

Pancreatic family history doesn't predict disease progression, but connotes alcohol consumption in adolescents and young adults with acute pancreatitis:



Analysis of an international cohort of 2,335 patients.

Márk Félix Juhász^{1,2}, Nelli Farkas¹, Andrea Szentesi^{1,2,3}, Andrzej Wedrychowicz⁴, Andreia Florina Nita⁵, Natália László⁶, Alexandra Tészás⁷, István Tokodi⁸, Áron Vincze⁹, Bálint Erőss^{1,2,10}, Ferenc Izbéki⁸, László Czakó¹¹, Mária Papp¹², Péter Hegyi^{1,2,10},
Andrea Párniczky^{1,2,13} on behalf of the Hungarian Pancreatic Study Group



¹Institute for Translational Medicine, Szentágotthai Research Centre, Medical School, University of Pécs, Pécs, Hungary; ²Centre for Translational Medicine, Semmelweis University, Budapest, Hungary; ³Centre for Translational Medicine, Department of Medicine, University of Szeged, Szeged, Hungary; ⁴Department of Pediatrics, Gastroenterology and Nutrition, Faculty of Medicine, Jagiellonian University Medical College, Krakow, Poland; ⁵Department of Paediatrics, Grigore Alexandrescu Emergency Hospital for Children, Bucharest, Romania; ⁶Department of Pediatrics, Szent János's Hospital and North Buda Unified Hospitals, Budapest, Hungary; ⁷Department of Paediatrics, University of Pécs Clinical Centre, Pécs, Hungary; ⁸Szent György University Teaching Hospital of Fejér County, Székesfehérvár, Hungary; ⁹Division of Gastroenterology, First Department of Medicine, Medical School, University of Pécs, Pécs, Hungary; ¹⁰Division of Pancreatic Diseases, Heart and Vascular Center, Semmelweis University, Budapest, Hungary; ¹¹Department of Medicine, University of Szeged, Szeged, Hungary; ¹²Department of Internal Medicine, Division of Gastroenterology, University of Debrecen, Debrecen, Hungary; ¹³Heim Pál National Pediatric Institute, Budapest, Hungary

INTRODUCTION

In pediatric acute pancreatitis (AP) a family history of pancreatic diseases is **prognostic for earlier onset of recurrent AP (ARP)** and chronic pancreatitis (CP). Adult CP guidelines also strongly recommend assessment (100% agreement), however, evidence is scarce for this recommendation – we failed to identify any clinical studies examining the connection between ARP, CP and pancreatic family history. **Age-specific reasons of familial aggregation are also unclear.**

We aimed to examine the prognostic role of pancreatic family history for ARP/CP and observe possible underlying mechanisms.

METHODS

We conducted a secondary analysis of the **Hungarian Pancreatic Study Group's (HPSG)** multicenter, international, prospective registry of **AP patients, both children and adults**. We compared those with positive and those with a negative family history of pancreatic diseases (AP, ARP, CP, autoimmune AP, pancreatic cancer), in different age groups, and analyzed trends of accompanying factors. Chi-square and Fisher exact tests were used.

DISCUSSION

While it is likely, that with higher patient numbers, the marked difference in ARP/CP in the pediatric subgroups would be retained, further increasing adult subgroups – while it could lead to significant results – would likely still be clinically irrelevant, and not suitable for ARP/CP prediction. In the positive group, we found an excess of idiopathic AP (likely reflecting genetic risk factors) in the pediatric age groups: peaking at 75% at 0-5 years then steadily decreasing to meet the negative group in adulthood. Next to the decline in idiopathic AP, we found a markedly higher rate of alcohol consumption / smoking among 12-29 year olds with positive family history. The most likely explanation is the well-documented association between parental and offspring alcohol consumption.

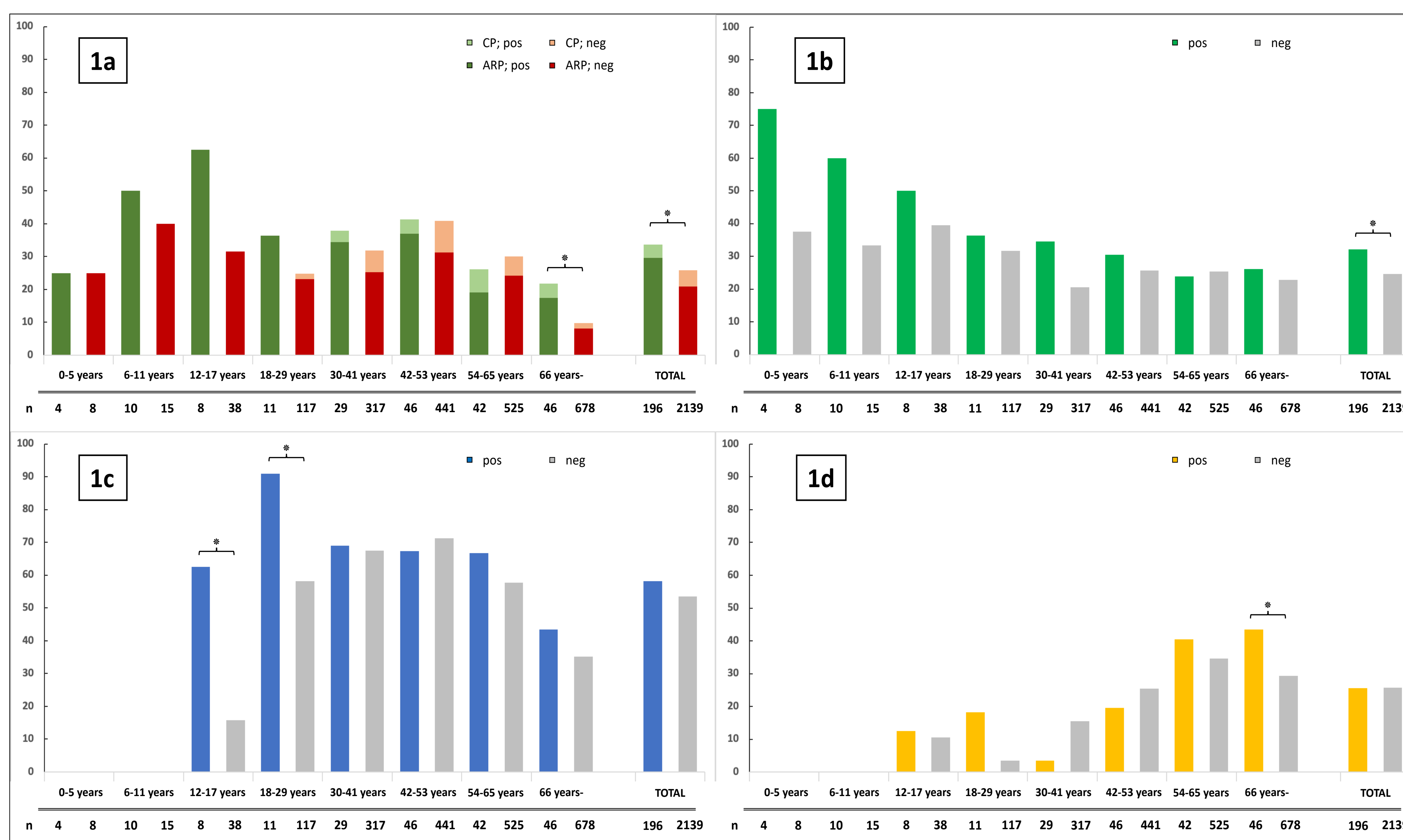


Figure 1: 1a: rate of acute recurrent pancreatitis (ARP) and chronic pancreatitis (CP) in different age groups of acute pancreatitis (AP) patients with positive (pos) and negative (neg) pancreatic family history; 1b: rate of idiopathic etiology at time of the index AP registry enrolment; 1c: rate of alcohol consumption and/or smoking; 1d: rate of diabetes and/or hyperlipidemia. *: statistically significant difference n: number of participants with available data on variables.

RESULTS

Higher rate of **ARP** was noted in childhood, even more so in the positive than the negative family history groups, but without statistical significance. Overall, a significantly higher rate of ARP or CP was found in the positive family history group (33.7% vs 25.9%, $p=0.018$) (Fig. 1a).

We found an **excess of idiopathic etiology in children with a positive family history** (75% 0-5 years, 60% 6-11 years) which decreased over time to meet the negative group (20-35%). Statistically significant difference was found overall (32.1% vs 24.6% in the positive vs negative groups, respectively, $p=0.020$) (Fig. 1b).

Current **alcohol consumption and/or smoking was significantly more common** in the positive family history group in ages: **12-17 years** (62.5% vs 15.8%, $p=0.013$), **18-29 years** (90.9% vs 58.1%, $p=0.049$) but not overall (58.2% vs 53.4%, $p=0.204$) (Fig. 1c).

Significant difference regarding the presence of **DM and/or hyperlipidemia** at the time of the index case was observed only in patients 66 years old or above (43.5% vs 29.4% respectively, $p=0.044$) but not overall (25.5% vs 25.7%, $p=0.950$) (Fig. 1d).

IMPLICATIONS

- **Positive family history most likely signifies genetic background in early childhood** (reassuring the literature)
- **During adolescence and early adulthood, alcohol consumption and smoking emerges – clinicians should be aware and turn to intervention in such cases.**
- **Contrary to current viewpoints positive pancreatic family history is not a prognostic factor for ARP and CP in adults, so it should not be used as such.**