D7.1 Report summarising results of survey of EU countries to identify volumes and trends in relation to the import and export of stem cells

Grant Agreement number:	785270
Project acronym:	SAVDON
Work Package number:	WP7

Period covered:

from 01<sup>st</sup> January to 31<sup>st</sup> December 2017

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### Abbreviations

- BM = Bone Marrow
- CBU = Cord Blood Unit
- DLI = Donor Lymphocyte Infusion
- HLA = Human Leukocyte Antigen
- HSC = Hematopoietic Stem Cells
- PBSC = Peripheral Blood Stem Cell
- WMDA = World Marrow Donor Association

## **Executive Summary**

This World Marrow Donor Association Annual Report describes the 2017 import and export data, trends and insight of unrelated blood stem cell products between the different EU member states and third countries. In 2017, almost 1,5 million volunteer individuals were recruited and HLA typed into unrelated stem cell donor registries within the EU member states and 10,000 cord blood units were added to the inventory increasing the total number of available stem cell donors within the EU member states to 13,344,277 and worldwide over 33 million.

In 2017: within the EU member states 3,707 adult stem cell donations were imported and 3,719 exported and 12 cord blood products were imported and 172 exported.

In 2017, there were 3,429 adult stem cell donations exported to 19 countries outside of the EU member states and 954 adult stem cell donations imported from these countries. Additionally, 292 cord blood products where exported to countries out of the EU member states and 134 cord blood products were imported.

### Introduction

#### Blood stem cell transplantation

Every year, thousands of patients around the world are diagnosed with leukemia and other bloodrelated diseases. In the past, this diagnosis was often fatal. Today, a blood stem cell transplant can be a potentially life-saving treatment for more than 70 different diseases. In a blood stem cell transplantation, a patient's defective stem cells are replaced with healthy ones from a donor. More than 50,000 patients worldwide receive transplants every year, and this number continues to rise.(1)

A blood stem cell is an immature cell that can develop into all types of blood cells, including white blood cells, red blood cells and platelets. Stem cells can be mostly found in the bone marrow (a spongy tissue inside the bones). Small numbers also are found in the peripheral blood (the flowing, circulation blood of the body) and the umbilical cord (the cord that connects a foetus to its mother's placenta). (2-4)

Stem cells develop into the three types of blood cells that the body needs:

- Red blood cells, which carry oxygen throughout the body
- White blood cells, which fight infections
- Platelets, which help the blood to clot

There are three possible sources of blood stem cells for donation:(4)



#### **Bone marrow**

(Collected from the pelvis) The pelvis contains the largest number of blood stem cells. For this reason, cells from the cavity in the pelvic bone are used for a bone marrow donation.



#### Peripheral blood

(collected from the blood stream) Normally, only very few stem cells are found in the blood. Hormone-like substances called growth factors can be given to blood stem cell donors a few days before the collection. This causes their blood stem cells to grow faster and to move from the bone marrow into the blood stream. Peripheral blood is collected from adult donors.



#### **Umbilical cord blood**

(collected from the placenta and umbilical cord)

After a baby is born, the blood that is left behind in the placenta and umbilical cord (known as cord blood) can be collected. It is then frozen and stored until required for transplantation. In addition to the blood stem cell products, Donor Lymphocyte Infusion (DLI) can be applied. This is a type of therapy in which lymphocytes from the blood of a donor are given to a patient, who has already received a stem cell transplant from the same donor. The donor lymphocytes may kill remaining cancer cells. DLI is used to treat chronic myelogenous leukaemia (CML) that has come back and myeloma. It is being studied in the treatment of other types of cancer.(5)

#### HLA matching

Patients receiving blood stem cell transplants are matched with a volunteer donor through a test called HLA tissue typing. The immune system uses HLA proteins found in the surface of cells to identify which cells belong to you own body and which not. For HLA matching the patient as well as the potential donors will have blood drawn. This blood is tested in a specialised lab to figure out the HLA type. This will be compared to potential donors to see if there is a match. It could be difficult to find a match because some HLA types are less common. Patients are more likely to match someone with a similar ethnic background or ancestry. Because HLA markers are inherited, half from the mother and half from the father, each brother and sister who shares the same parents has a 25% chance of being a close HLA match. Unfortunately, many people do not have a good match within their family. A close match is important since it improves the chances for a successful transplant, it helps the donor cells engraft (grow and make new blood cells in the body) and reduces the risk of complications like graft-versus-host disease (GVHD). GVDH happens when the immune cells from the donated cells (the graft) attack the patient's body (the host). (6, 7) New donors should from the beginning be HLA typed in high resolution to avoid extra costs and time. Advanced resolution and precision of the exact HLA typing will lead to a better match and a faster donor choice in the Search and Match system.

#### Global network

Millions of volunteers have signed up to be a donor worldwide, and tens of thousands of patients receive blood stem cells from a healthy individual every year. Unfortunately, not every patient is able to find a potential match in the global database. This is more volunteers are needed to sign up as donors. WMDA services the global network by providing a centralised database of all potential donors around the world.

#### Import / export

This report shows the importance of global exchange of stem cell products. Nowadays people have spread all over the world which means that certain populations do not live in the same region anymore as they use to. Unfortunately, not every country has an established network for recruiting donors. Furthermore, smaller countries may not have enough volnteer donors or CBUs themselves. They rely on the international network of blood stem cell donor registries to find a suitable stem cell source.

The list of third countries who export from and import to the EU member states is shown in Appendix 1.

#### Questionnaire

In Appendix 2 the WMDA Questionnaire is shown with the questions regarding adult donor and CBU numbers and import and export data that were answered by the different registries worldwide.

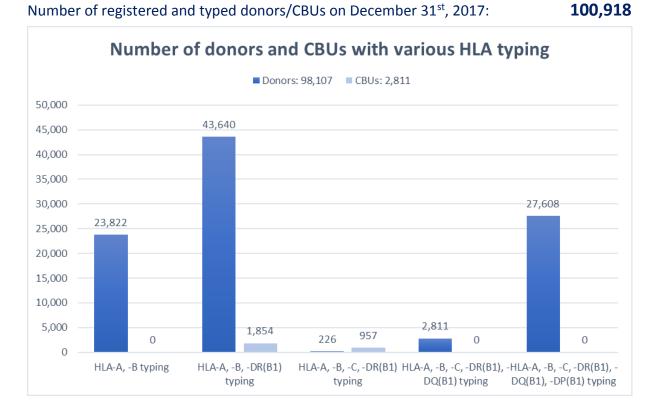
Data from Latvia, Estonia and Malta is missing since these countries do not have a donor registry or a cord blood bank. Estonia has a collaboration with Finland to find their donor matches, so they import stem cell sources from other countries. Lithuania, Luxembourg, Portugal, Romania, Bulgaria, Denmark, Hungary and Ireland do not have a cord blood bank, so they import cord blood products from other countries.

In Appendix 3 the Global Trends movie is shown with a summary of the achievements over the past 20 years worldwide.

The following pages show the report per EU member state including the import and export data of the different stem cell products.

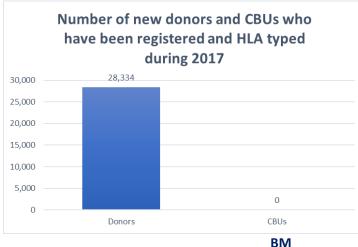
Austria





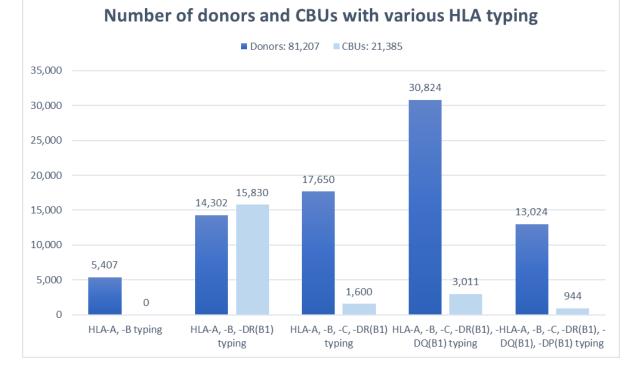
### Adult volunteer donors/CBUs added to the database during 2017:

28,334



	BM	PBSC	CBU	DLI
National	1	3	0	3
Export to EU member states	1	7	1	2
Export to third countries	5	12	0	0
Import from EU member states	14	94	7	6
Import from third countries	3	7	7	0

Belgium

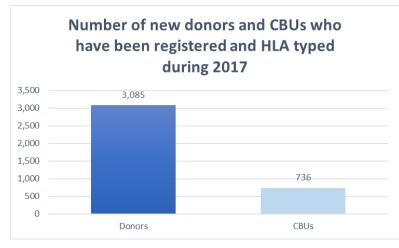


## Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:

### Adult volunteer donors/CBUs added to the database during 2017:

3,821

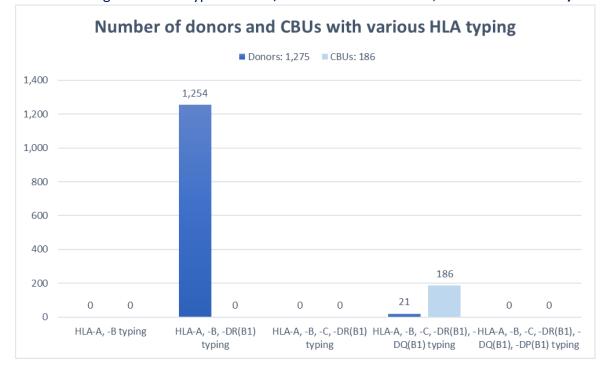
102,592



	BM	PBSC	CBU	DLI
National	0	9	3	1
Export to EU member states	1	21	10	3
Export to third countries	1	2	2	0
Import from EU member states	21	155	5	24
Import from third countries	4	36	3	5

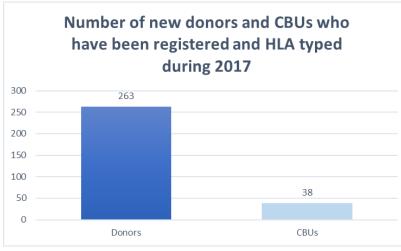
Bulgaria





### Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017: **1,461**

### Adult volunteer donors/CBUs added to the database during 2017:



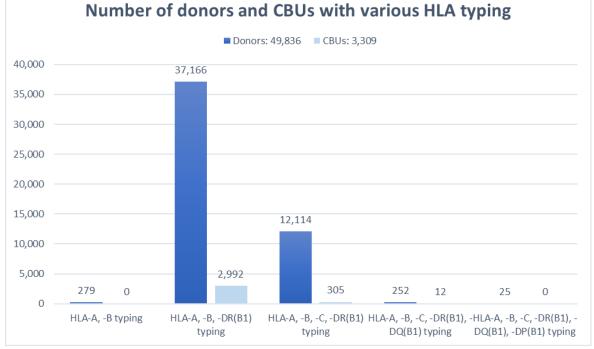
	BM	PBSC	CBU	DLI
National	0	0	-	0
Export to EU member states	0	0	-	0
Export to third countries	0	0	-	0
Import from EU member states	6	15	-	0
Import from third countries	0	1	-	0

263

Croatia

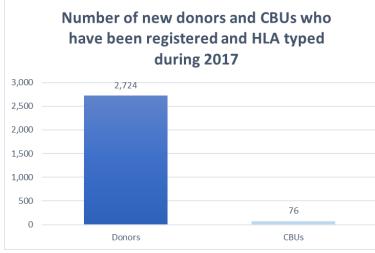






Adult volunteer donors/CBUs added to the database during 2017:





	BM	PBSC	CBU	DLI
National	1	12	0	0
Export to EU member states	1	0	0	0
Export to third countries	1	0	1	0
Import from EU member states	7	22	0	2
Import from third countries	1	2	0	0

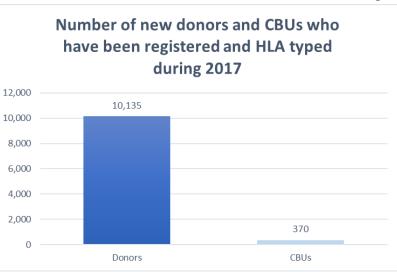
Cyprus



#### Number of donors and CBUs with various HLA typing Donors: 158,104 CBUs: 3,069 70,000 61,759 60,000 50,000 44,093 40,000 30,000 19,591 18,485 20,000 14,176 10,000 3,066 0 3 0 0 0 HLA-A, -B typing HLA-A, -B, -DR(B1) HLA-A, -B, -C, -DR(B1) HLA-A, -B, -C, -DR(B1), -HLA-A, -B, -C, -DR(B1), typing typing DQ(B1) typing DQ(B1), -DP(B1) typing

### Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017: **161,173**

### Adult volunteer donors/CBUs added to the database during 2017:

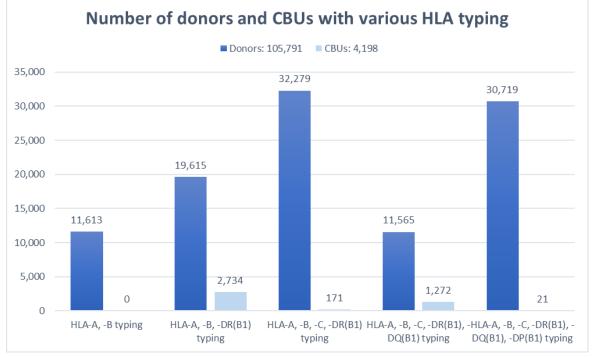


BM	PBSC	CBU	DLI
2	7	0	1
1	23	0	1
1	18	0	1
0	0	0	0
0	0	0	0
	BM 2 1 1 0 0	2 7 1 23	2     7     0       1     23     0

10,505

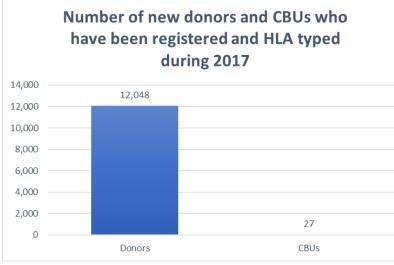
**Czech Republic** 





Adult volunteer donors/CBUs added to the database during 2017:



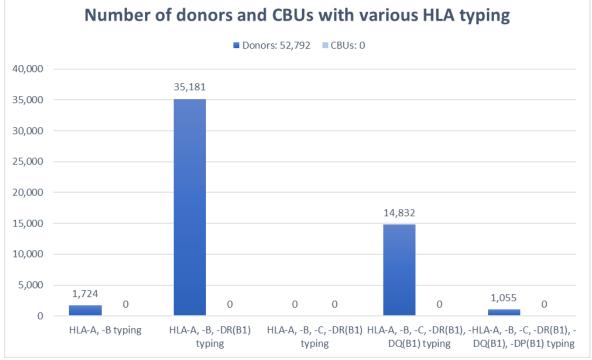


BM	PBSC	CBU	DLI
4	24	1	7
3	5	0	1
2	2	0	1
12	106	0	12
1	10	0	2
	BM 4 3 2 12 1	4     24       3     5       2     2       12     106	4 24 1   3 5 0   2 2 0   12 106 0

Denmark

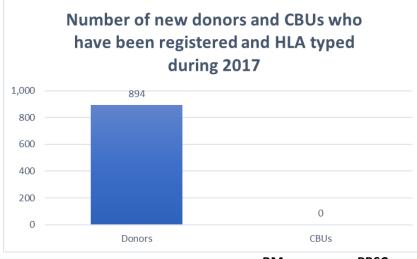






Adult volunteer donors/CBUs added to the database during 2017:





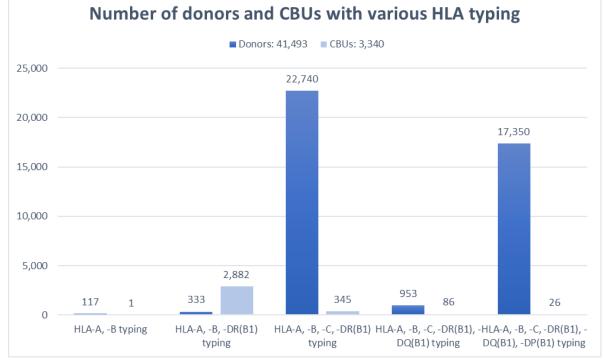
	BM	PBSC	CBU	DLI
National	0	0	-	0
Export to EU member states	0	9	-	1
Export to third countries	0	0	-	0
Import from EU member states	24	82	2	0
Import from third countries	6	10	0	0

# Finland



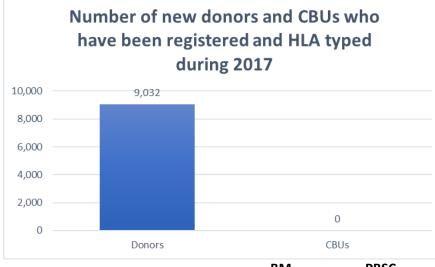


£ .::



Adult volunteer donors/CBUs added to the database during 2017:



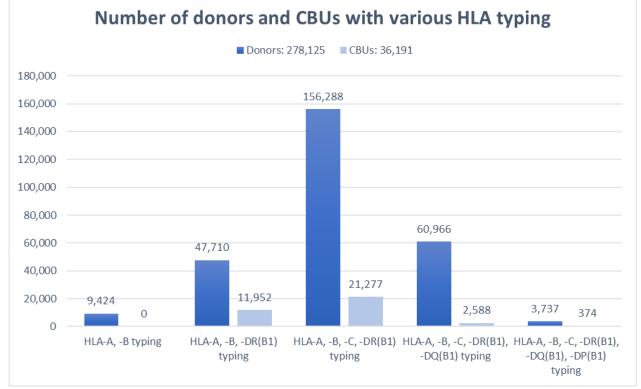


	BM	PBSC	CBU	DLI
National	7	20	1	2
Export to EU member states	3	5	0	1
Export to third countries	0	3	1	0
Import from EU member states	10	58	0	6
Import from third countries	1	7	0	1

France

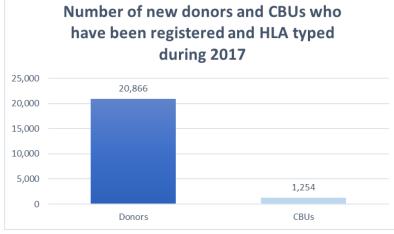






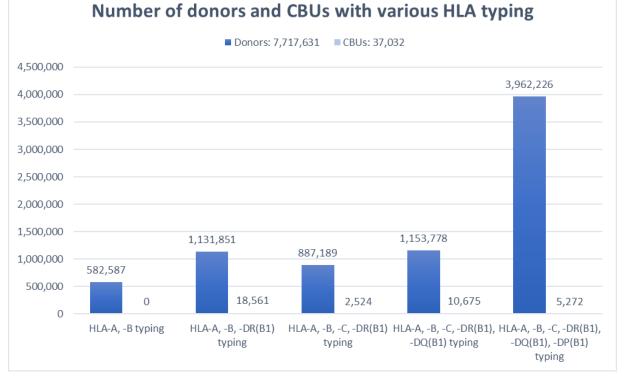
Adult volunteer donors/CBUs added to the database during 2017:





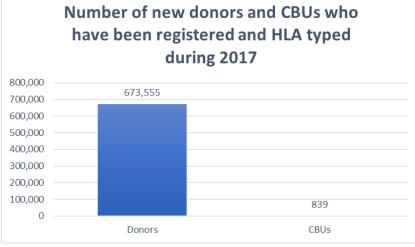
	BM	PBSC	CBU	DLI
National	26	96	45	19
Export to EU member states	13	37	40	4
Export to third countries	5	18	67	1
Import from EU member states	126	538	43	77
Import from third countries	41	98	22	15

## Germany



Adult volunteer donors/CBUs added to the database during 2017:





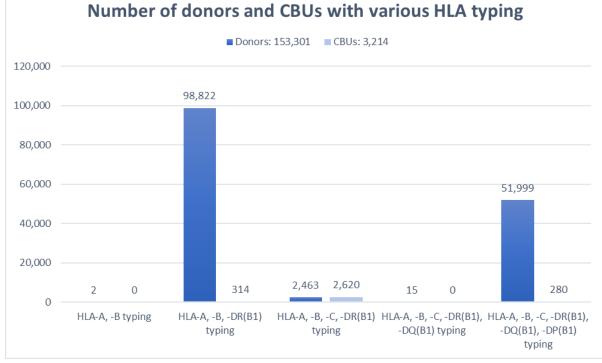
	BM	PBSC	CBU	DLI
National	228	1,674	1	277
Export to EU member states	452	2,090	28	254
Export to third countries	557	1,915	50	135
Import from EU member states	23	307	0	43
Import from third countries	23	169	0	13

7,754,663

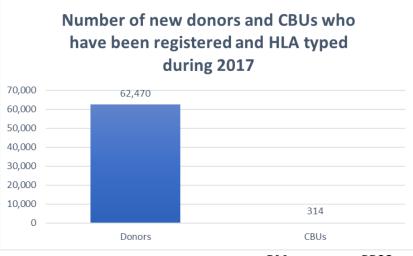
Greece



#### Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:



#### Adult volunteer donors/CBUs added to the database during 2017:



	BM	PBSC	CBU	DLI
National	4	17	1	2
Export to EU member states	0	16	0	2
Export to third countries	0	6	0	0
Import from EU member states	8	46	4	2
Import from third countries	1	9	3	5

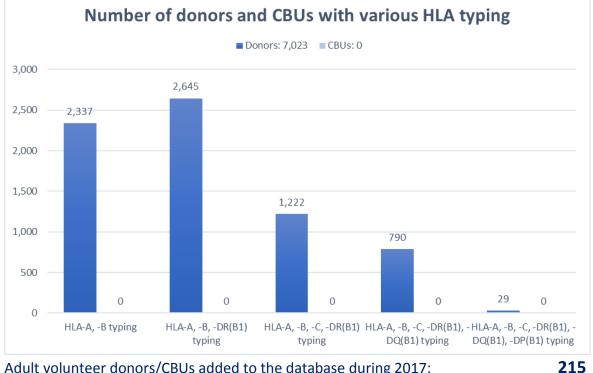


Hungary

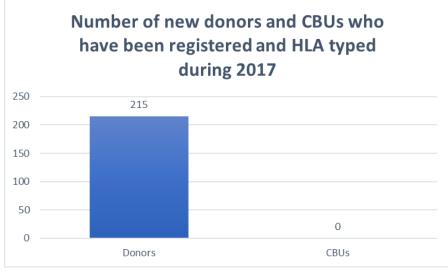








Adult volunteer donors/CBUs added to the database during 2017:

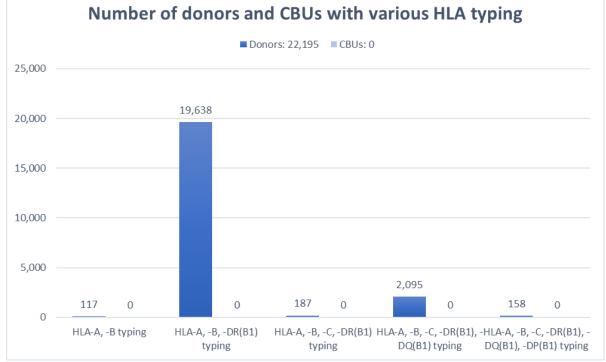


	BM	PBSC	CBU	DLI
National	0	0	-	0
Export to EU member states	0	0	-	0
Export to third countries	0	0	-	0
Import from EU member states	11	49	4	6
Import from third countries	2	4	0	1

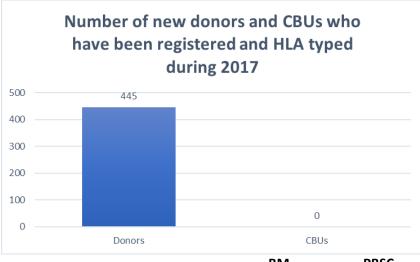
Ireland



#### Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:



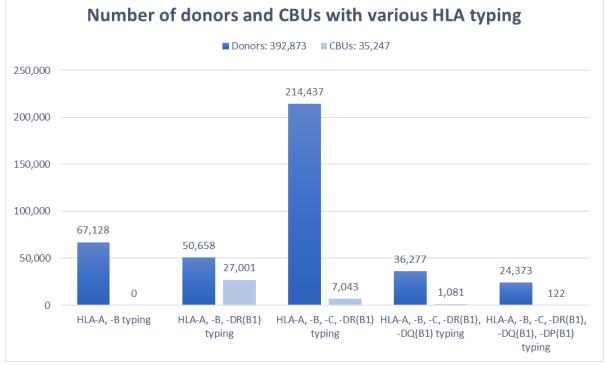
Adult volunteer donors/CBUs added to the database during 2017:



	BM	PBSC	CBU	DLI
National	1	2	-	1
Export to EU member states	0	1	-	0
Export to third countries	0	2	-	0
Import from EU member states	14	31	0	4
Import from third countries	1	6	0	0

445

Italy

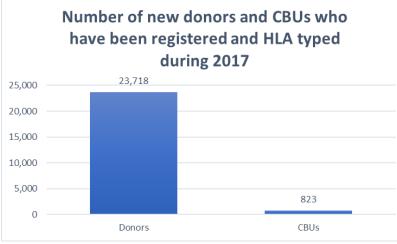


### Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:

Adult volunteer donors/CBUs added to the database during 2017:



428,120

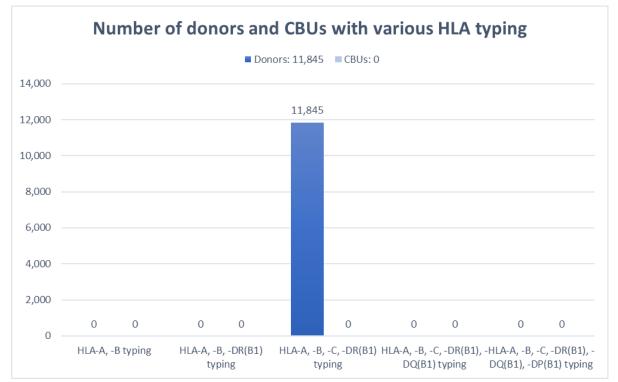


	BM	PBSC	CBU	DLI
National	48	111	2	15
Export to EU member states	8	37	12	7
Export to third countries	9	15	15	2
Import from EU member states	146	357	13	42
Import from third countries	42	126	13	12

# Lithuania

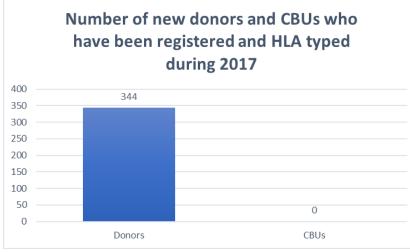


11,845



Adult volunteer donors/CBUs added to the database during 2017:



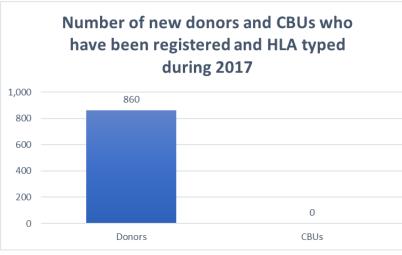


	BM	PBSC	CBU	DLI
National	0	10	-	0
Export to EU member states	0	3	-	0
Export to third countries	0	1	-	0
Import from EU member states	7	25	0	1
Import from third countries	0	3	0	1

# Luxembourg

#### Number of donors and CBUs with various HLA typing Donors: 7,941 CBUs: 0 2,500 2,338 2,264 2,000 1,474 1,391 1,500 1,000 474 500 0 0 0 0 0 0 HLA-A, -B, -C, -DR(B1) HLA-A, -B, -C, -DR(B1), -HLA-A, -B, -C, -DR(B1), -HLA-A, -B typing HLA-A, -B, -DR(B1) DQ(B1) typing DQ(B1), -DP(B1) typing typing typing

Adult volunteer donors/CBUs added to the database during 2017:



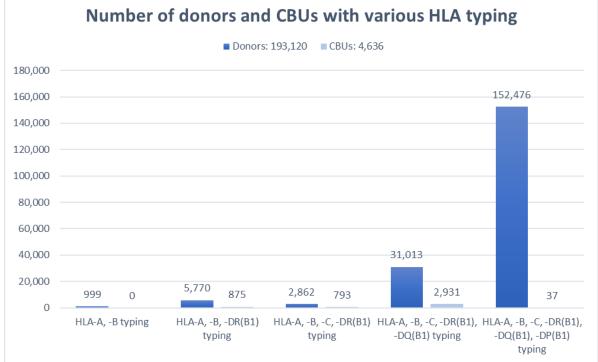
	BM	PBSC	CBU	DLI
National	0	0	-	0
Export to EU member states	0	1	-	0
Export to third countries	1	0	-	1
Import from EU member states	0	0	0	0
Import from third countries	0	0	0	0



860

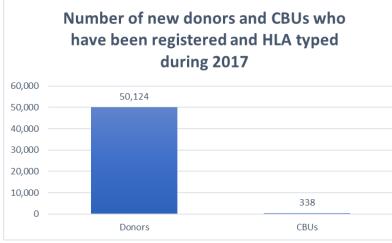


# Netherlands



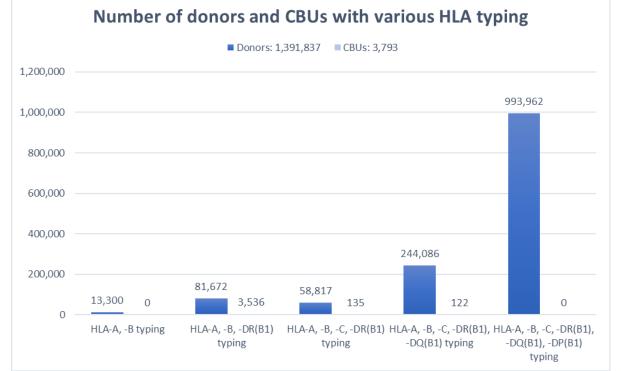
Adult volunteer donors/CBUs added to the database during 2017:





	BM	PBSC	CBU	DLI
National	3	13	3	3
Export to EU member states	14	41	1	3
Export to third countries	11	17	2	1
Import from EU member states	23	254	49	46
Import from third countries	5	39	27	8

Poland

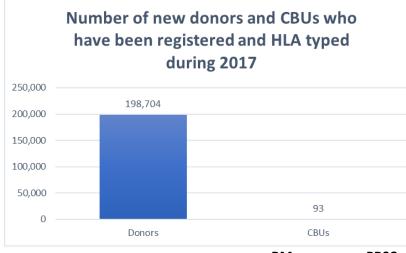


Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:





1,395,630

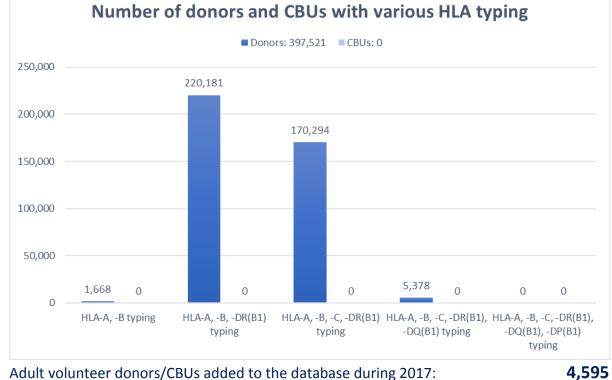


	BM	PBSC	CBU	DLI
National	23	250	0	15
Export to EU member states	77	487	0	55
Export to third countries	75	315	0	18
Import from EU member states	19	121	1	3
Import from third countries	2	21	0	0

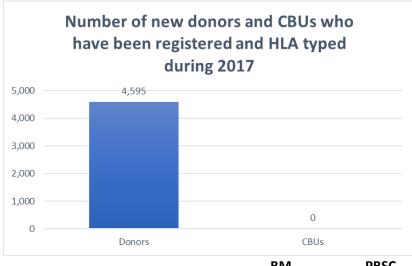
# Portugal

Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:





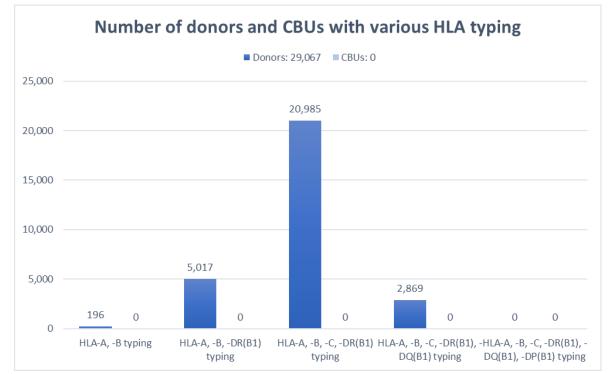
Adult volunteer donors/CBUs added to the database during 2017:



BM	PBSC	CBU	DLI
8	32	-	0
4	26	-	5
6	13	-	0
10	22	0	3
4	4	1	0
	8 4 6	8     32       4     26       6     13       10     22	8     32     -       4     26     -       6     13     -       10     22     0

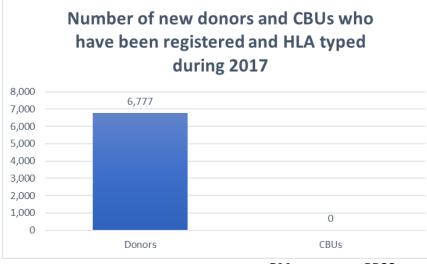
# Romania





Adult volunteer donors/CBUs added to the database during 2017:



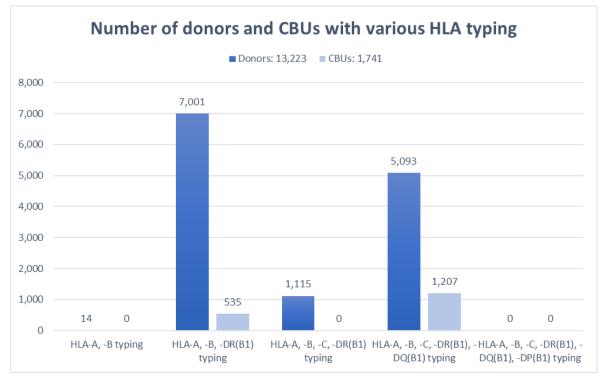


	BM	PBSC	CBU	DLI
National	0	1	-	0
Export to EU member states	1	2	-	0
Export to third countries	0	0	-	0
Import from EU member states	1	33	0	2
Import from third countries	0	1	0	0

Slovakia

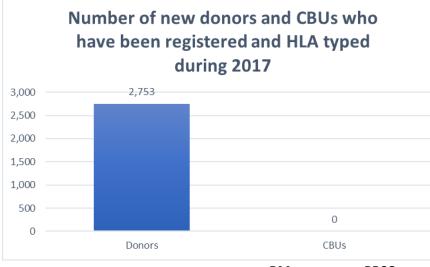


14,964



Adult volunteer donors/CBUs added to the database during 2017:



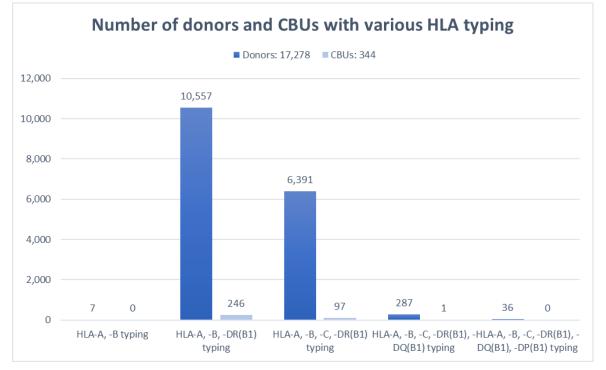


	BM	PBSC	CBU	DLI
National	0	1	0	0
Export to EU member states	0	0	3	0
Export to third countries	0	1	0	0
Import from EU member states	8	34	0	0
Import from third countries	0	2	0	0

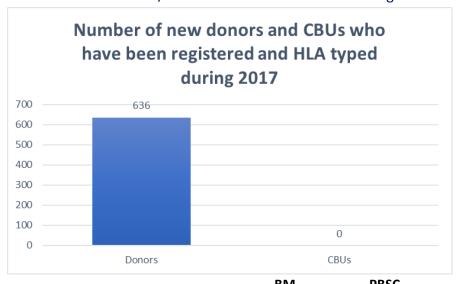
Slovenia



636



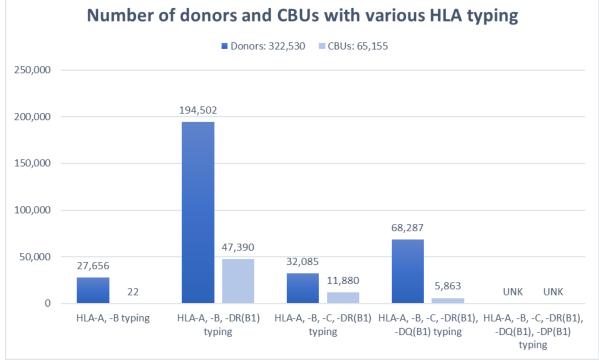
Adult volunteer donors/CBUs added to the database during 2017:



	BM	PBSC	CBU	DLI
National	0	1	0	0
Export to EU member states	0	1	0	0
Export to third countries	0	2	0	0
Import from EU member states	0	24	1	1
Import from third countries	0	2	1	1

Spain



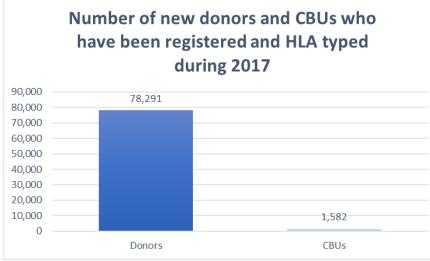


Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:

387,685

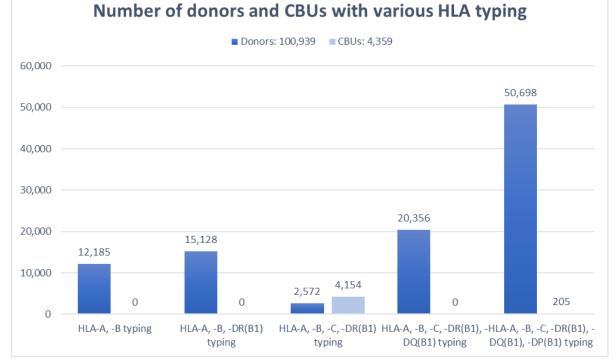
Adult volunteer donors/CBUs added to the database during 2017:





	BM	PBSC	CBU	DLI
National	14	69	21	1
Export to EU member states	8	31	53	1
Export to third countries	3	21	103	0
Import from EU member states	46	200	11	12
Import from third countries	14	81	15	2

Sweden

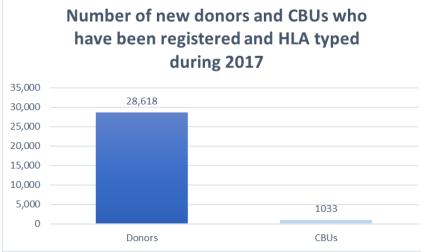


Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:





105,298

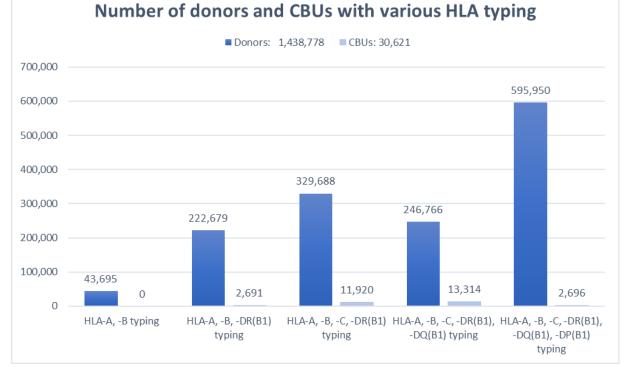


BM	PBSC	CBU	DLI
0	31	0	2
3	15	5	0
2	14	4	1
26	117	0	7
8	17	1	0
	0 3 2	0     31       3     15       2     14	0     31     0       3     15     5       2     14     4





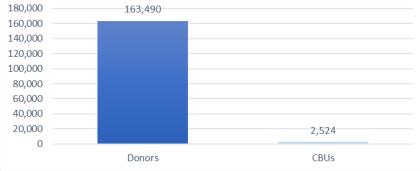
# United Kingdom



Number of registered and typed donors/CBUs on December 31<sup>st</sup>, 2017:

### Adult volunteer donors/CBUs added to the database during 2017:





	BM	PBSC	CBU	DLI
National	49	392	28	59
Export to EU member states	36	217	19	23
Export to third countries	77	292	46	8
Import from EU member states	63	392	28	59
Import from third countries	22	116	44	7

166,014

## Appendix 1: List of countries that export to EU member states

- Argentina
- Australia
- Brazil
- Canada
- China
- India
- Iran
- Israel
- Japan
- Korea
- Norway
- Russia
- Singapore
- South Africa
- Switzerland
- Taiwan province of China
- Thailand
- Turkey
- United States of America

## Appendix 2: WMDA Questionnaire

#### QUESTIONNAIRE

## Online Questionnaire - Year 2017 blank questionnaire in PDF NOTE: Please do not fill in the green boxes, these boxes will be automatically calculated based on your response. Only fill out 0 if the answer is 0 • If the answer is unknown, fill out UNK If the answer is not applicable, fill out NA • It is not possible to leave any answers blanc (except for the country list) General registry and cord blood registry/bank data hide questionnaire 1) Number of registered and typed donors/CBUs on December 31 2017 (please give number of donors/CBUs with specific HLA typing profile regardless of HLA typing resolution, please list a donor/CBU only once) 1a) Number of adult volunteer donors (all donors) 1a1) Number of donors with HLA-A, -B typing 1a2) Number of donors with HLA-A, -B, -DR(B1) typing 1a3) Number of donors with HLA-A, -B, -C, -DR(B1) typing 1a4) Number of donors with HLA-A, -B, -C, -DR(B1), -DQ(B1) typing 1a5) Number of donors with HLA-A, -B, -C, -DR(B1), -DQ(B1), -DP(B1) typing 1b) Number of HLA typed CBUs available for transplantation for unrelated patients 1b1) Number of CBUs with HLA-A, -B typing 1b2) Number of CBUs with HLA-A, -B, -DR(B1) typing 1b3) Number of CBUs with HLA-A, -B, -C, -DR(B1) typing 1b4) Number of CBUs with HLA-A, -B, -C, -DR(B1), -DQ(B1) typing 1b5) Number of CBUs with HLA-A, -B, -C, -DR(B1), -DQ(B1), -DP(B1) typing Adult volunteer donors/CBUs added to the database during 2017 hide questionnaire 2) Count all new donors/CBUs recruited and HLA typed during 2017 2a) Count the donors who have been registered and HLA typed during 2017 2b) Count the number of new HLA typed CBUs added during 2017

istribution of HDC products abipted to various countries for unrelated transplantation during 2017	hido questionnaire
istribution of HPC products shipped to various countries for unrelated transplantation during 2017	hide questionnaire
Number of HPC products provided by your donors shipped to international patients for unrelated transplantation	
47) HPC-Marrow (international)	
48) HPC-Apheresis (international)	
49) HPC-Cord (international)	
50) DLI (international)	
Number of HPC products provided by your donors shipped to national patients for unrelated transplantation	
51) HPC-Marrow (national)	
52) HPC-Apheresis (national)	
53) HPC-Cord (national)	
54) DLI (national)	
Number of HPC products provided by your donors shipped to the following countries for unrelated transplantation	
55) HPC-Marrow shipped to the following countries	
Own country	
Afghanistan	
Argentina	
Armenia	
Australia	
Austria	

Belarus	
Belgium	
Brazil	
Bulgaria	
Canada	
Chile	
China	
China Hong Kong	
Colombia	
Costa Rica	
Croatia	
Cuba	
Cyprus	
Czech Republic	
Denmark	
Ecuador	
Egypt	
Estonia	
Finland	
France	

Georgia	
Germany	
Greece	
Hungary	
India	
Iran	
Ireland	
Israel	
Italy	
Japan	
Jordan	
Korea	
Latvia	
Liechtenstein	
Lithuania	
Malaysia	
Mexico	
Netherlands	
New Zealand	
Norway	
Oman	

Pakistan	
Philippines	
Poland	
Portugal	
Romania	
Russia	
San Marino	
Saudi Arabia	
Serbia	
Singapore	
Slovakia	
Slovenia	
South Africa	
Spain	
Sri Lanka	
Sweden	
Switzerland	
Taiwan, province of China	
Thailand	
Turkey	

United Kingdom		
United States		
Uruguay		
Venezuela		
Vietnam		
Please specify	No.:	
	Text:	
56) HPC-Apheresis shipped to the following countries		
Own country		
Afghanistan		
Argentina		
Armenia		
Australia		
Austria		
Belarus		
Belgium		
Brazil		
Bulgaria		
Canada		
Chile		
China		

China Hong Kong	
Colombia	
Costa Rica	
Croatia	
Cuba	
Cyprus	
Czech Republic	
Denmark	
Ecuador	
Egypt	
Estonia	
Finland	
France	
Georgia	
Germany	
Greece	
Hungary	
India	
Iran	
Ireland	

Israel	
Italy	
Japan	
Jordan	
Korea	
Latvia	
Lebanon	
Liechtenstein	
Lithuania	
Malaysia	
Mexico	
Netherlands	
New Zealand	
Norway	
Oman	
Pakistan	
Philippines	
Poland	
Portugal	
Romania	
Russia	

San Marino	
Saudi Arabia	
Serbia	
Singapore	
Slovakia	
Slovenia	
South Africa	
Spain	
Sri Lanka	
Sweden	
Switzerland	
Taiwan, province of China	
Thailand	
Turkey	
United Kingdom	
United States	
Uruguay	
Venezuela	
Vietnam	
Please, specify country names No.	:
Tex	t

57) HPC-Cord shipped to the following countries	
Own country	
Afghanistan	
Argentina	
Armenia	
Australia	
Austria	
Belarus	
Belgium	
Brazil	
Bulgaria	
Canada	
Chile	
China	
China Hong Kong	
Colombia	
Costa Rica	
Croatia	
Cuba	
Cyprus	

Czech Republic	
Denmark	
Ecuador	
Egypt	
Estonia	
Finland	
France	
Georgia	
Germany	
Greece	
Hungary	
India	
Iran	
Ireland	
Israel	
Italy	
Japan	
Jordan	
Korea	
Latvia	
Liechtenstein	

Lithuania	
Malaysia	
Mexico	
Netherlands	
New Zealand	
Norway	
Oman	
Pakistan	
Philippines	
Poland	
Portugal	
Romania	
Russia	
San Marino	
Saudi Arabia	
Serbia	
Singapore	
Slovakia	
Slovenia	
South Africa	
Spain	

Sri Lanka	
Sweden	
Switzerland	
Taiwan, province of China	
Thailand	
Turkey	
United Kingdom	
United States	
Uruguay	
Venezuela	
Vietnam	
Unknown, Please specify	No.:
	Text:
58) DLI shipped to the following countries	
Own country	
Afghanistan	
Argentina	
Armenia	
Armenia Australia	
Australia	

Belgium	
Brazil	
Bulgaria	
Canada	
Chile	
China	
China Hong Kong	
Colombia	
Costa Rica	
Croatia	
Cuba	
Cyprus	
Czech Republic	
Denmark	
Egypt	
Equador	
Estonia	
Finland	
France	
Georgia	
Germany	

Greece	
Hungary	
India	
Iran	
Ireland	
Israel	
Italy	
Japan	
Jordan	
Korea	
Latvia	
Liechtenstein	
Lithuania	
Malaysia	
Mexico	
Netherlands	
New Zealand	
Norway	
Oman	
Pakistan	

Philippines	
Poland	
Portugal	
Romania	
Russia	
San Marino	
Saudi Arabia	
Serbia	
Singapore	
Slovakia	
Slovenia	
South Africa	
Spain	
Srilanka	
Sweden	
Switzerland	
Taiwan	
Thailand	
Turkey	
United Kingdom	
United States	
Uruguay	
Venezuela	
Vietnam	
Unknown	

## Appendix 3: Global Trends 1997 - 2017

The Global Trends movie with a summary of all the achievements over the past 20 years can be viewed via the following website:

https://drive.google.com/file/d/1RsoIj-JMkBMBfDAmvXL29QinmB 9Nsnf/view

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8. Images of countries by This Photo by Unknown Author is licensed under CC BY-SA (Word Office)