 2 : 13-year-old boy	Patient 3 : 13-year-old girl	Patient 4 : 4-year-old girl
Clinical presentation		<ul> <li>Obesity</li> <li>Polyuria-polydipsia x 1 week</li> <li>Vomiting x 24 hours</li> <li>Severe pancreatitis</li> </ul>	<ul> <li>Polyuria-polydipsia x few months</li> <li>Weight loss</li> <li>Vomiting x 24 hours</li> <li>Mild pancreatitis</li> </ul>	<ul> <li>Polyuria-polydipsia x 8 months</li> <li>No vomiting but abdominal pain</li> <li>No pancreatitis</li> </ul>	<ul> <li>Polyuria-polydipsia         x 2 weeks</li> <li>Vomiting x 24         hours</li> <li>Biochemical         pancreatitis</li> </ul>
Biochemistry					
	Glucose (mmol/L)	23,3	24	22,1	22,1
	Blood gas (pH/PCO2/Bicarb)	7,35/25/13,6	6,8/7,5/1,5	7,12/11,6/3,7	7,0/22/5
	Triglycerides (N : 0.4-1.3 mmol/L)	114,96	75,65	159,8	101
	Lipase (N : 4-39 U/L)	1725	375	82	451
Management		<ul> <li>Classical insulin infusion 0,1U/kg/h</li> <li>Plasmapheresis</li> </ul>	<ul> <li>High dose of insulin infusion 0,2U/kg/h</li> <li>Fenofibrates</li> </ul>	- High dose of insulin infusion 0,2U/Kg/h - Plasmapheresis	- Classical insulin infusion 0,1U/kg/h
Evolution		Normalization of clinical and biological parameters	Normalization of clinical and biological parameters	Normalization of clinical and biological parameters	Normalization of clinical and biological parameters
Genetics		Negative panel	VUS in the APOA5 gene	Negative panel	Heterozygous mutation of LPL gene : p.Gly215Glu

#### Conclusion

- Triad DKA-HyperTG-Pancreatitis is rare, particulary in pediatric population
- Some individual susceptibility factors like genetics are probably important for the development of the triad.
- But mechanisms still unclear: in our 4 patients, no correlation between the TG levels and the genetic risk factors.
- No consensus guidelines for the management

# References

<sup>1</sup>Hegele RA, et al. Lancet Diabetes Endocrinol. 2014 Aug;2(8):655-666. <sup>2</sup>Berberich AJ, et al. J Intern Med. 2019 Dec;286(6):644-650.

Literature review

PATHOPHYSIOLOGY of the triad 3-4-5

Prolonged lack of insulin

Lipolysis

Inhibition of lipoprotein lipase activity

Increase FFA and TG

Individual susceptibilit y factors

Pancreatic cell free radical injury

Edema

Capillary hyperviscosity

Pancreatitis

#### MANAGEMENT<sup>2</sup>: NO CONSENSUS

- IV hydratation
- Insulin infusion: 0,1 to 0,2 U/Kg/h
- Heparin infusion?
- Plasmapheresis: no clear guidelines
  - TG lowering 50-90% after one session
  - No benefit over conservative approach after 72 hours
  - No benefit on morbidity-mortality
  - Risk at central line insertion or transfusion reaction
- Fenofibates : not recommended in acute management
- <sup>3</sup> Wolfgram PM, et al. Journal of Pediatric Intensive Care. 2013 Jan;2(2):77-80.
- <sup>4</sup> Zaher FZ, et al. Case Rep Endocrinol. 2019 Jan 6;2019:8974619.

<sup>5</sup> Sharma PK, et al. Indian J Crit Car Med. 2017 March;21:176-178