

28/12/2020 ACTION-STATUS

Clinical/Lab center	Israel	Germany	Italy	Turkey	Poland
Action					
Collaboration agreement	done	done	done	done	done
Amendment No1 to the Collaboration Agreement	done	done	done	done	done
Money transfer to financial departments of each center (1 st part)	done	done	done	done	done
Settlement of expenses – Appendix 1 (modified)	done	done	done	done	done
Money transfer to financial departments of each center (2 nd part)	done	done	done	done	done
Use and settlement of funds	done	in progress	done	done	in progress
Recruitment procedures	-	-	done (Oct/Nov 2018)	done (May/June 2018)	done (June 2018)
1st year intervention (LifeStyle modification -LSM)	-	-	finished (Oct/Nov 2019)	finished (May/June 2019)	finished (June 2019)
2nd year con. intervention (only LSM) – according to appendix 2 sent previously (with previous checklist)	-	-	expected (Oct/Nov 2020) difficulties due to Covid-19	expected (May/June 2020) difficulties due to Covid-19	expected (June 2020) difficulties due to Covid-19
Database /readiness for analysis	1. 'CRF Questionnaire data/Physical exam/Lab tests (V0-V3-V6-V12) – DONE , database ready for analysis, sent to Michael Shmoish - part of basic data were presented - abstract ESPE 2019 2. Microbiome data: DONE , to transfer the data from Biobank to Michael Shmoish 3. Bile acids (urine) results: DONE , to transfer the data to Michael Shmoish 4. Steroid metabolites (urine) results: DONE . 'Clustering' based on steroid metabolome - DONE 5. Transcriptome analysis: Biobank is waiting for selecting the samples (~50 samples/representatives of clusters based on steroid metabolome clusters or/and clinical profile?) – is being prepared 6. Adipokines: DONE , the database was modified.				
Drop out V3/V6/V12	-	-	V3:..... V6:..... V12:~50%	V3:..... V6:..... V12: ~ -25%	V3:-5 V6: -14 V12: -20 (-25%) (in total)

Transport and analysis of the samples					
2 urine samples (V0) from each patients +10 representative ¹ samples (V6) to Giessen	-	-	done	done	done
1 fecal sample (V0) to Biobank/ Lodz ²	-	-	done (70 samples were sent, in IT08, IT30, IT48, IT58, IT70, IT71, IT 77 the repeated isolations were not successful – backup samples are not available)	done (81 samples were sent, in TR56, TR66,TR72, TR78 the repeated isolations were not successful – backup samples are not available)	done (83 samples were sent, in PL16 the repeated isolation was not successful)
50 representative PAXgene samples (tanscriptome: total RNA+miRNA V0) ³ –to Biobank/Lodz (the same lab like fecal samples for microbiome)	-	-	<ul style="list-style-type: none">- samples collected – transport to Biobank/Lodz: DONE- the selection of 50 representative samples – IN PROGRESS		
Samples of the serum for leptin/adiponectin to Katowice ²	-	-	DONE, results are added to database		
Genomics			<ul style="list-style-type: none">-Waiting for decisions concerning funds – despite repeated contact Sandoz did not respond (Roberta Luzzi)-examination of polymorphisms in the same samples prepared for transcriptome analysis -Biobank (Lodz) will check if it possible ⁴		
1 st Report to ESPE	Done: The financial (interim) report was sent in JUNE 2020, the next one will be sent in JUNE 2021				
2 nd /Final Report to ESPE (should be after two years)	The ESPE Science Committee agreed to my proposed delay on the project under the condition that a short report is received every 6 months in order to update the Science Committee on progress				
Qs?/Risks?: <ul style="list-style-type: none">- Patients drop out- money for transcriptome/adipokines- money for material analysis (stored) after intervention (collected at V6, V12,...V24?)	<ul style="list-style-type: none">- <u>Drop out</u> – 163/240 (67.9%) finished 12-month follow-up- <u>Transcriptome</u>: to do the analyses of total mRNA (together with miRNA) in 50 samples; Medical University of Silesia will add ~21K EUR from its budget- <u>Leptin/adiponectin (V0)</u> – done based on Medical University of Silesia budget- <u>Databases format</u>:<ul style="list-style-type: none">-all details should be established with Michael Shmoish				

- databases format	(email address added). -have to be careful about all details in adding data (excel cells' format, save the same format of document etc.) - <u>All collected and stored samples (V6, V12) stay in the local centers – seeking funding</u>
Qs? - next meeting /workshops? - type of contact: e-mails; skype? Whatsapp group	- Workshops/online meetings are planning: to discuss all clinical / lab results
Dissemination/Popularization of the project's idea/ project's results	- ESPE meeting Sep 2019 (abstract P1-53) - ECOG meeting Nov 2019 – 1h session: - to present general idea of MULTI-OMICS project (Aneta Gawlik) - to present part of clinical data (Malgorzata Wasniewska) (the discussion concerned mainly how detailed is our questionnaire, they were impressed☺) - <u>A review article regarding our project is being prepared</u> - <u>ESPE 2021: to present data</u>

V0- 1st visit (recruitment visit), V6 – visit after 6 months, V12- V after 12 months, Q-query/Qs - queries

¹ V6- the samples of urine (24hrs) for steroid metabolites/bile acids are in Giessen (Prof S.Wudy)

² - We collected two types of samples (fecal/adipokines) in one transport to Łódź/Poland – the samples for adipokines were transferred in dry ice from Łódź to Katowice by our team.

³ - Waiting for decision about miRNA – not sure it will be technically possible to have total RNA + miRNA from the same sample – waiting for Biobank answer

⁴ – examination of polymorphisms using the same samples prepared for transcriptome analysis (by Biobank) – all PAXgene samples were sent to Biobank

Details concerning transport of the samples (please see all details in e-mail sent 14th Feb)

Address for fecal samples+ adipokines (serum):

**Biobank Lab, Head
University of Lodz
Pilarskiego 14 Street
90- 237 Lodz, Poland**

phone +48426655702

(www.biobank.uni.lodz.pl)

Delegate of Polish National Node to the Assembly of Members BBMRI-ERIC)

Address for steroid metabolites/metabolome +bile acids (urine):

**Prof. Dr. Stefan Wudy and Dr. Michaela Hartmann
Steroid Research and Mass Spectrometry Laboratory
Division of Pediatric Endocrinology and Diabetology
Children's Hospital
Feulgenstrasse 12
35392 Giessen
Germany**