The Fates of Endocrine Disruptors in Consumer Products: Bisphenol A

Prepared by Dr. Matthew A. DeNardo for the One Health One Planet Conference on March 8, 2018 at Phipps Conservatory



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THE INSTITUTE FOR **GREEN SCIENCE** CARNEGIE MELLON UNIVERSITY



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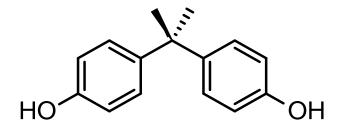
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THE INSTITUTE FOR **GREEN SCIENCE** CARNEGIE MELLON UNIVERSITY A multidisciplinary investigation of the technical and environmental performances of TAML/peroxide elimination of Bisphenol A compounds from water. *Green Chemistry* doi:10.1039/C7GC01415E

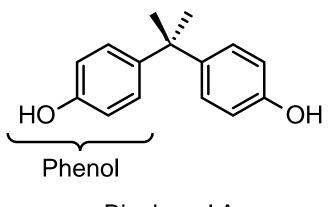


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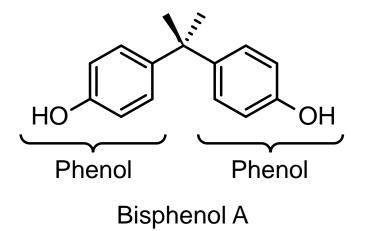
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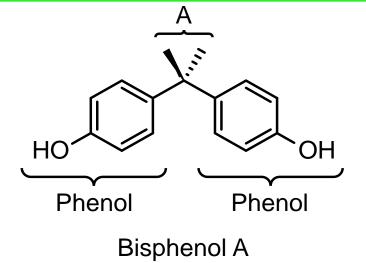


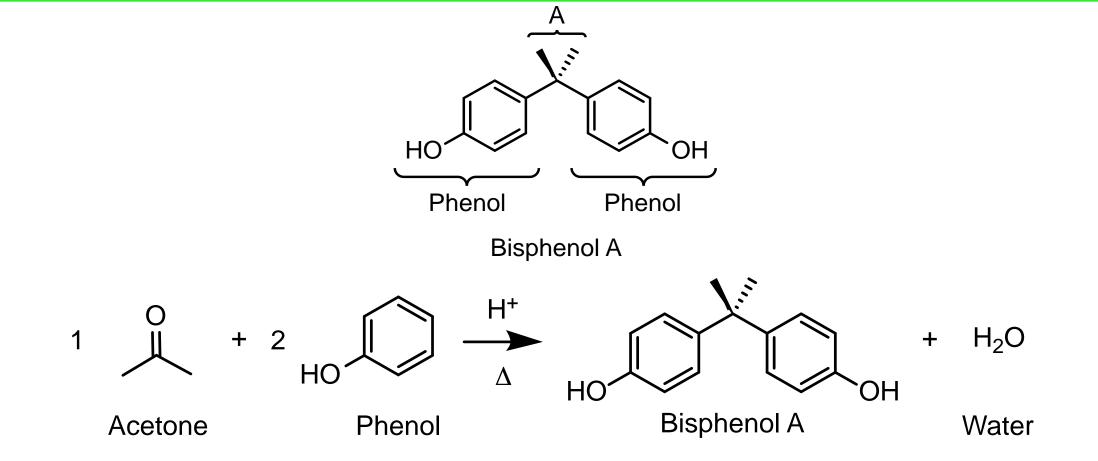
Bisphenol A

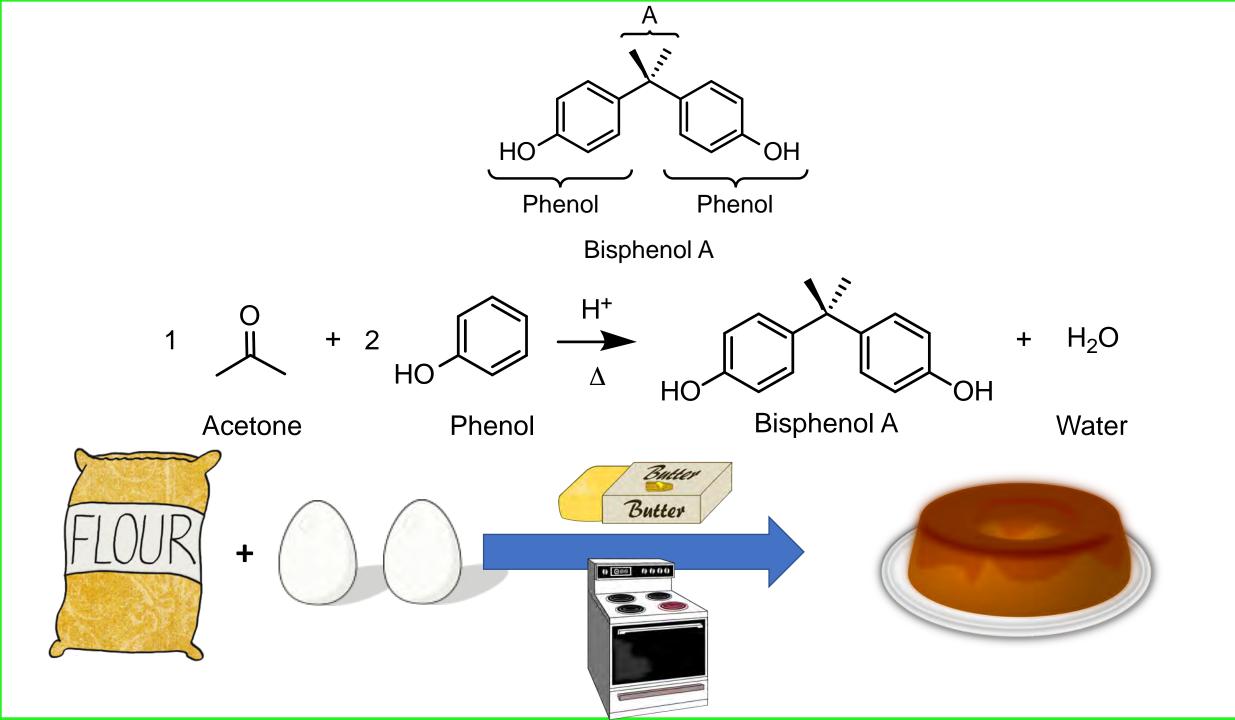


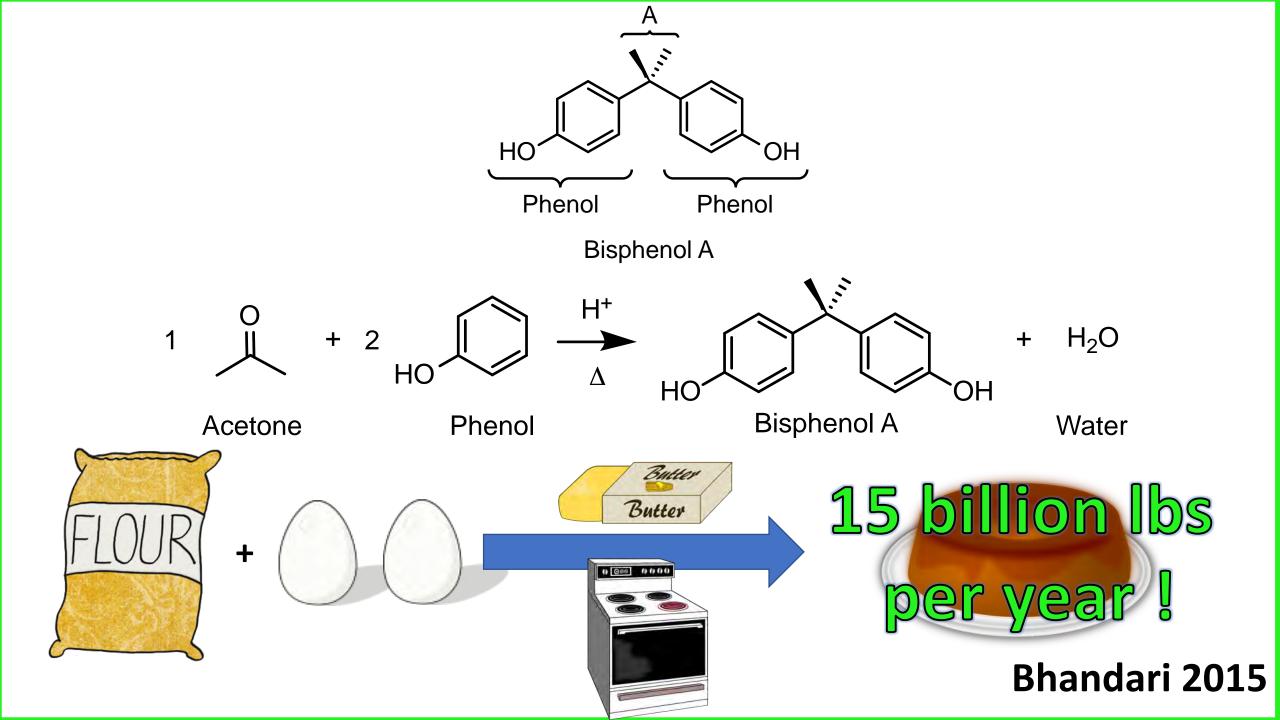
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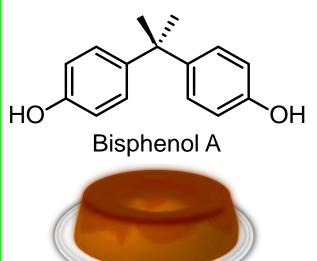


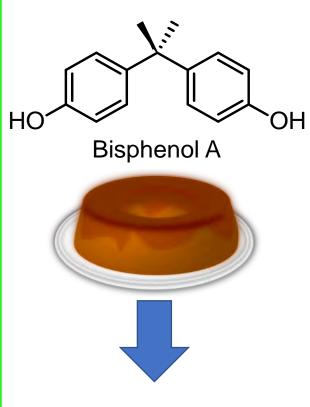


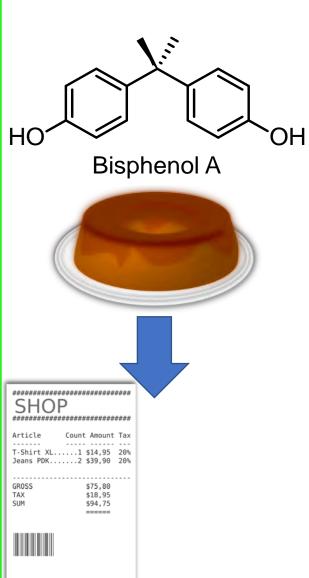


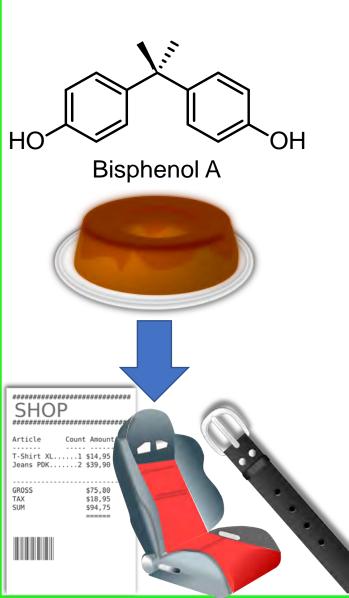


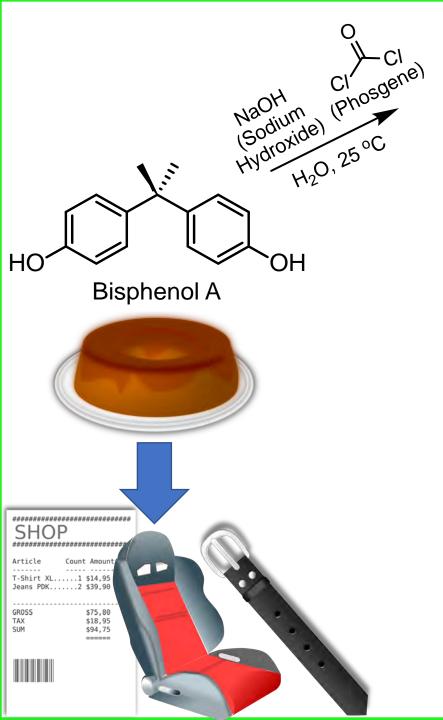


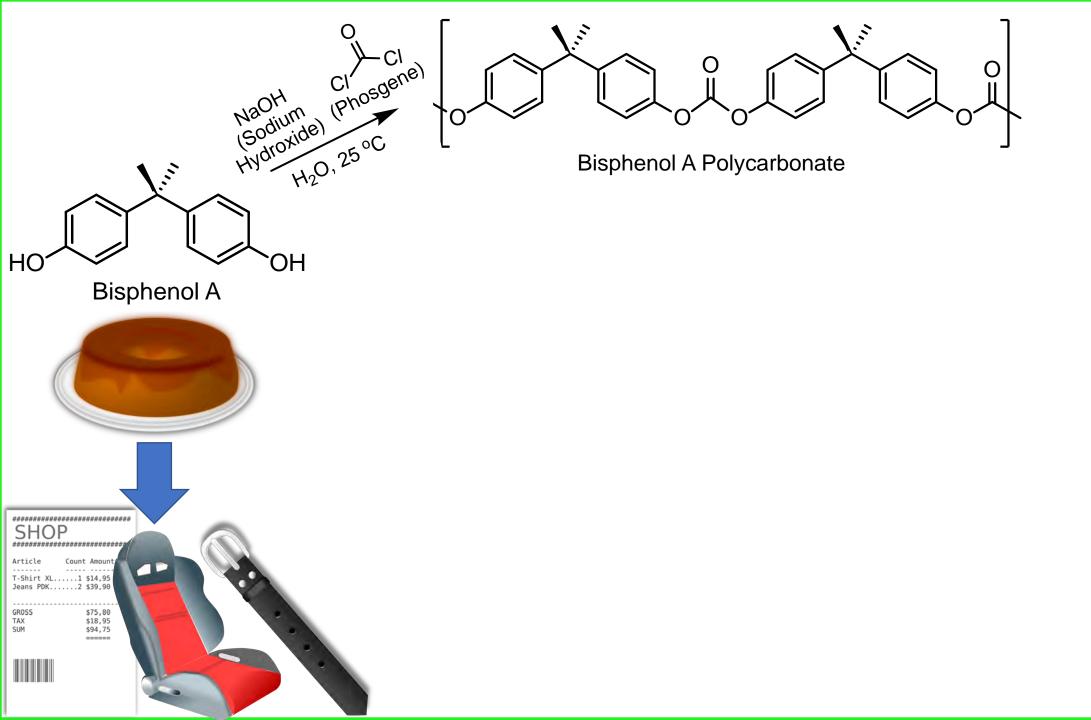


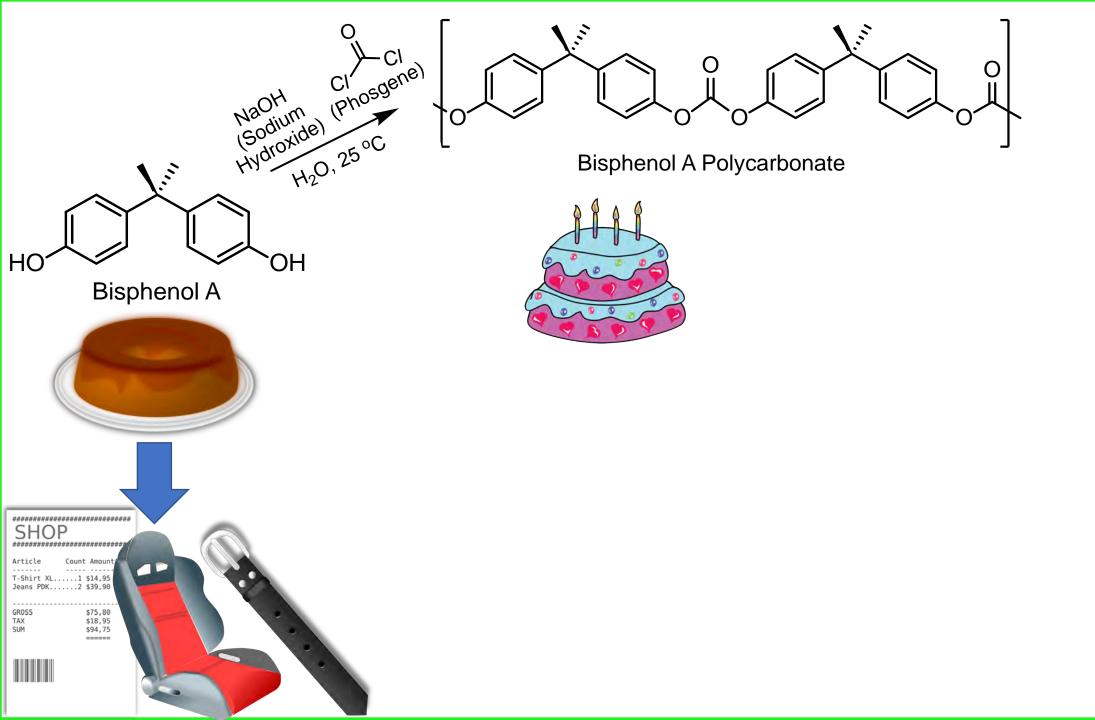


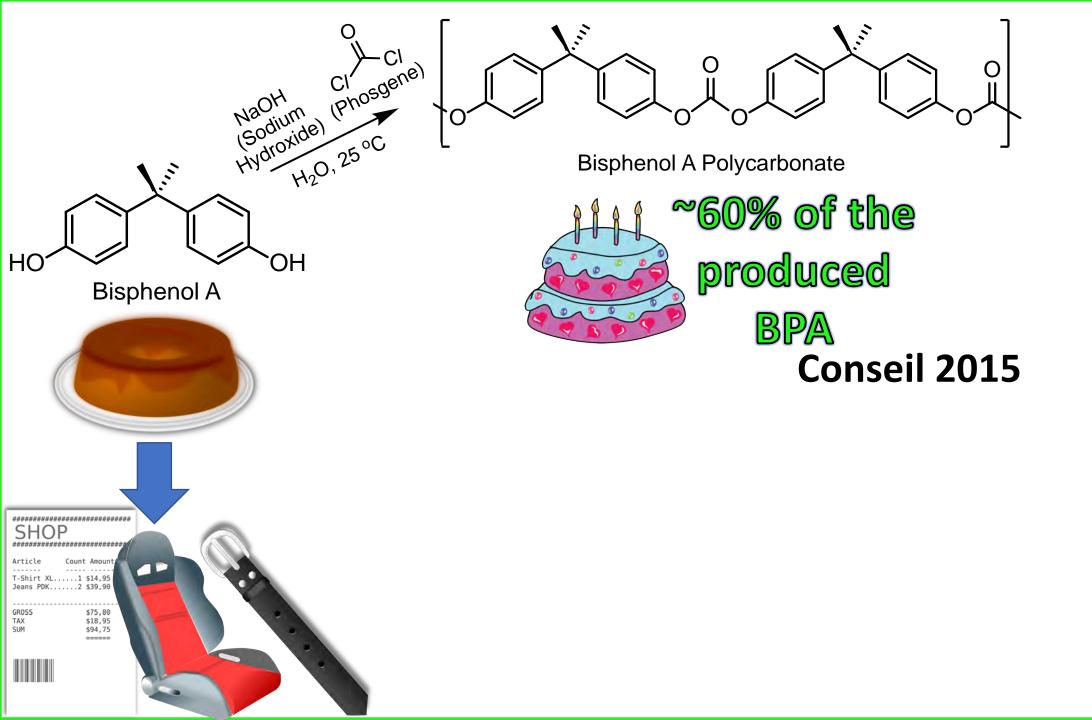


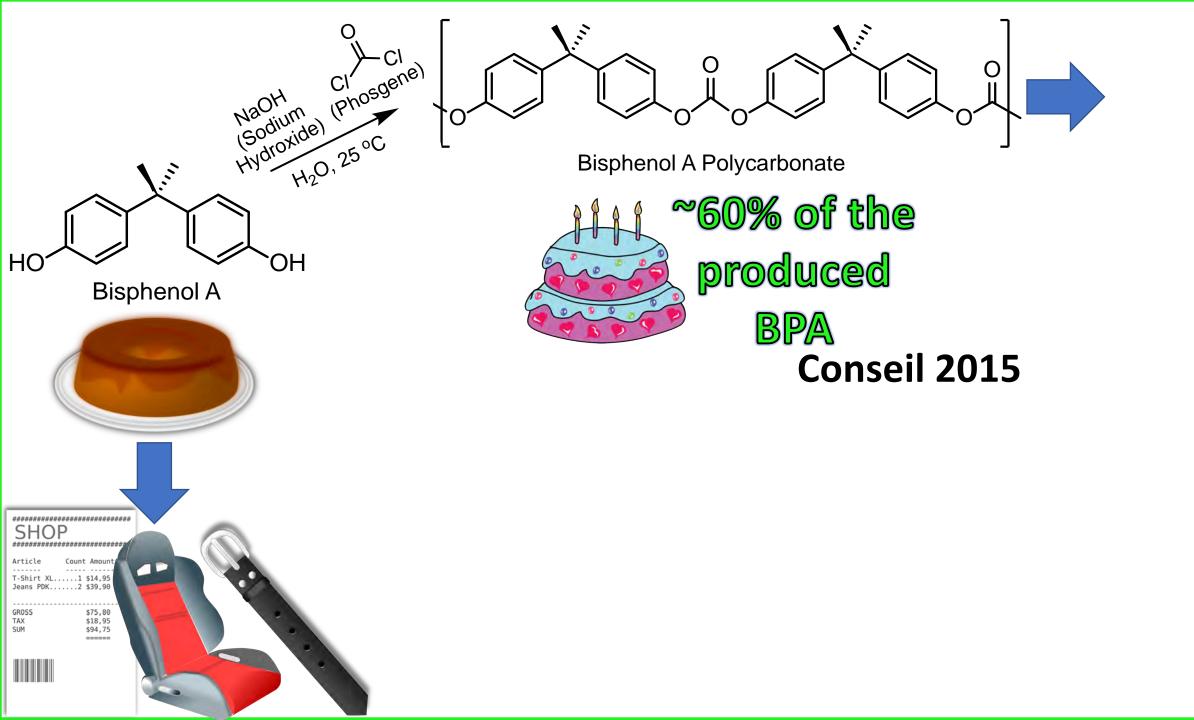


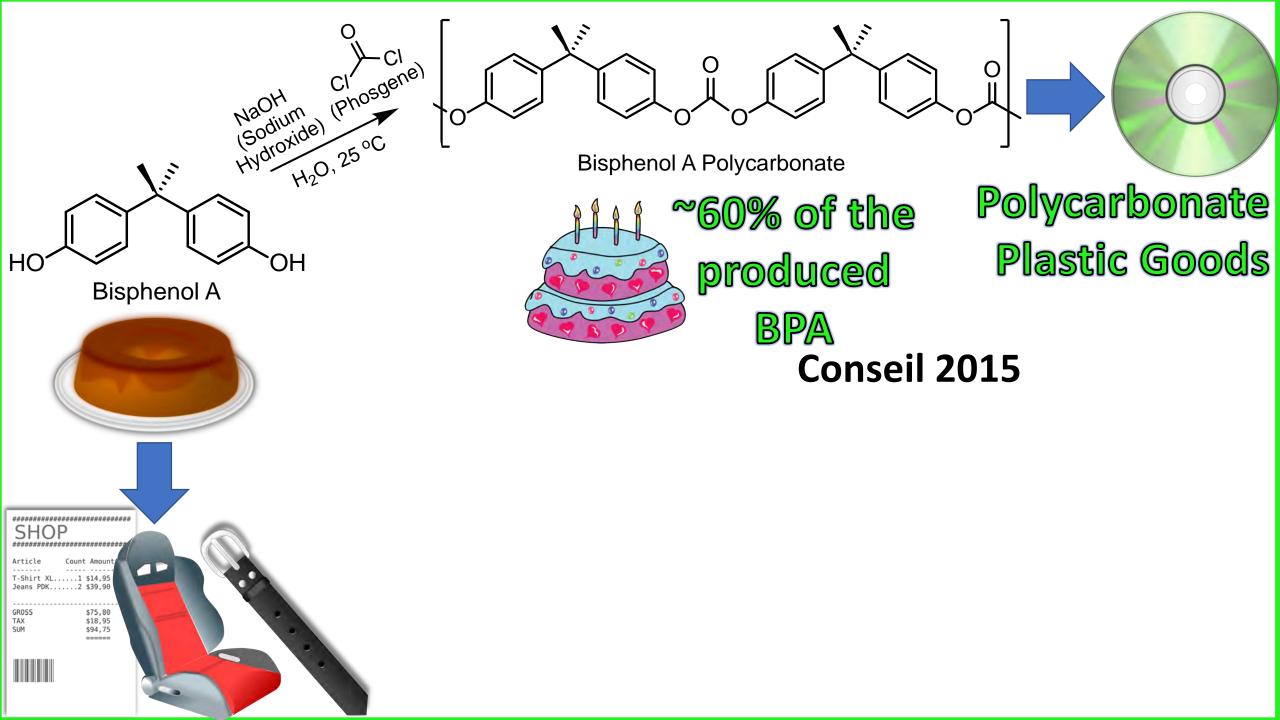


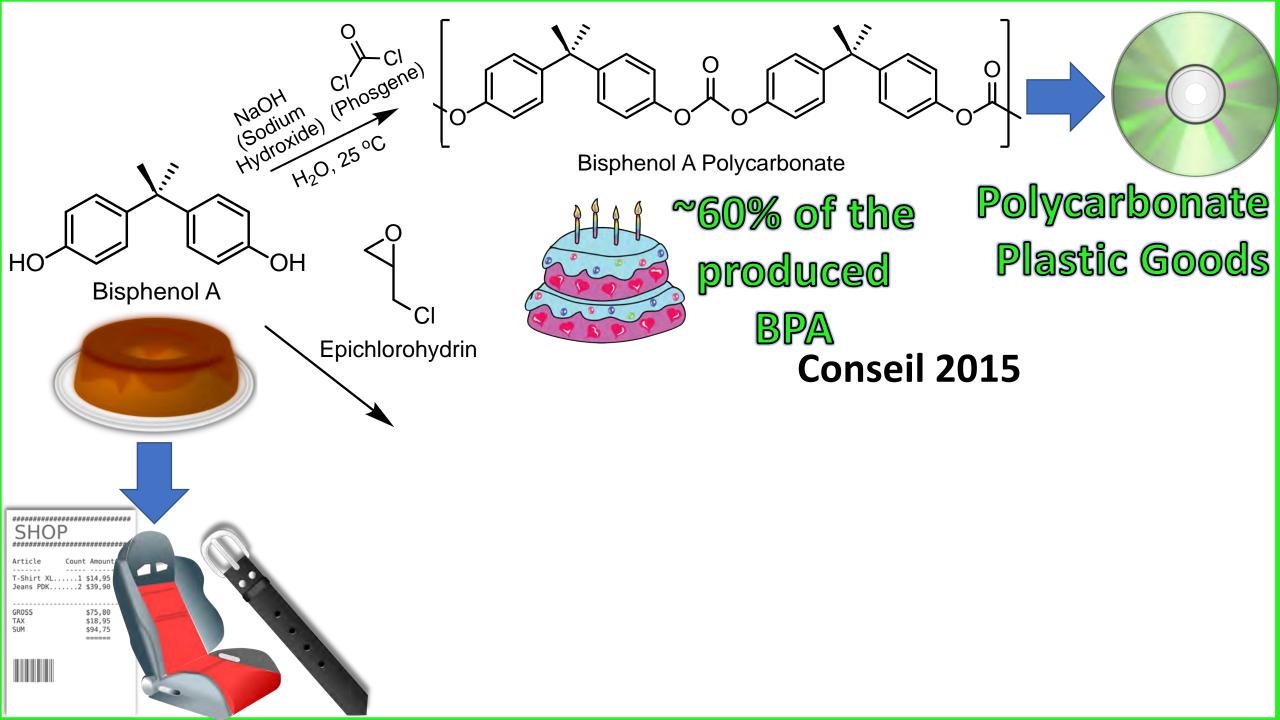


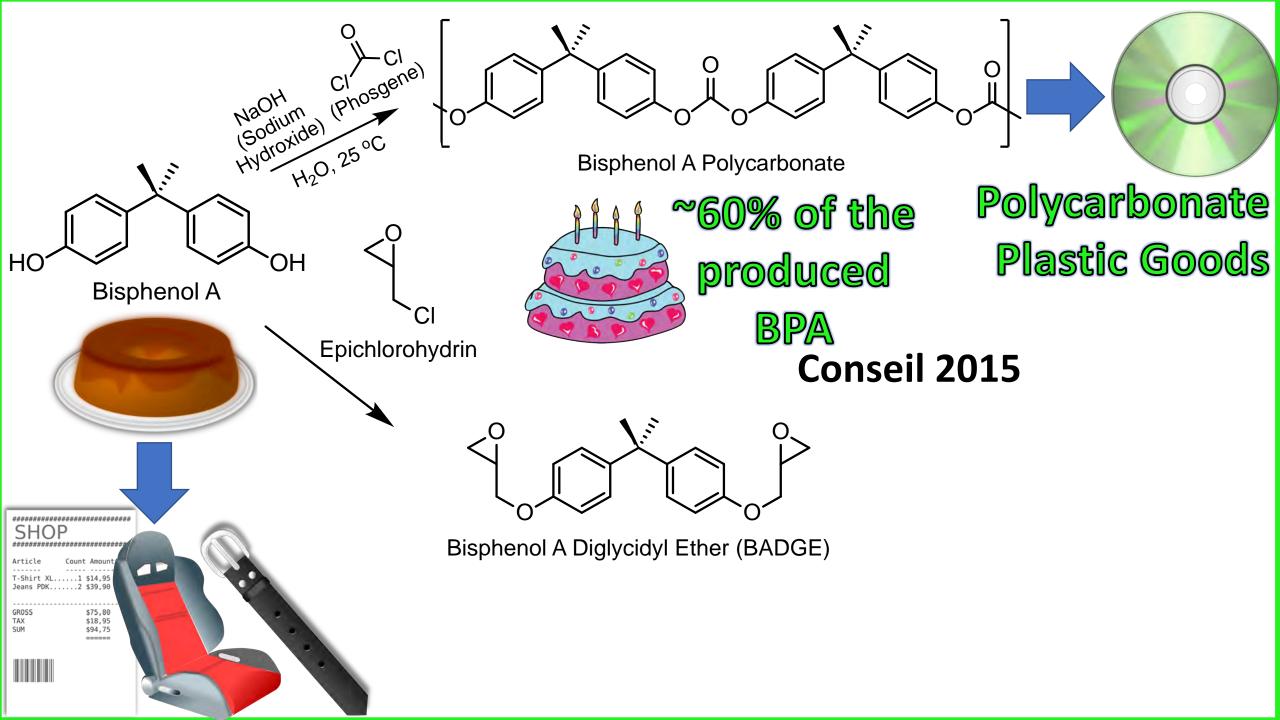


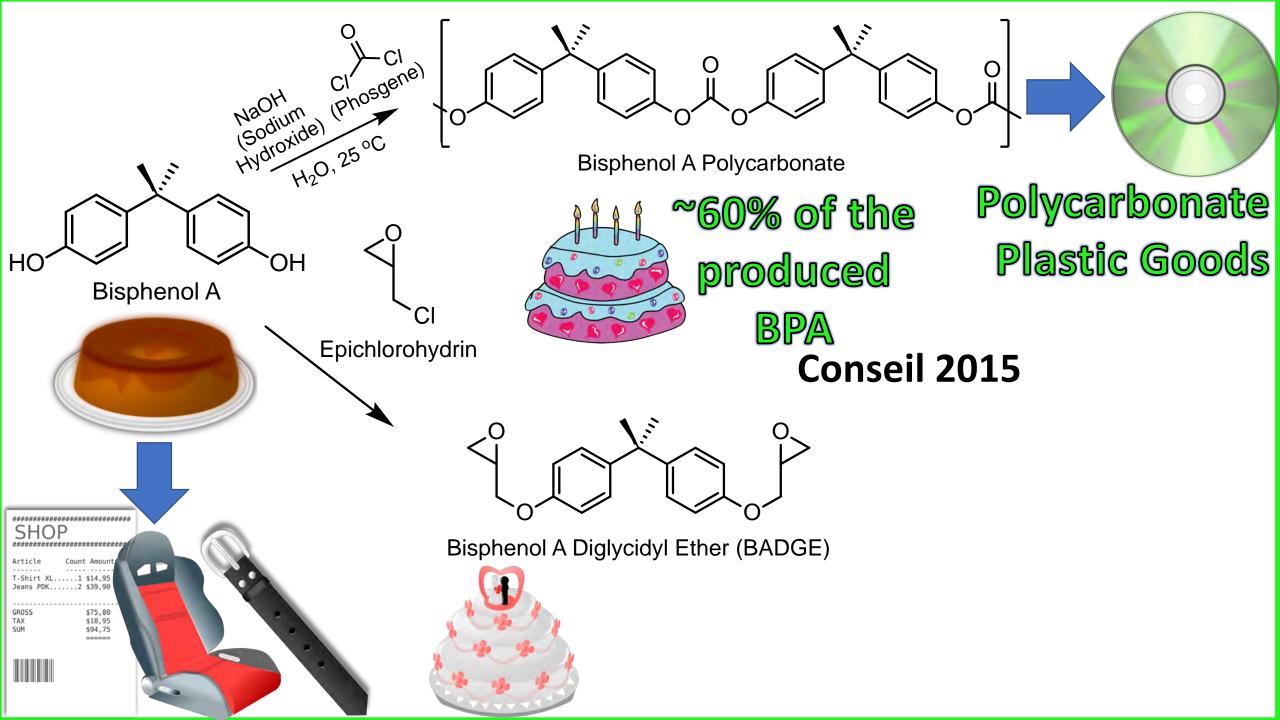


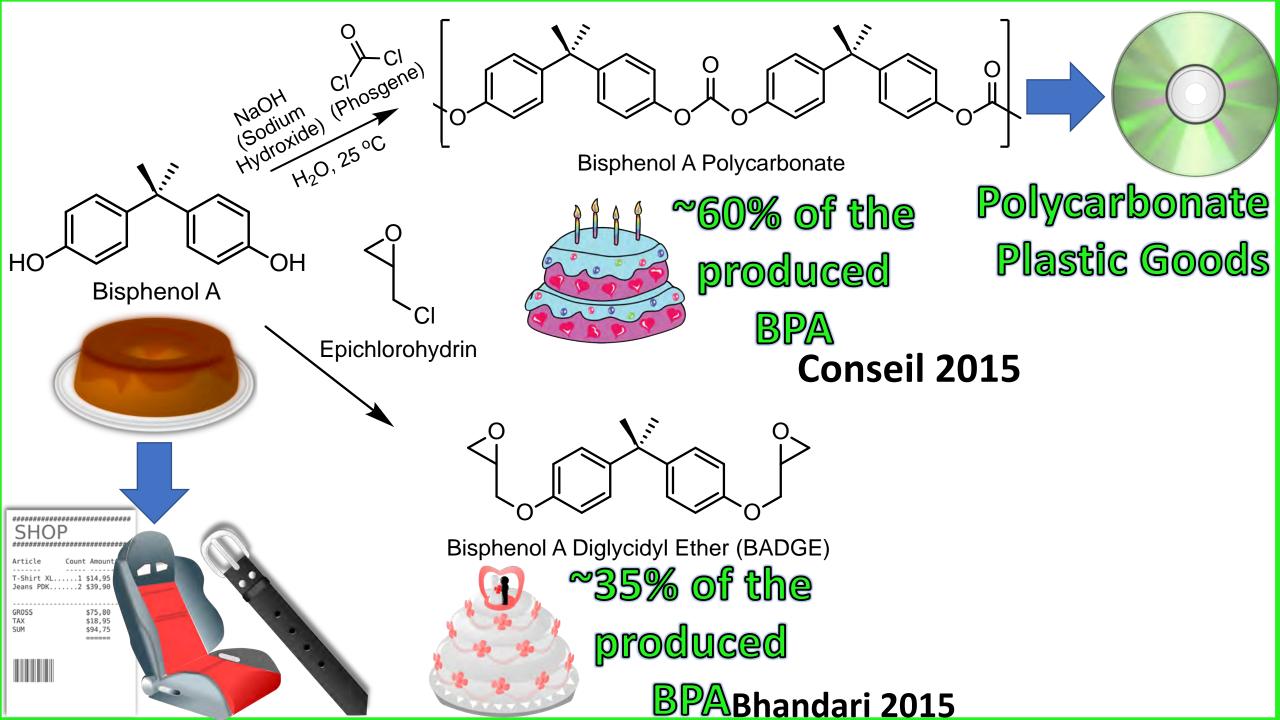


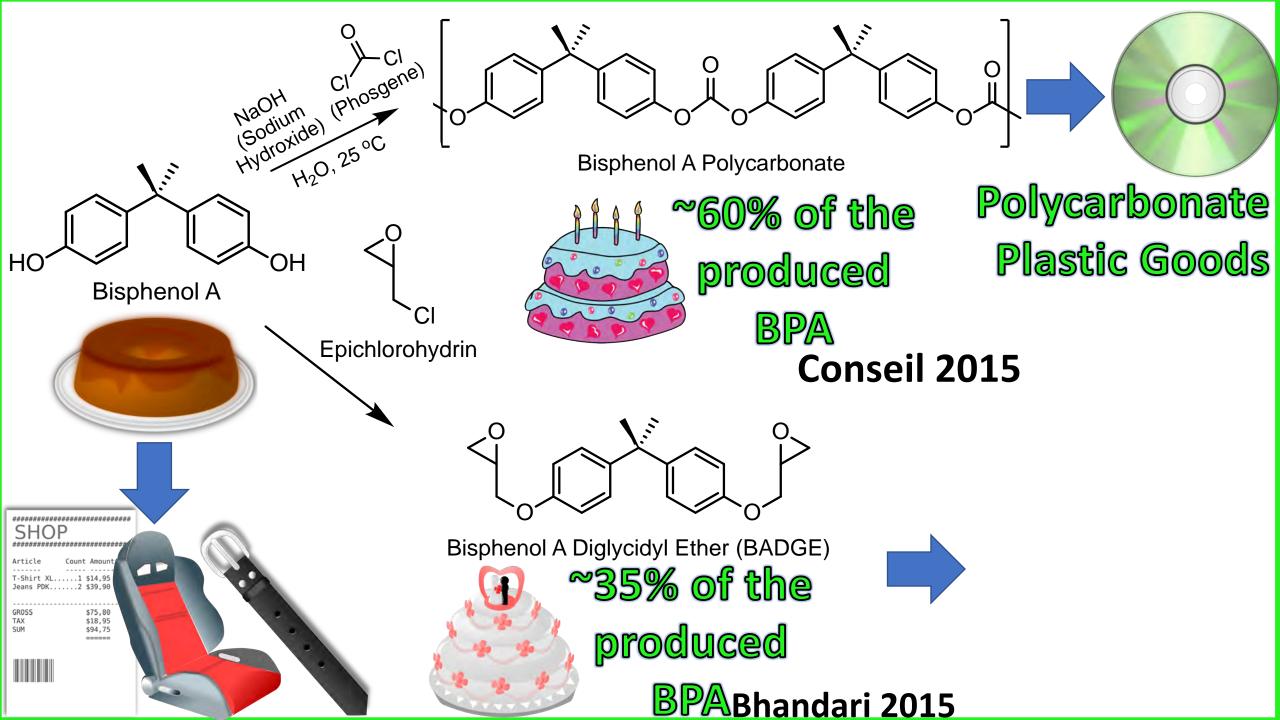


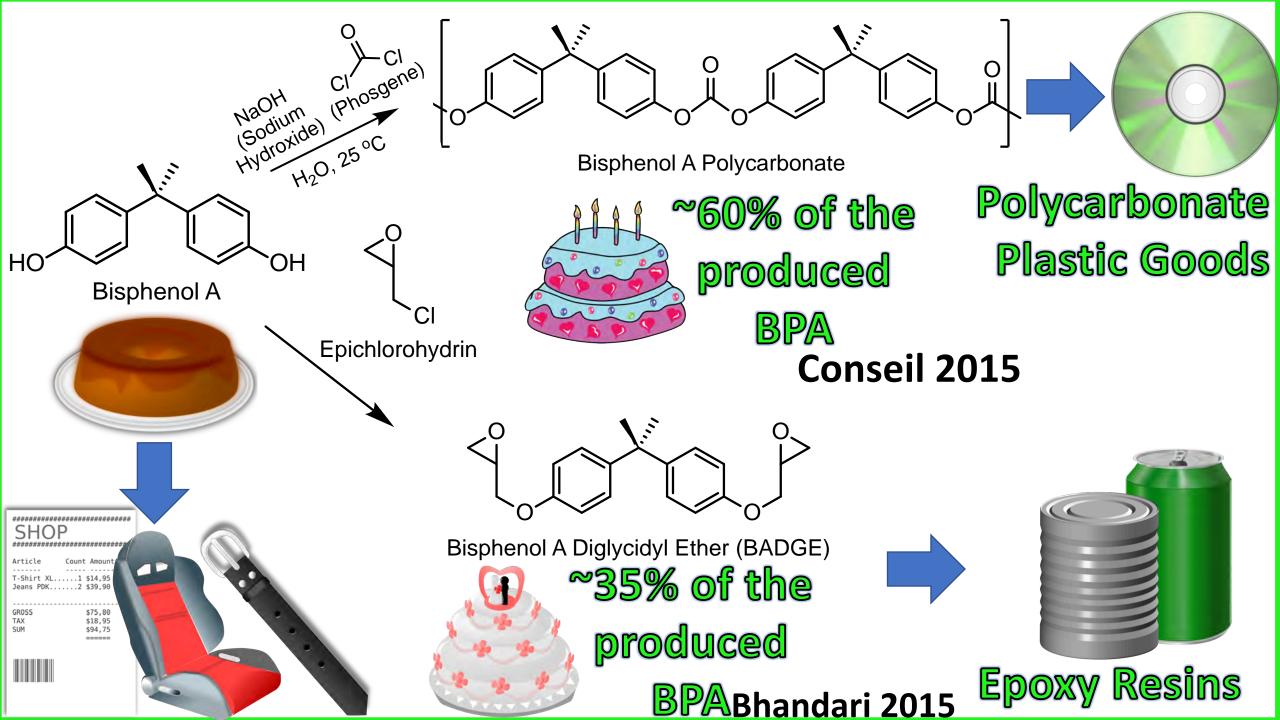


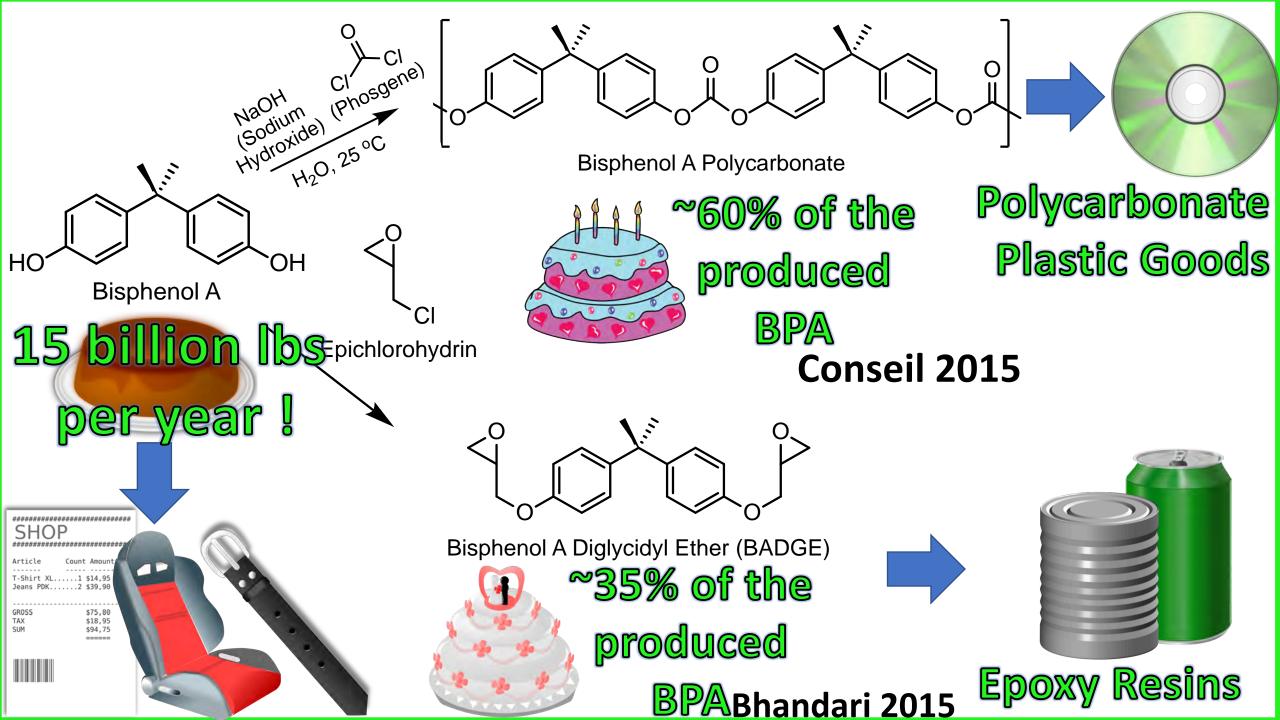












What Are the Consequences of The Mass Production of BPA?



















Contamination of Air



- Contamination of Air
- Contamination of Water

What Are the Concentrations of BPA in Industrial Effluents?

Effluent	Country	Year	Range (µg L ⁻¹)	Mean (µg L ⁻¹)	Ref.
Chemical Industry	Austria	2000	2.5–50	18	Furhacker 2000
Paper Production and Recycling	Austria	2000	28–72	41	Furhacker 2000
Paper Production and Recycling	Japan	2002	0.2–370	59	Fukazawa 2002
Plastics Manufacturing and Recycling	Nigeria	2015	108– 163	130	Mackinwa 2015

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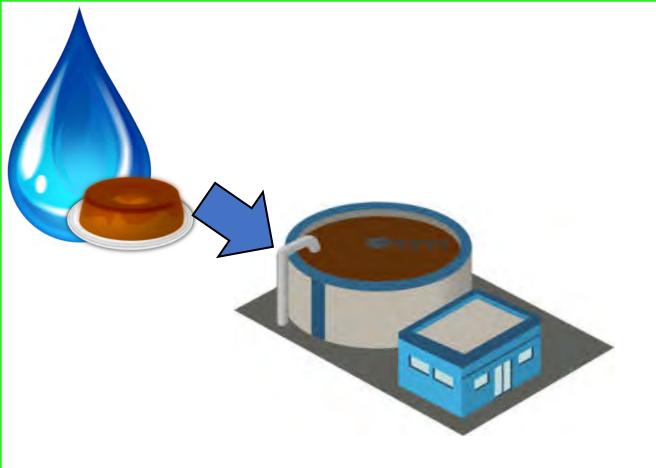
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What Are the Concentrations of BPA in Municipal Influents?

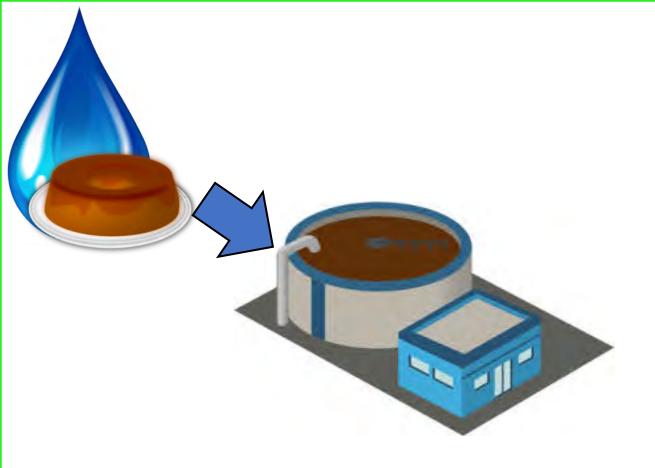
Country	Year	Range (µg L ⁻¹)	Mean (µg L ⁻¹)	Ref.
Austria	2000	10–37	21	Furhacker 2000
Canada	2004	0.16-28.1		Lee 2004
Germany	2008	<0.02-12.2	3.67	Hohne 2008

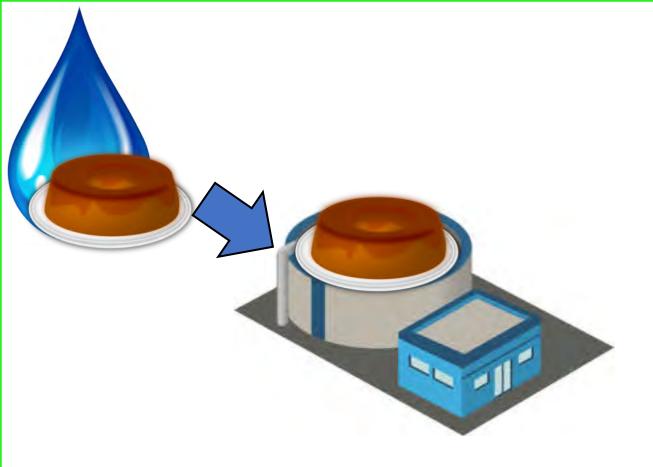
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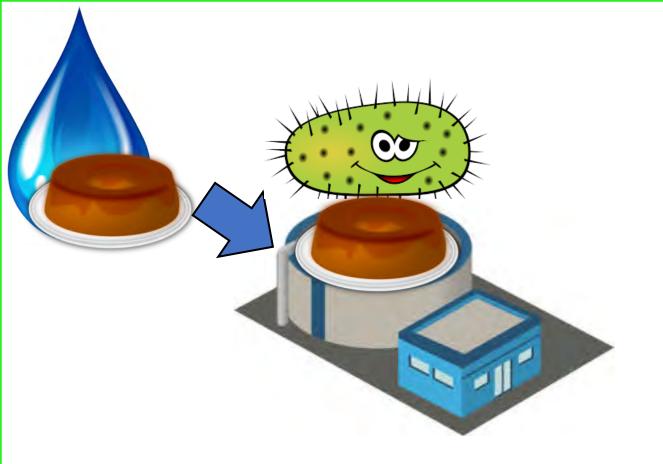
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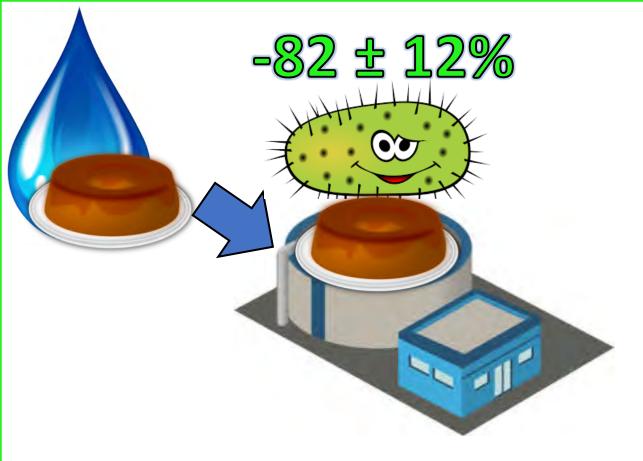
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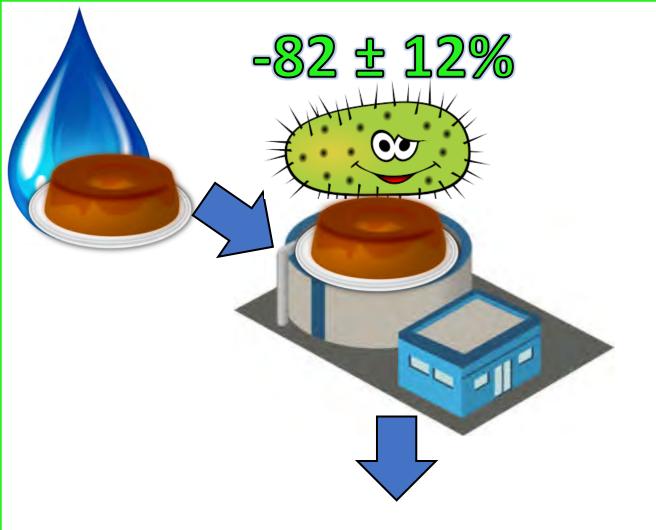
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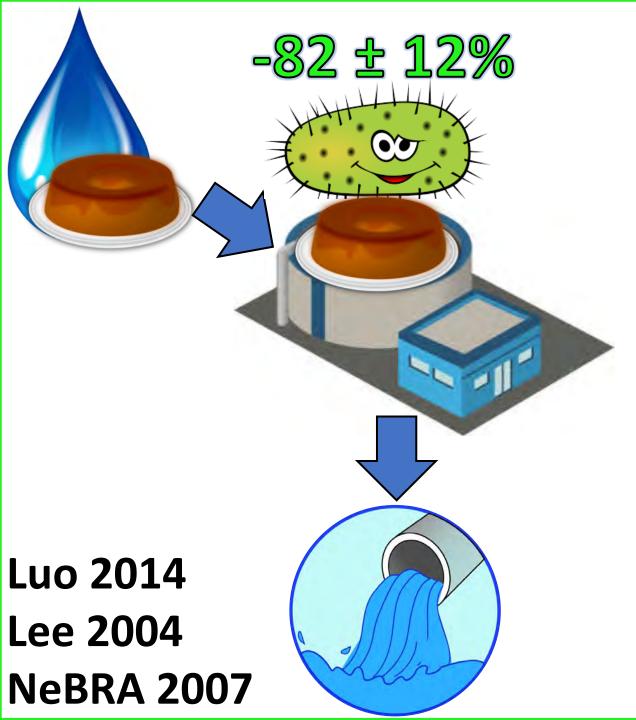


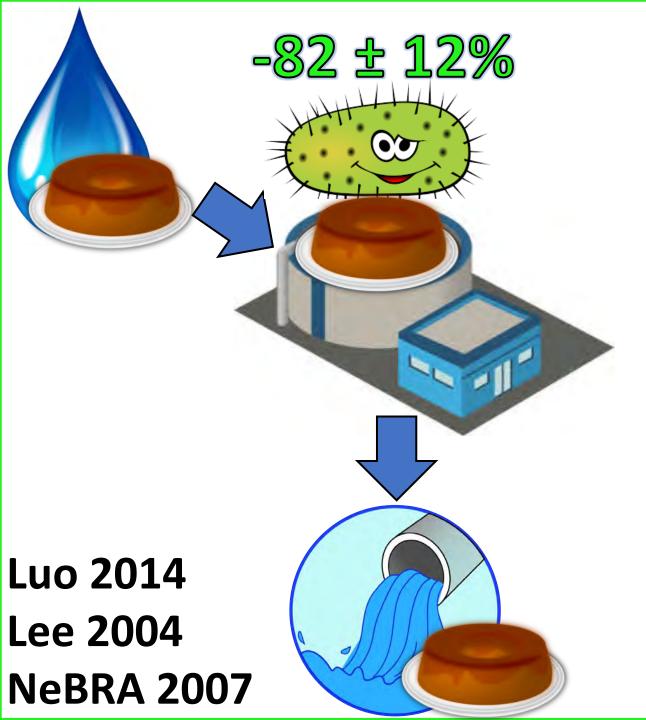












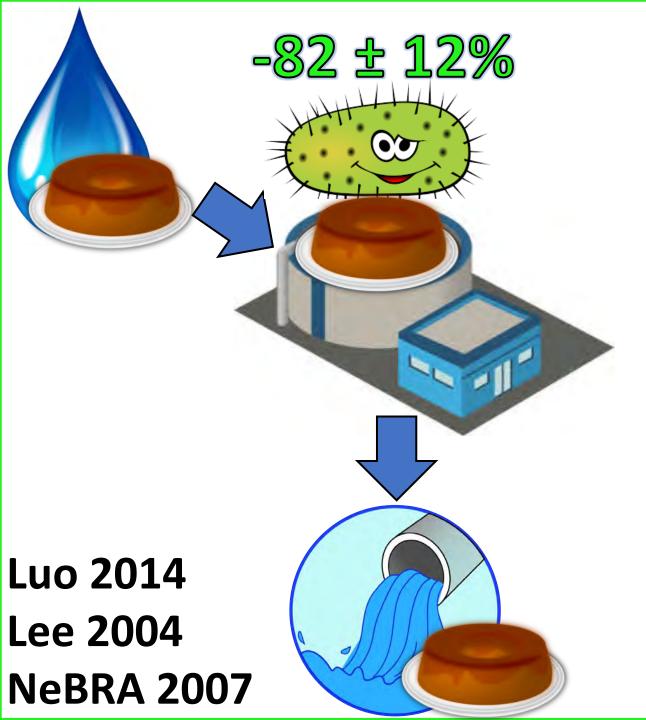
What Are the Concentrations of BPA in WWTP Effluents?

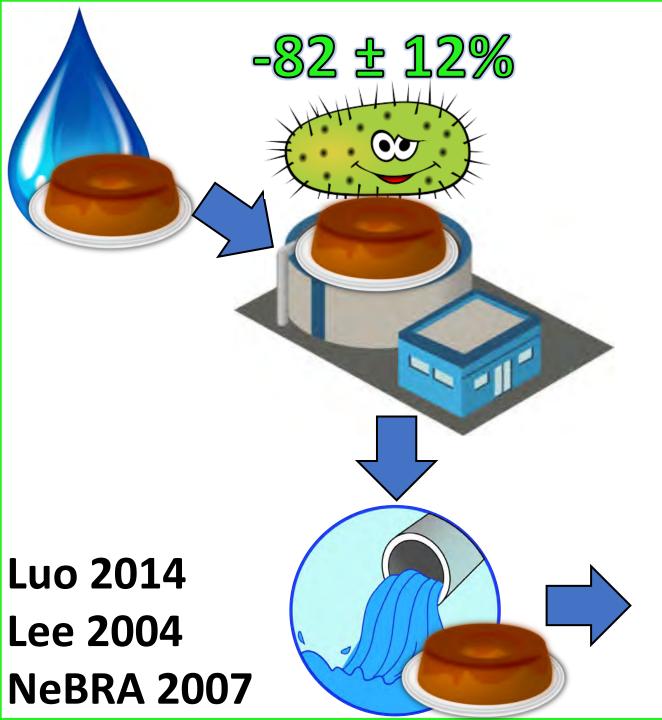
Country	Year	Range (µg L ⁻¹)	Mean (μg L ⁻¹)	Ref.
Austria	2000	<0.5-2.5	1.5	Furhacker
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Canada	2004	0.01-17.3		Lee 2004
EU	2008	3.13–45		EU 2008
EU	2008	<0.02-7.6	0.52	Hohne
US	1999	<0.01-2.7		Barber 1999

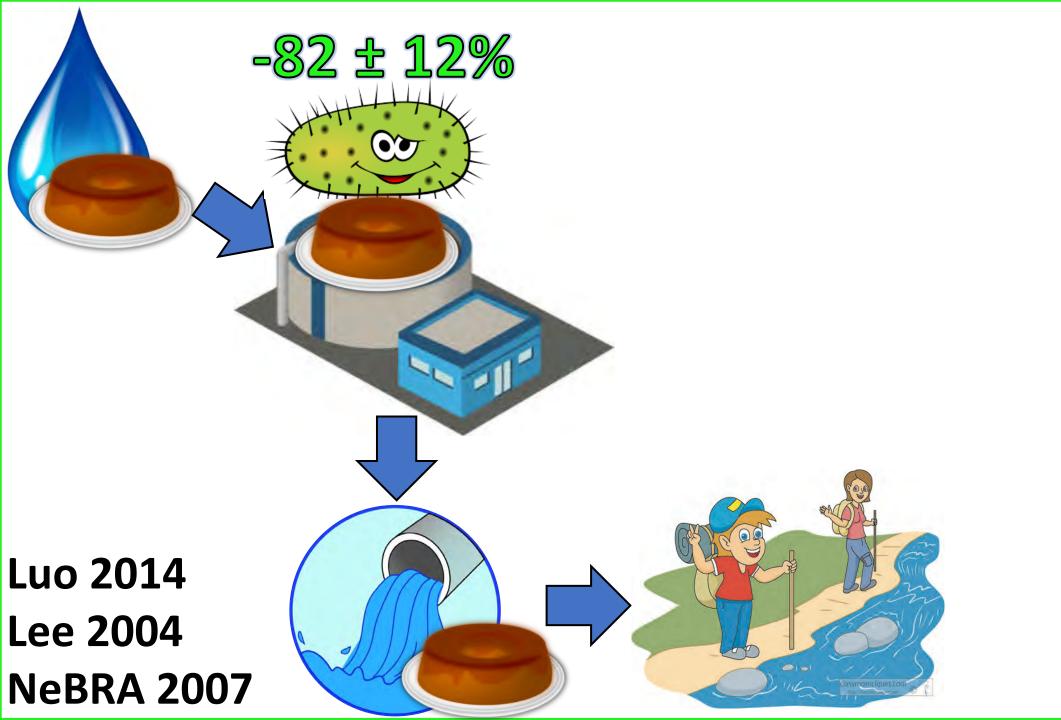
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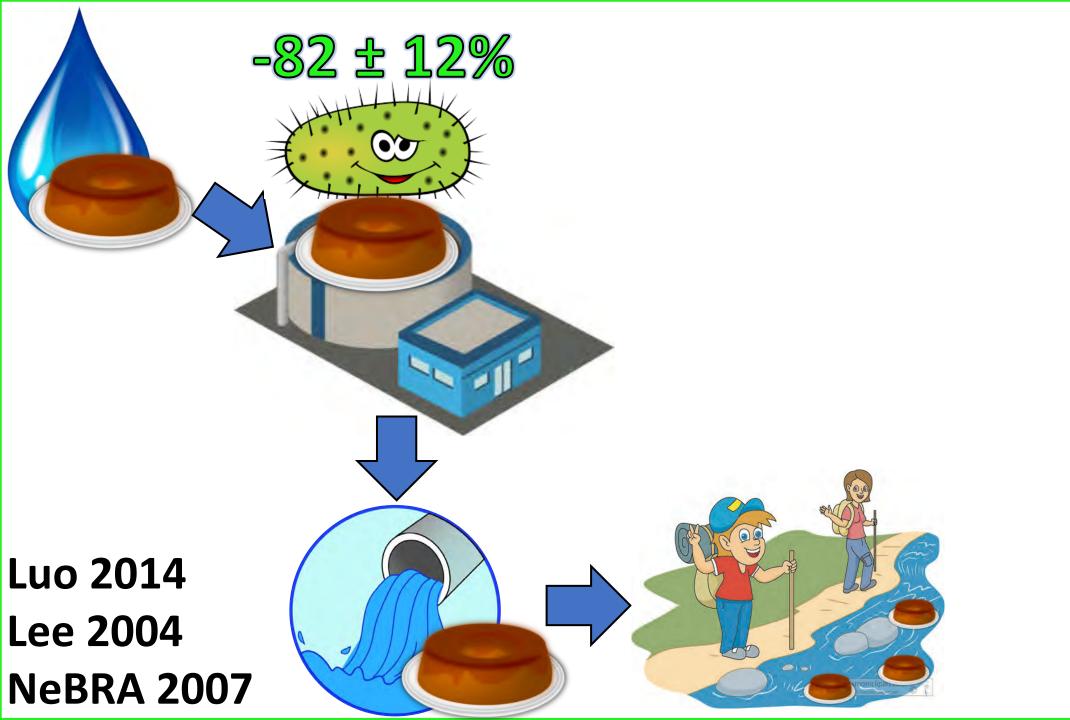
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What Are the Concentrations of BPA in Surface Waters?

Surface Water Concentrations of BPA				
Country	Watershed	Max (µg L ⁻¹)	Median / Avg / Range (µg L ⁻¹)	Ref.
US	Multiple	12	Med: 0.14	Kolpin 2002
Brazil	Atibaia	13	Avg: 4.6	Montagner
China	Dongguan	56	Avg: 6.5	Tang 2012
Japan	Nagara	22.2	Avg: 4.8	Funakoshi
Portugal	Multiple		Range: 0.07–4	Azevedo 2001

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What Are the Effects of Fish Exposures to These BPA Concentrations?



- Metcalfe 2001
 - Saili 2012
 - Lee 2012
 - Wang 2010
 - **Chung 2011**
 - Lam 2011
 - Gibert 2011

• Alteration of Gene Expression

- Metcalfe 2001
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- Alteration of Gene Expression
- Reduction of Heart Rate

Metcalfe 2001 Saili 2012 Lee 2012 Wang 2010 Chung 2011

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- Alteration of Gene Expression
- Reduction of Heart Rate
- Decreased Eye Pigmentation Density

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- Testis-Ova in Males

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- Reduction of Heart Rate
- Decreased Eye Pigmentation Density
- Accelerated Development
- Delayed Hatching of Embryos
- Hyperactivity in Larva
- Testis-Ova in Males
- Learning Deficits in Adult Males



Adult Fish

Crain 2007 Sohoni 2001 Lahnsteiner 2005 Hatef 2012 Hayashi 2007 Villenueve 2012 Liu 2012 Qin 2013

Crain 2007 Sohoni 2001 Lahnsteiner 2005 Hatef 2012 Hayashi 2007 Villenueve 2012

- Liu 2012
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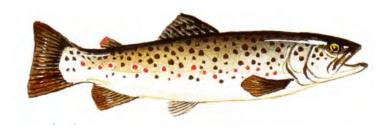
Adult Fish

 Alteration of gene expression including stimulation of vitellogenin synthesis in males (vtg is a precursor of egg yolk protein and is a biomarker for exposure to oestrogens) Crain 2007 Sohoni 2001 Lahnsteiner 2005 Hatef 2012

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Adult Fish

- Alteration of gene expression including stimulation of vitellogenin synthesis in males (vtg is a precursor of egg yolk protein and is a biomarker for exposure to oestrogens)
- Reduction in male sperm quality

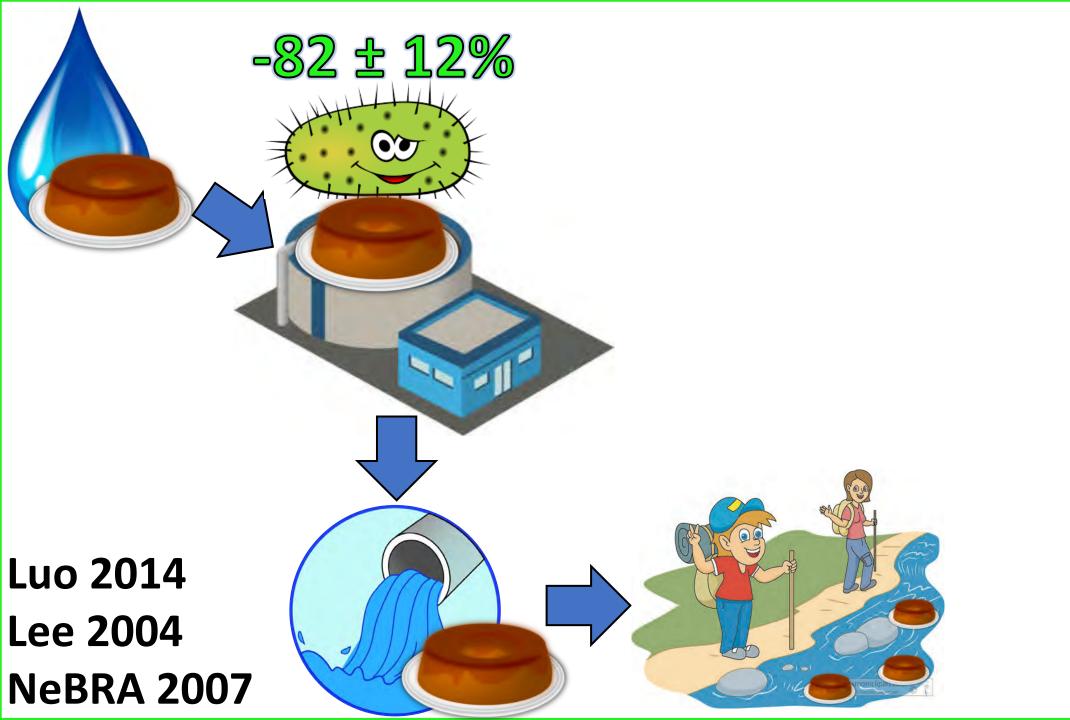
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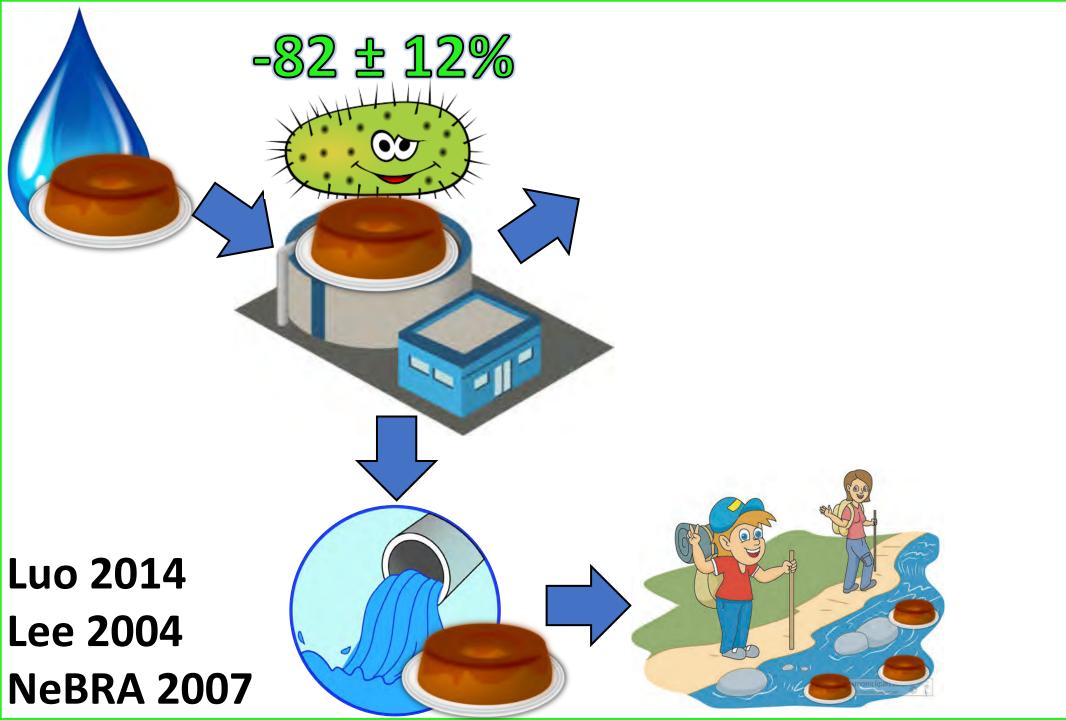
- Villenueve 2012
- Liu 2012
- Qin 2013

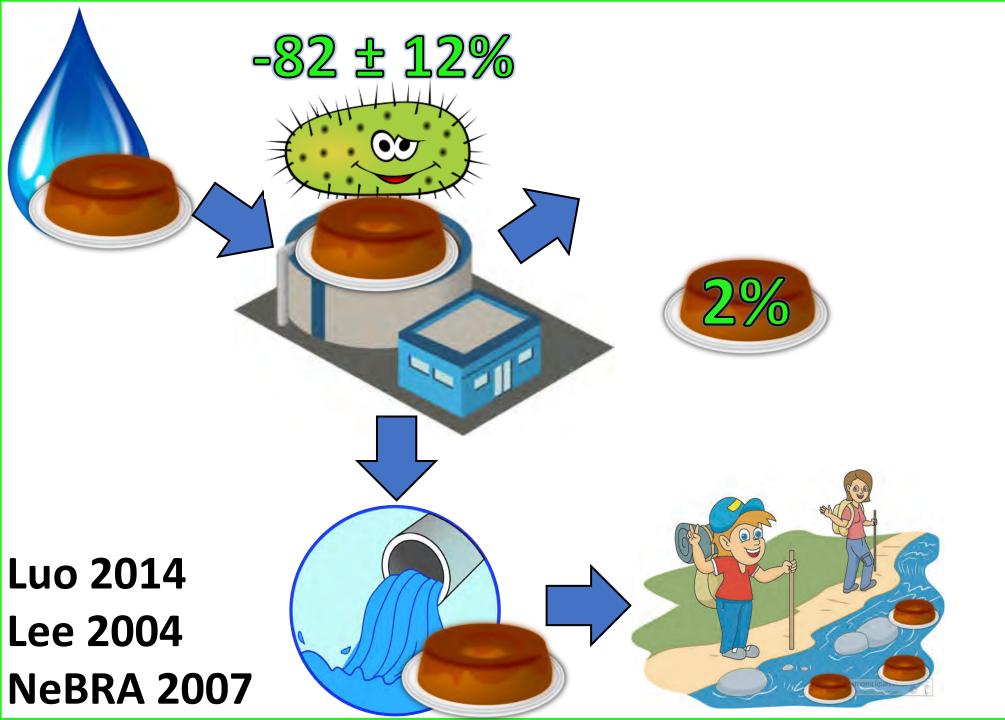


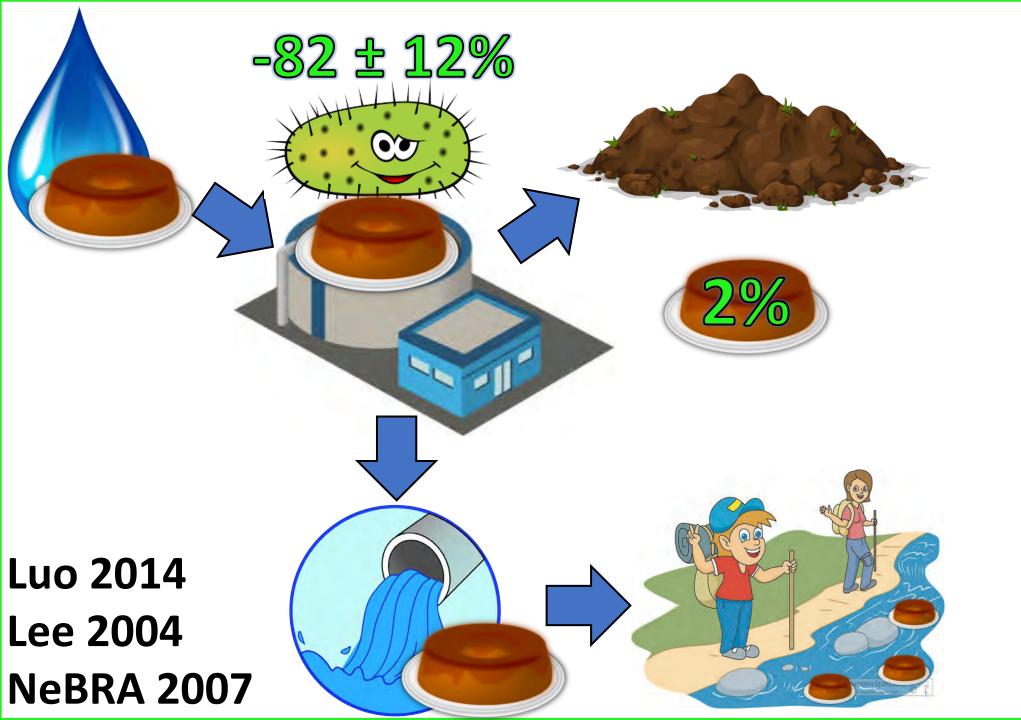
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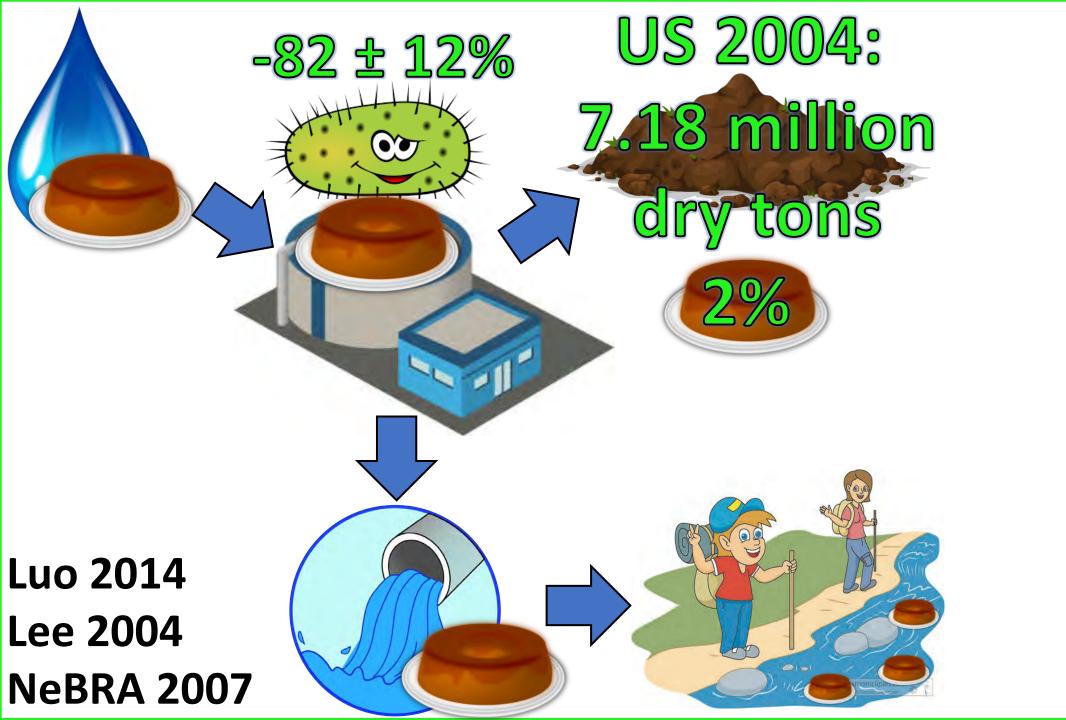
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- Delayed or no ovulation in females

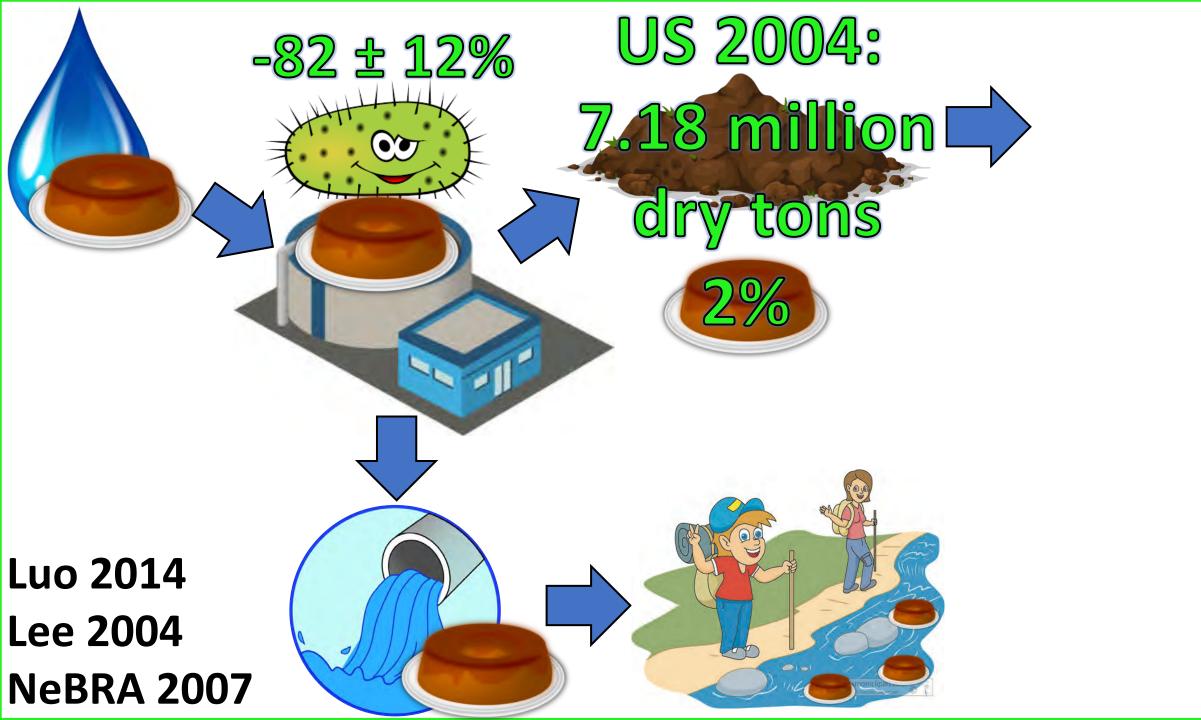


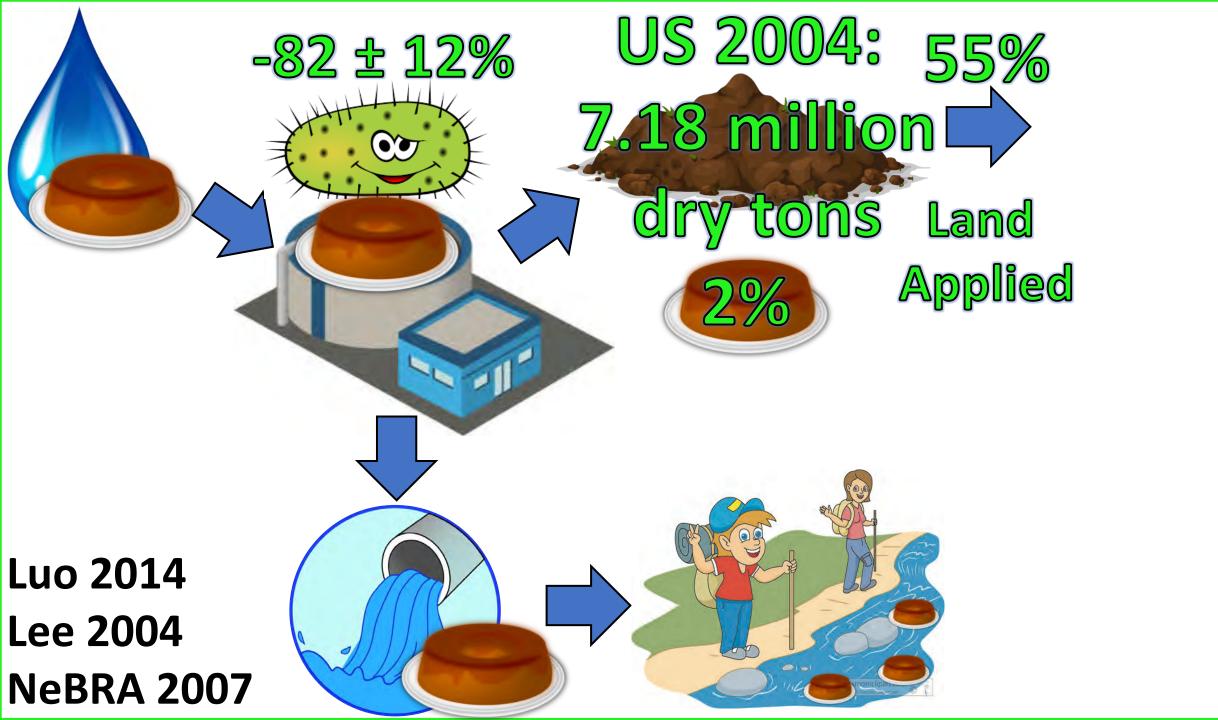


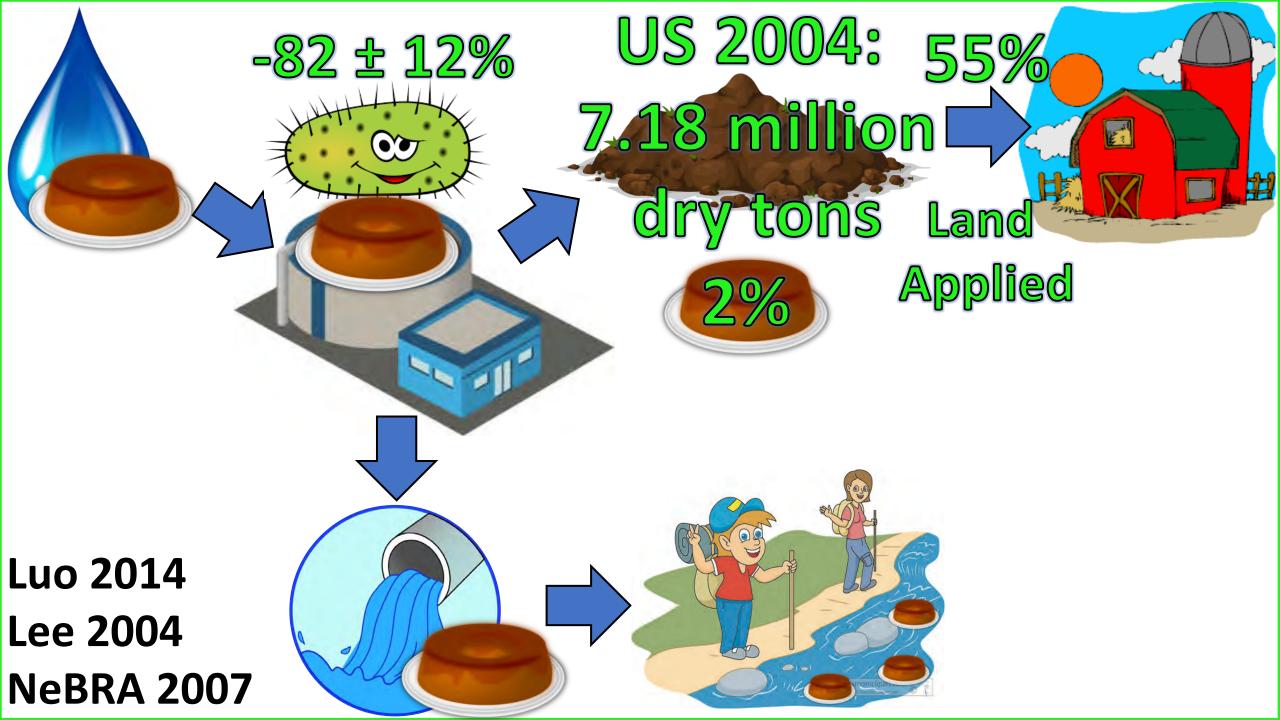


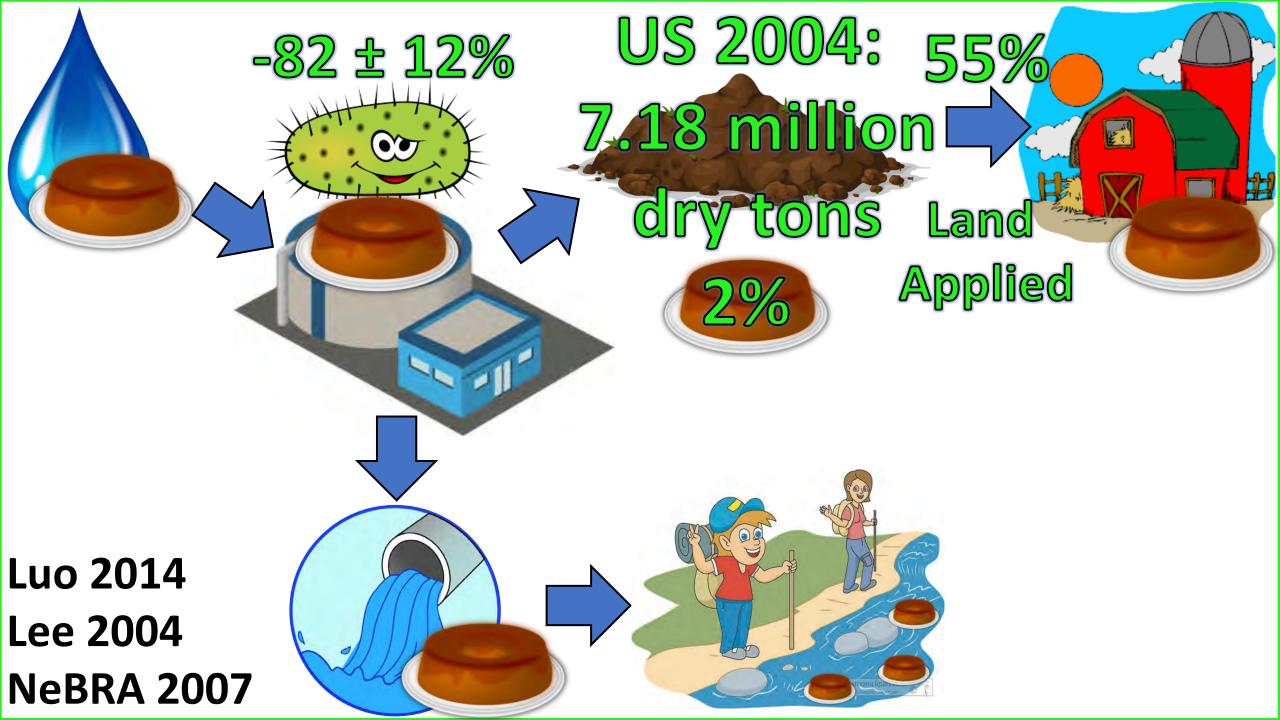


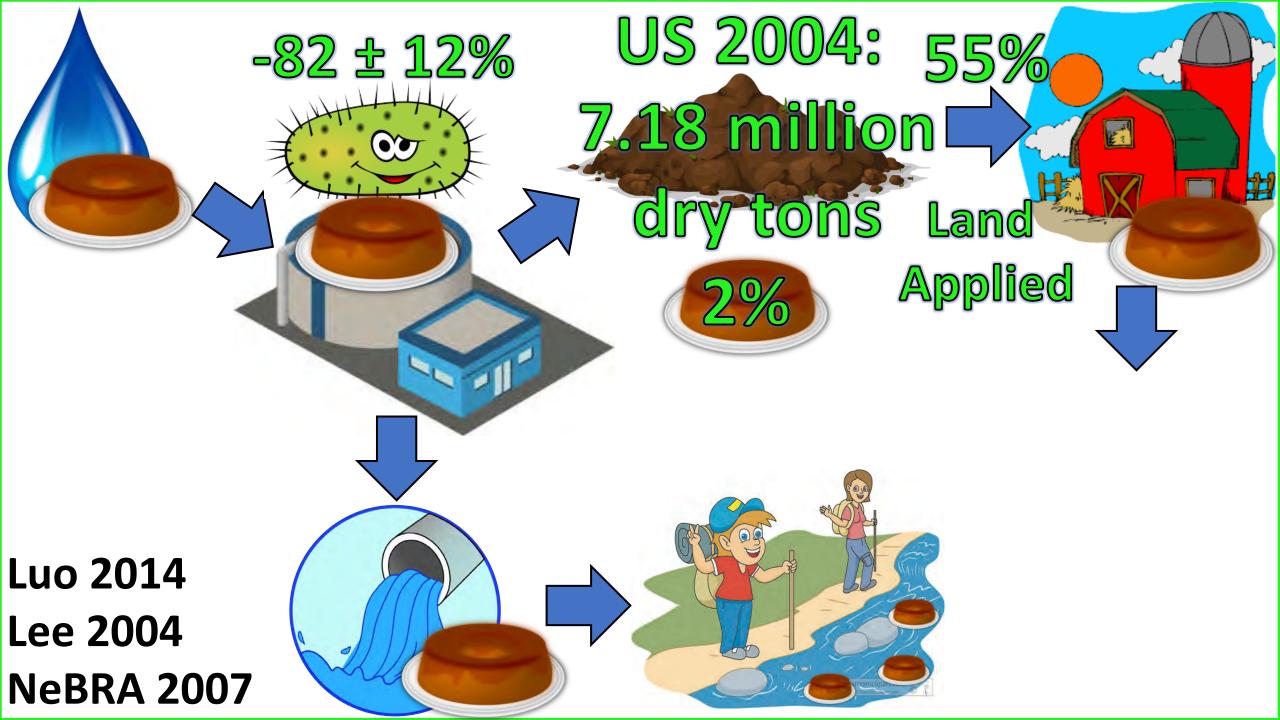


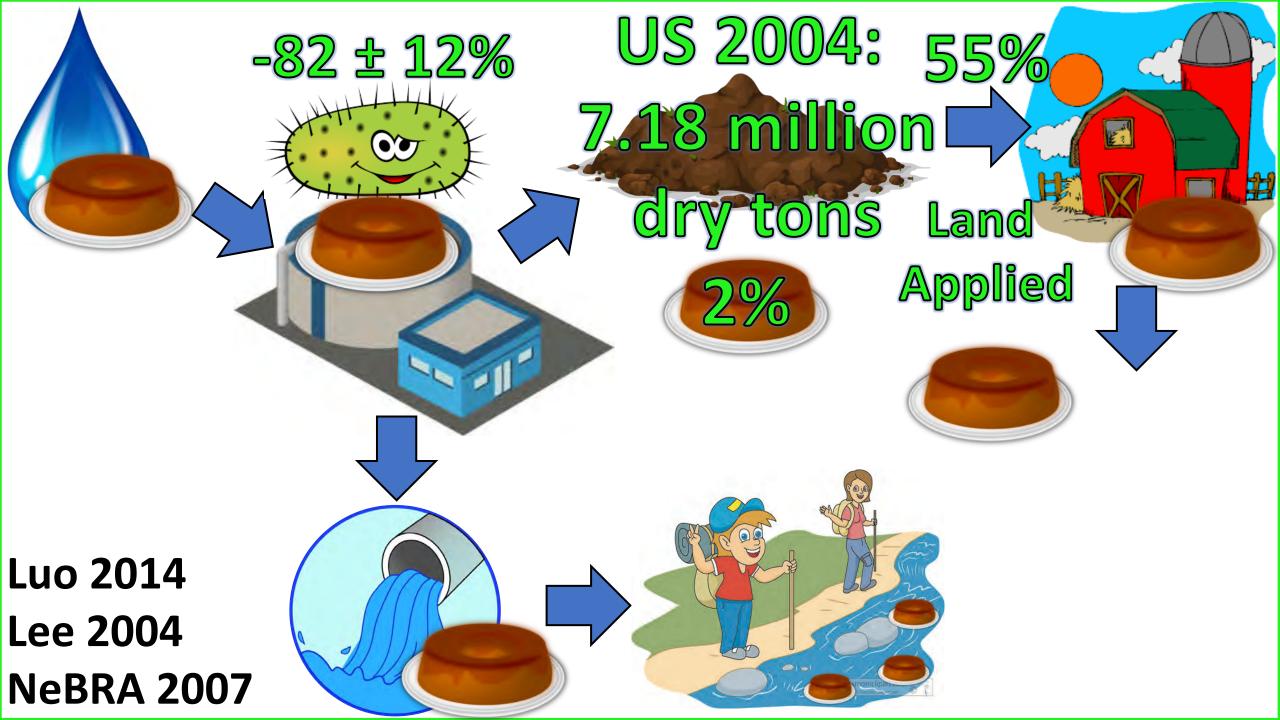


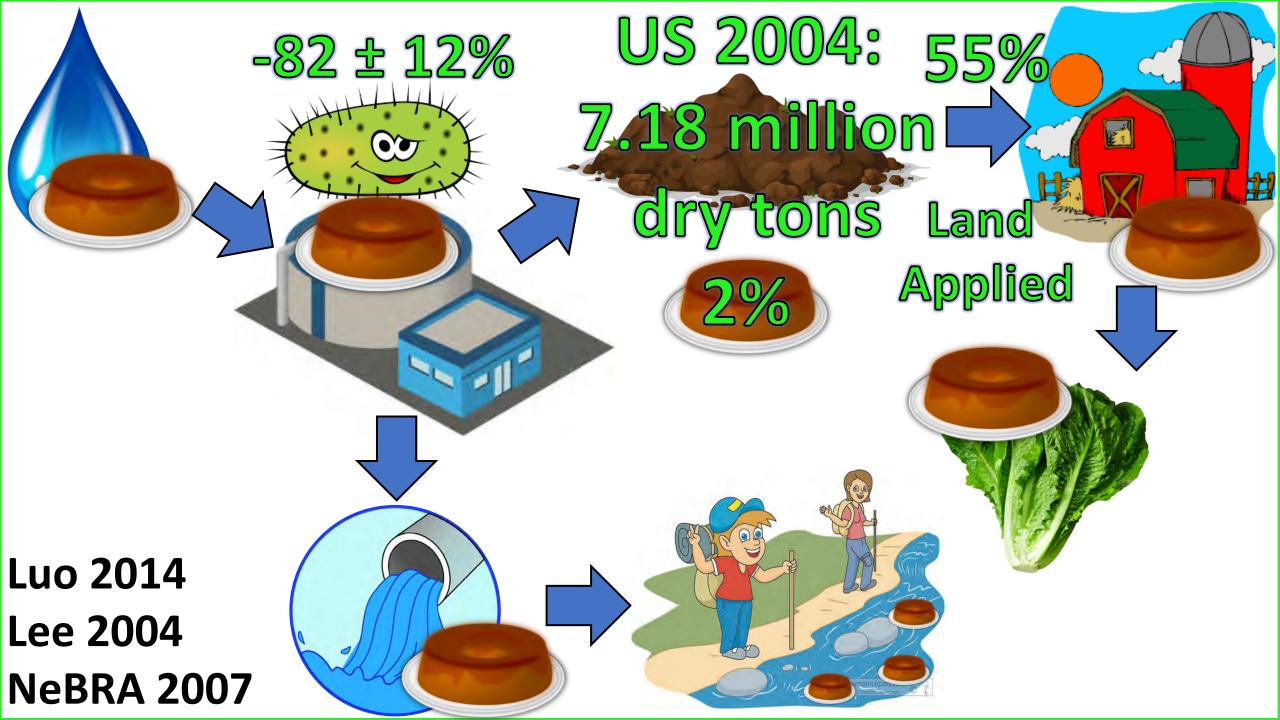


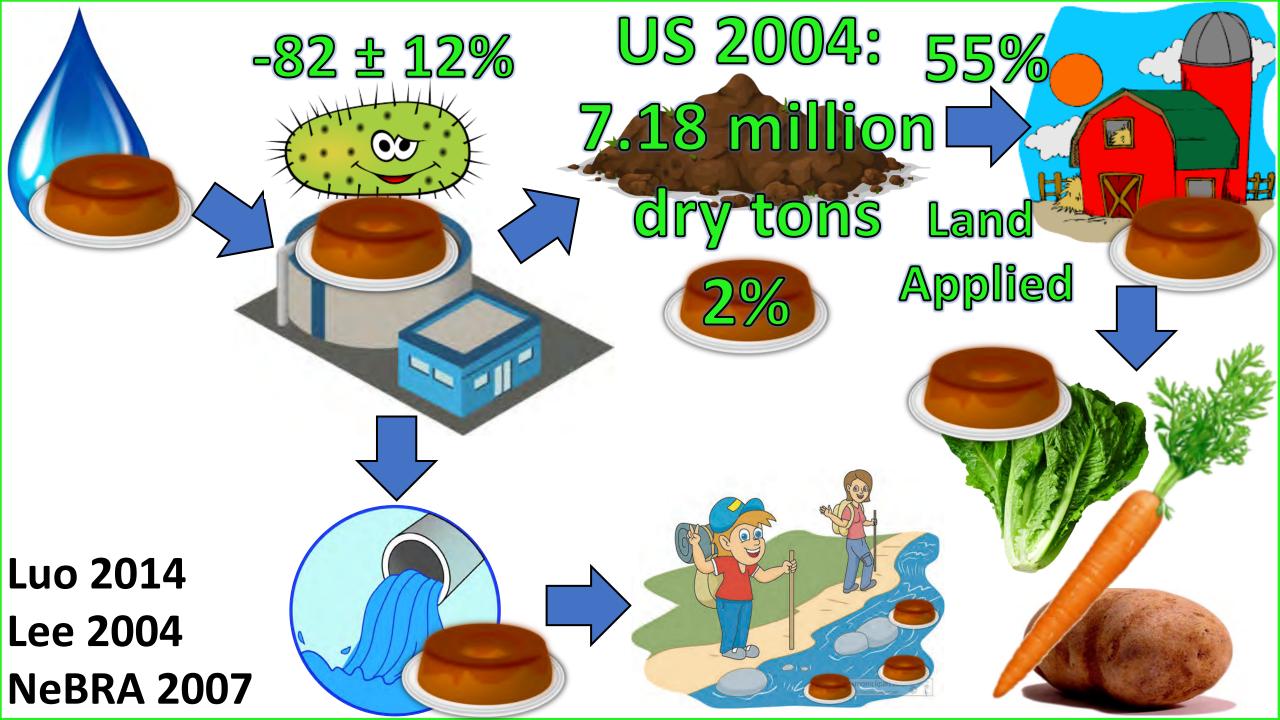


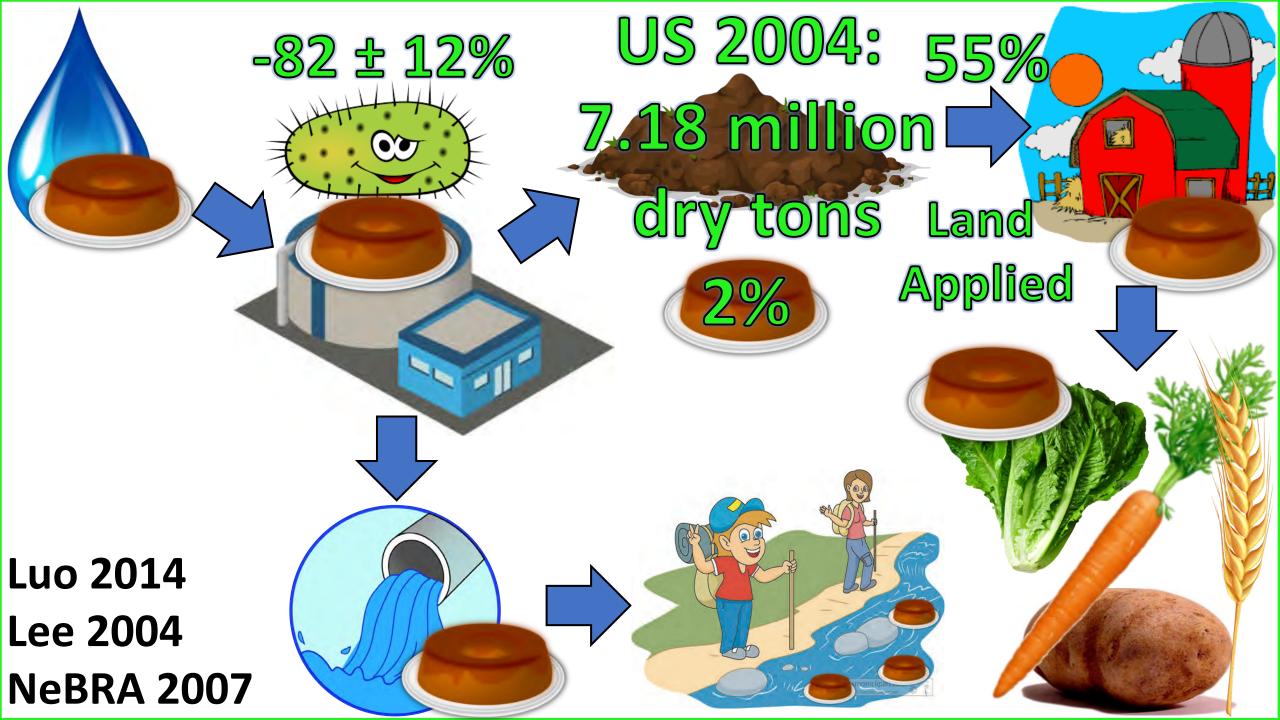










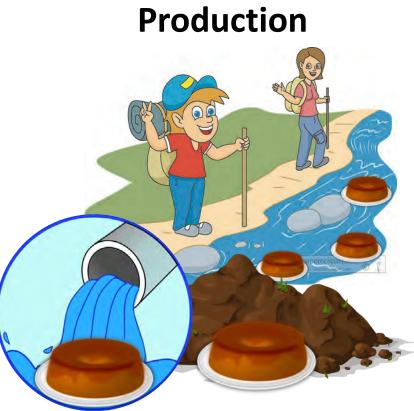




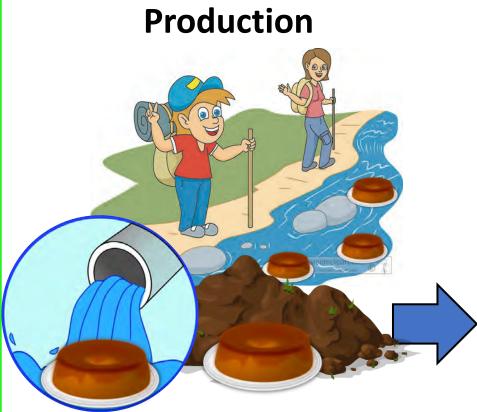
- Contamination of Air
- Contamination of Water



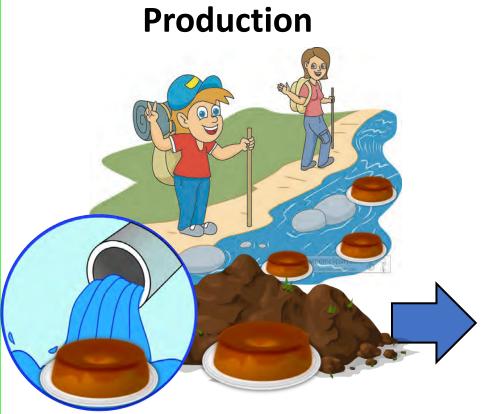
- Contamination of Air
- Contamination of Water
- Contamination of Soil



- Contamination of Air
- Contamination of Water
- Contamination of Soil

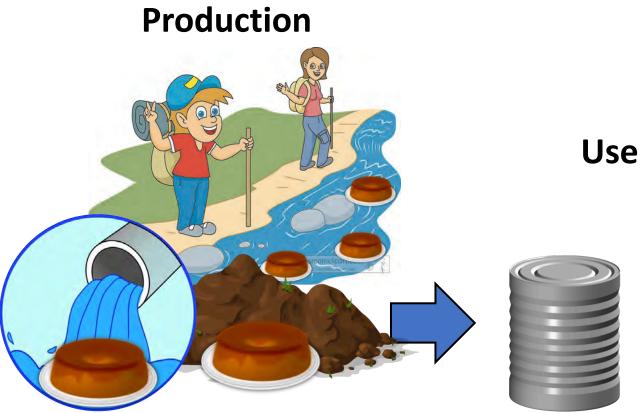


- Contamination of Air
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- Contamination of Soil

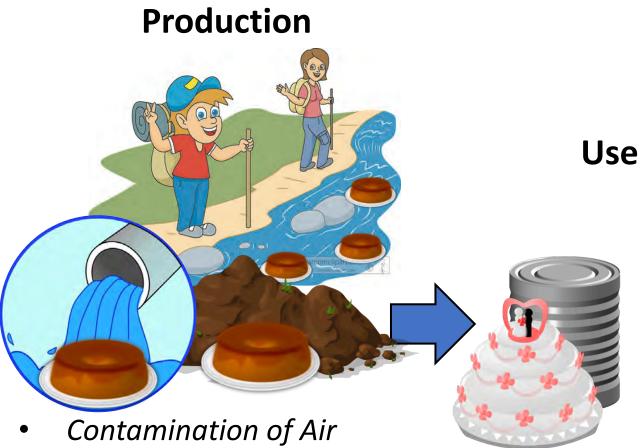


Use

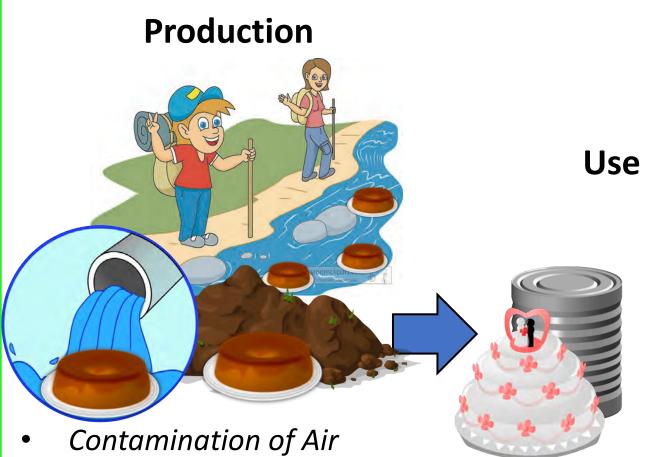
- Contamination of Air
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- Contamination of Air
- Contamination of Water
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- Contamination of Water
- Contamination of Soil



- Contamination of Water
- Contamination of Soil
- Contamination of Foods

Canned Goods: the liquid phases of canned artichokes, green beans, corn, mushrooms, peas, and mixed vegetables, the homogenized liquid and solid contents of canned fruit products including coconut cream, coconut milk, lychees, mangoes, olives, peaches, light pineapples, tomatoes, tomato juice and tomato paste, and fruit pieces and cocktails, the homogenized liquid and solid contents of canned vegetable products including asparagus, baked beans in tomato sauce, green beans, beetroot, carrots, corn, mount elephants, mushrooms, peas, jalapeño peppers, potatoes, and goulash, the homogenized liquid and solid contents of canned soups including cream of chicken, chicken and white wine, potato, tomato, and Tom Kha, the homogenized liquid and solid contents of canned sauces of many varieties including demi-glace, fond de volaille, gratin, meat, tomato, and white, the homogenized liquid and solid contents of canned pastas in tomato sauce, the homogenized liquid and solid contents of canned seafood including Japanese sand lance, mackerel, pilchards in tomato sauce, salmon, sardine, sardine in tomato sauce shrimp, squid, tuna, and fish and vegetable mixtures, the homogenized liquid and solid contents of canned meats including chicken, corned beef, fish balls, ham, hot dogs, and pork, the homogenized liquid and solid contents of canned quail eggs, the homogenized liquid and solid contents of desserts including evaporated milk and creamed rice, the solids of canned crushed tomatoes, young peas, corn, haricot beans, red kidney beans, lentils, mushrooms, tuna in oil, and sardines in oil, mackerel filet in tomato sauce, and canned dinners.

Brotons 1995 **Braunrath 2005** Thomson 2005 Yonekubo 2008 Goodson 2002 Grumetto 2008 Rastkari 2011 Mungia-Lopez 2002 Maragou 2006 Yoshida 2001 Sakhi 2014

Canned Goods: the liquid phases of canned artichokes, green beans, corn, mushrooms, peas, and mixed verets bleste homogenized awid and solid pontents of can truit prod ncluding v ves, peq Ut milk, bees, maa coconut cre itoes, an fru pieces of ocktor, t l h moge tomato jurca ato pust eaetable b beans, beetroot, carrots, corn, mount elephants_mushrooms, peas, jalapeño peppers, potatoes, and goulash, the homogenized liquid and solid ntents of canned soups includin cream of ta and mile Cand solid chicker tomato, and white, the homogenized liquid and solid Use ts of canned pastas in tomato sauce, the homogenized liquid and solid contents of caned seafood nd lance. a in dine ne zeo iquid mackerel, pilch dine np, squid, tuna, and fish nd canned enized liquid meats including and solid contents of canned quail eggs, the homogenized liquid and solid contents of desserts including evaporated milk and creamed rice, the solids of canned crushed tomatoes, young peas, corn, haricot beans, red kidney beans, lentils, mushrooms, tuna in oil, and sardines in oil, mackerel filet in tomato sauce, and canned dinners.

Canned Goods: the liquid phases of canned artichokes, green beans, corn, mushrooms, peas, and mixed vere tables the homogenized liquid and solid ontents of cancel fruit products coconut creph, out milk, thees, mr good ves, pear of point pointers, pr tomato juice no pust and fruit pieces in ocktors, the mogen line of the contents of cancel fruit pieces in the point of th ncluding beans, beetroot, carrots, corn, mount elephants_mushrooms, peas, jalapeño peppers, potatoes, and goulash, the homogenized liquid and solid ntents of canned soups includin cream of chicker (hicken a m v re) of the of and of the the of the of and solid content of a local of the of tomato, and white, the homogenized liquid and solid *Solid* ts of canned pastas in tomato sauce, the homogenized liquid and solid contents of caoed seafood i and lance, i eto le his rectification de seujoou ingresties parla lance, le parla de la parla de la porte seujoou ingresties parla lance, le parla de la porte seujoou ingresties parla lance, le parla de la porte seujoou ingresties parla lance, le parla de la porte seujoou ingresties parla lance, le parla de la porte seujoou ingresties parla lance, cetter parla de la porte seujoou ingresties parla lance, cetter parla de la porte seujoou ingresties parla lance, cetter parla de la porte seujoou ingresties parla lance, le parla de la porte seujo parla de la porte seujo parla la porte mackerel, pilch tuna, and fish meats including and solid contents of canned quail eggs, the homogenized liquid and solid contents of desserts including everyor ted milk and creamed ice, the solids of canned crisher tomatoes. Fung peas, cor of the contract of the solids of canned crisher tomatoes. Fung peas, and canned dinners.

Basheer 2004, Sakhi 2014 Vivacqua 2003, Cao 2009

baby food
 products in glass
 jars with metal
 lids

Basheer 2004, Sakhi 2014 Vivacqua 2003, Cao 2009



- baby food
 products in glass
 jars with metal
 lids
- whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

• fresh cherries



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants
 - medlars



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

Seafoods

• white clams



baby food
 products in glass
 jars with metal
 lids

 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

- white clams
 - crabs



baby food
 products in glass
 jars with metal
 lids

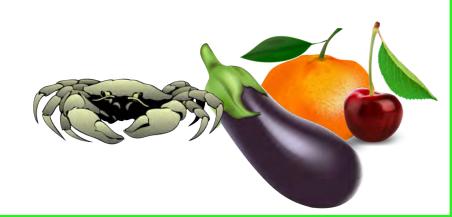
 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

- white clams
 - crabs
- blood cockles



baby food
 products in glass
 jars with metal
 lids

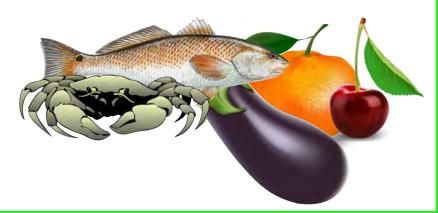
 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

- white clams
 - crabs
- blood cockles
 - fish



baby food
 products in glass
 jars with metal
 lids

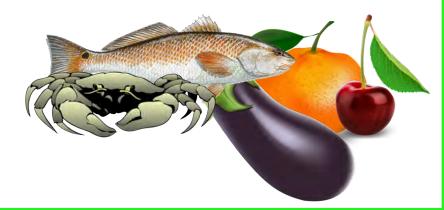
 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009



Randomly Selected Fresh Foods

- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

- white clams
 - crabs
- blood cockles
 - fish
 - prawn



baby food
 products in glass
 jars with metal
 lids

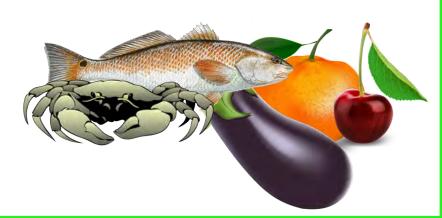
 whole eggs packaged in cardboard
 Basheer 2004, Sakhi 2014
 Vivacqua 2003, Cao 2009

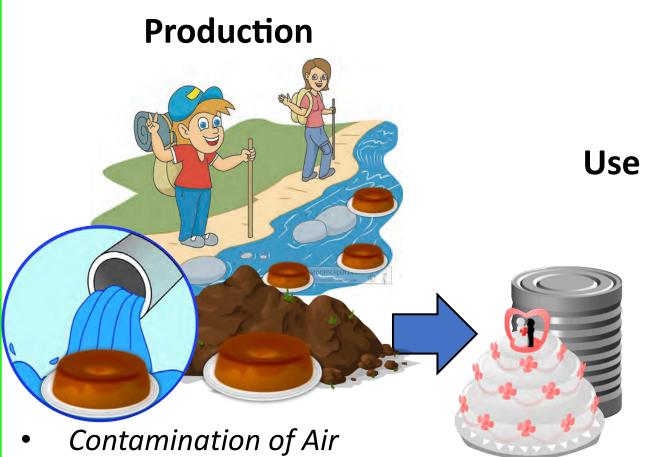


Randomly Selected Fresh Foods

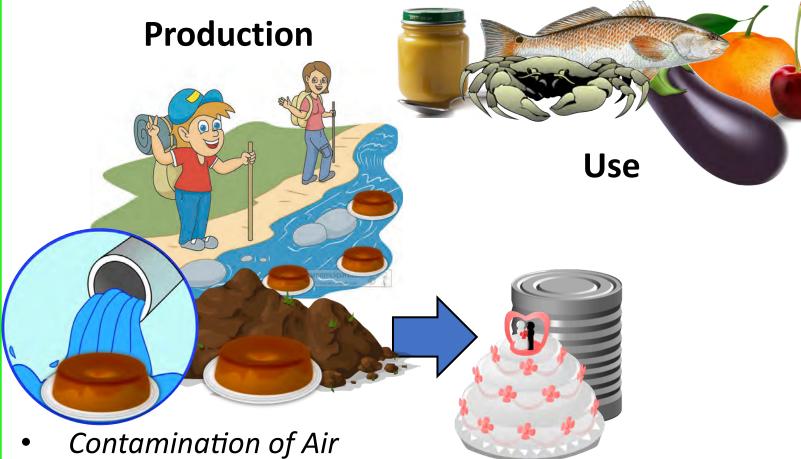
- fresh cherries
 - courgettes
 - eggplants
 - medlars
 - oranges
 - peaches
 - peppers
 - tomatoes

- white clams
 - crabs
- blood cockles
 - fish
 - prawn
 - squid





- Contamination of Water
- Contamination of Soil
- Contamination of Foods



- Contamination of Water ullet
- Contamination of Soil •
- Contamination of Foods •



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

• canned beers

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- *decaffeinated and non-decaffeinated coffees*

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated coffees
- soft drinks including diet and regular ginger ales

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

canned beers

- energy drinks
- decaffeinated and non-decaffeinated coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated orange and lemon soft drinks coffees
- soft drinks including diet and regular *ginger ales*
- diet and regular root beers •
- diet and regular colas

energy drinks

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

- energy drinks
- orange and lemon soft drinks
- *flavoured and unflavoured soda waters*
- teas and tonic waters

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated •
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

- energy drinks
- orange and lemon soft drinks
- *flavoured and unflavoured soda waters*
- teas and tonic waters
- infant and follow up formulas

Tateoka 2014, Braunrath 2005, Takao 1999, Kang 2002

- canned beers
- decaffeinated and non-decaffeinated •
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

<u>Tateoka 2014</u>,

Braunrath 2005, Takao 1999, Kang 2002



- energy drinks
 - orange and lemon soft drinks
- *flavoured and unflavoured soda waters*
- teas and tonic waters
- infant and follow up formulas

- canned beers
- decaffeinated and non-decaffeinated •
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

s A

6.2 μg L⁻¹ of BPA in her breastmilk

<u>Tateoka 2014,</u>

Braunrath 2005,

Takao 1999, Kang 2002

- energy drinks
 - orange and lemon soft drinks
- *flavoured and unflavoured soda waters*
- teas and tonic waters
- infant and follow up formulas

- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

6.2 μg L⁻¹ of BPA in her breastmilk

<u>Tateoka 2014,</u>

Braunrath 2005,

Takao 1999, Kang 2002

- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas



- canned beers
- decaffeinated and non-decaffeinated •
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

6.2 μg L⁻¹ of BPA in her breastmilk

<u>Tateoka 2014,</u>

Braunrath 2005,

Takao 1999, Kang 2002

- energy drinks
 - orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas



- canned beers
- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger ales
- diet and regular root beers
- diet and regular colas

6.2 μg L⁻¹ of BPA in her breastmilk

<u>Tateoka 2014,</u>

Braunrath 2005,

Takao 1999, Kang 2002

- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas

ca. 30 μg L⁻¹ of BPA in her breastmilk

Cao 2009, Sakhi 2014, Biles 1997, Ackerman 2010, Cao 2008, Kuo 2004

canned beers

diet

die

Tateoka 2014,

Braunrath 2005,

Takao 1999, Kang 2002

- decaffeinated and non-decaffeinated
 coffees
- soft drinks including diet and regular ginger

ar root beers

 $6.2 \ \mu g \ L^{-1} \ of \ BPA$

in her breastmilk

r colas

- energy drinks
- orange and lemon soft drinks
- *flavoured and unflavoured soda waters*
- teas and tonic waters
- infant and follow up formulas

ca. 30 μg L⁻¹ of BPA in her breastmilk

Cao 2009, Sakhi 2014, Biles 1997, Ackerman 2010, Cao 2008, Kuo 2004

canned beers

diet

die

- decaffeinated and non-decaffeinated coffees
- soft drinks including diet and regular ginger

ar root beers

r colas

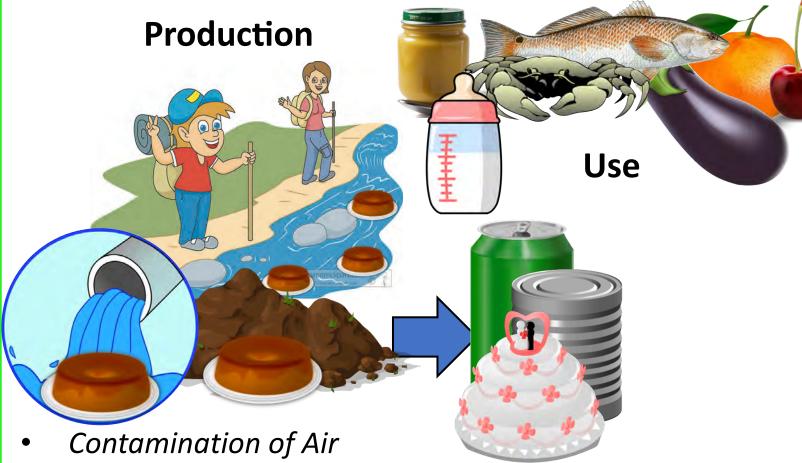
- energy drinks
- orange and lemon soft drinks
- flavoured and unflavoured soda waters
- teas and tonic waters
- infant and follow up formulas

ca. 30 μ g L⁻¹ of BPA in her breastmilk

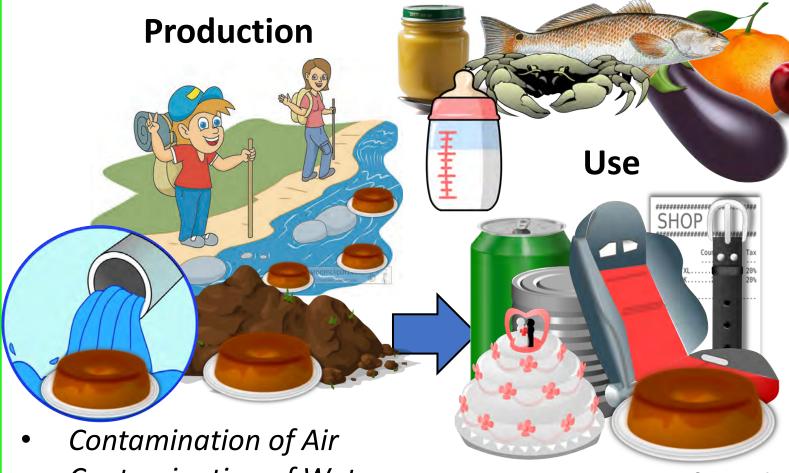
 $6.2 \ \mu g \ L^{-1} \ of \ BPA$ 37.4 µg in her breastmilk Cao 2009, Sakhi 2014, **Tateoka 2014,** Biles 1997, Ackerman 2010, Braunrath 2005, Cao 2008, Kuo 2004 Takao 1999, Kang 2002



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products

Vandenberg 2013, Zalko 2011, Biedermann 2010, Dodson 2012

SHOP	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,
Article	Count Amount Tax
T-Shirt XL	1 \$14,95 20%
Jeans PDK	2 \$39,90 20%
GR0SS	\$75,80
TAX	\$18,95
SUM	\$94,75



• When we touch receipts, BPA is transferred to the skin

Vandenberg 2013, Zalko 2011, Biedermann 2010, Dodson 2012

SHOP)) ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	
Article	Count Amount	Тах
T-Shirt XL Jeans PDK	1 \$14,95 2 \$39,90	
GROSS TAX SUM	\$75,80 \$18,95 \$94,75 ======	



- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy

Vandenberg 2013, Zalko 2011, Biedermann 2010, Dodson 2012

SHO)) ;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;	
Article	Count Amount	Тах
	1 \$14,95	
Jeans PDK	2 \$39,90	20%
GROSS	\$75,80	
TAX	\$18,95	
SUM	\$94,75	

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin

Vandenberg 2013, Zalko 2011, Biedermann 2010, Dodson 2012

Article Count Amount Tax T-Shirt XL1 \$14,95 20% Jeans PDK2 \$39,90 20% GROSS \$75,80 TAX \$18,95 SUM \$94,75 ======	SHOP)) ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	
T-Shirt XL1 \$14,95 20% Jeans PDK2 \$39,90 20% 	Article	Count Amount	Тах
Jeans PDK2 \$39,90 20% GROSS \$75,80 TAX \$18,95			
GROSS \$75,80 TAX \$18,95	T-Shirt XL	1 \$14,95	20%
TAX \$18,95	Jeans PDK	2 \$39,90	20%
	TAX	\$18,95	

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann
 2010. Dodson 2012

Article Count Amount Tax
GROSS \$75,80 TAX \$18,95 SUM \$94,75

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann





- When we touch receipts, BPA is transferred to the skin
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- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann



- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann





- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann





Cleansers

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann

2010, Dodson 2012



- Cleansers
- Conditioner

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann

- Cleansers
- Conditioner
- Shaving cream



- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann

- Cleansers
- Conditioner
- Shaving cream
- Lotions



- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann



- Cleansers
- Conditioner
- Shaving cream
- Lotions
- Shampoo

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann



- Cleansers
- Conditioner
- Shaving cream
- Lotions
- Shampoo
- Bar soap

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann



- Cleansers
- Conditioner
- Shaving cream
- Lotions
- Shampoo
- Bar soap
- Sunscreen

- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann

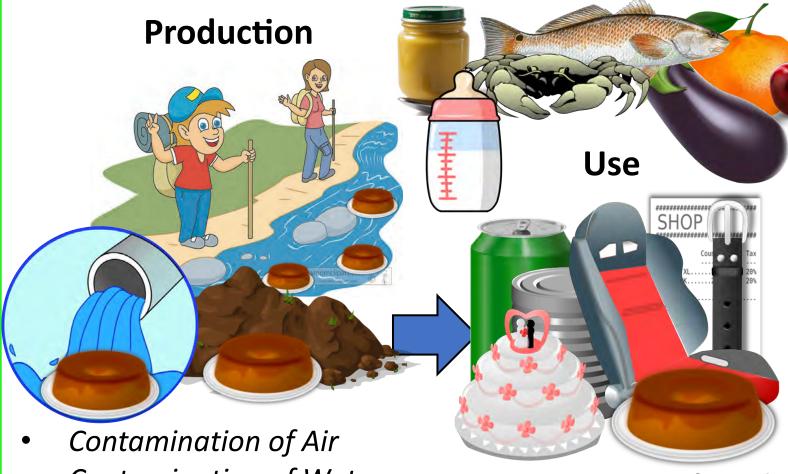


- Cleansers
- Conditioner
- Shaving cream
- Lotions
- Shampoo
- Bar soap
- Sunscreen
- Toothpaste

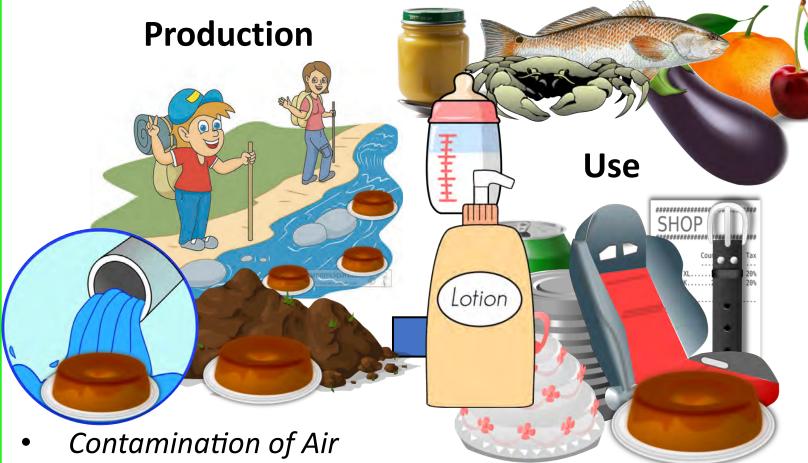
- When we touch receipts, BPA is transferred to the skin
- The amount transferred to the skin increases by a factor of 10 when skin is wet or greasy
- The BPA can then migrate into the skin
- Migration increases when a vector such as hand cream or lotion is present
 Vandenberg 2013, Zalko 2011, Biedermann



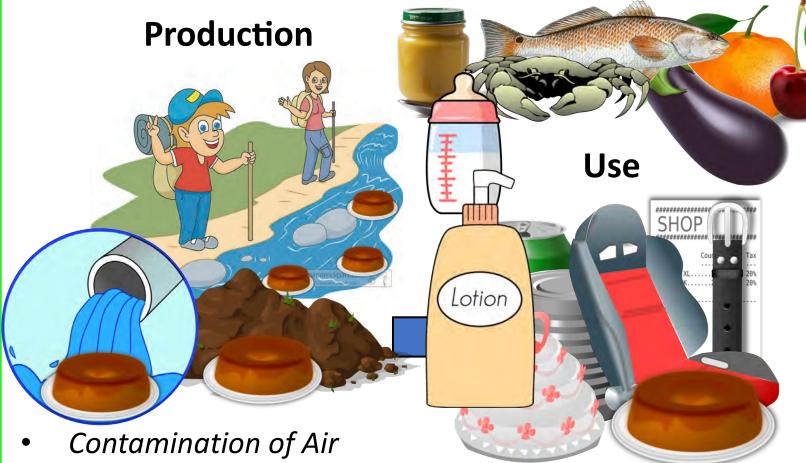
- Cleansers
- Conditioner
- Shaving cream
- Lotions
- Shampoo
- Bar soap
- Sunscreen
- Toothpaste
- Face and body wash



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products



- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products
- Contamination of People

What Are the Concentrations of BPA in People?

Human Tissue and Fluid Concentrations of BPA and Total BPA. Refs: vom Saal 2007, NHANES 2003, Kuruto-Niwa 2007, Ye 2006						
Fluid or Tissue	n	% Positive	Range (µg L ⁻¹)	Central Tendency / Mean (μg L ⁻¹)		
Foetal, Child, and Adult Fluids and Tissues				Central Tendency: 0.3–4.4 µg L ⁻¹ BPA		
Urine of Americans Over 6 Years Old	2,517	92.6%	0.3−149 µg L ⁻¹ Total BPA	Mean: 5.2 μg L ⁻¹ Total BPA		
Colostrum of Japanese Mothers	101	100%	1–7 μg L ⁻¹ Total BPA	Mean: 3.41 µg L ⁻¹ Total BPA		
Breast Milk of American Mothers	20	90%	<0.3–6.3 µg L ⁻¹ of BPA	Mean: 1.3 μg L ⁻¹ BPA		

Human Tissue and Fluid Concentrations of BPA and Total BPA. Refs: vom Saal 2007, NHANES 2003, Kuruto-Niwa 2007, Ye 2006						
Fluid or Tissue	n	% Positive	Range (µg L ⁻¹)	Central Tendency / Mean (μg L ⁻¹)		
Foetal, Child, and Adult Fluids and Tissues				Central Tendency: 0.3–4.4 μg L ⁻¹ BPA		
Urine of Americans Over 6 Years Old	2,517	92.6%	0.3–149 μg L ⁻¹ Total BPA	Mean: 5.2 μg L ⁻¹ Total BPA		
Colostrum of Japanese Mothers	101	100%	1–7 μg L ⁻¹ Total BPA	Mean: 3.41 μg L ⁻¹ Total BPA		
Breast Milk of American Mothers	20	90%	<0.3–6.3 µg L ⁻¹ of BPA	Mean: 1.3 μg L ⁻¹ BPA		

Human Tissue and Fluid Concentrations of BPA and Total BPA. Refs: vom Saal 2007, NHANES 2003, Kuruto-Niwa 2007, Ye 2006						
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What Are Human Health Effects of These Exposures?





 "Confident" of human effects on the male reproductive tract arising from adult exposures

- *"Confident" of human effects on the male reproductive tract arising from adult exposures*
 - "Confident" of effects on the organization of the reproductive tract of males, the brain, and metabolism arising from developmental exposures

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 - "Confident" of effects on the organization of the reproductive tract of males, the brain, and metabolism arising from **developmental exposures**

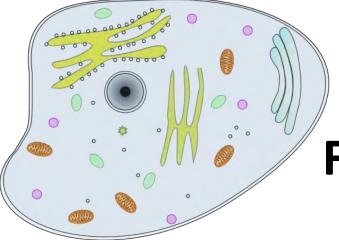
US National Toxicology Program Panel and the US Food and Drug Administration

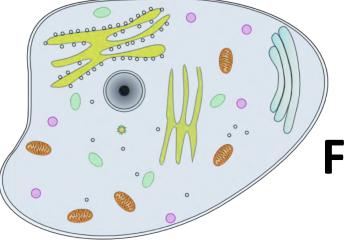
- "Confident" of human effects on the male reproductive tract arising from adult exposures
- "Confident" of effects on the organization of the reproductive tract of males, the brain, and metabolism arising from developmental exposures

US National Toxicology Program Panel and the US Food and Drug Administration

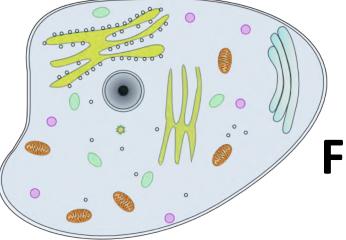
 "Some concern" for effects on the brain, behavior, and male prostate gland arising from foetus, infant, and child exposures

What Effects Have Been Reported To Arise From Exposures Of Human Cells To These BPA Concentrations?

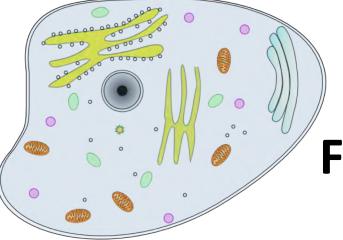




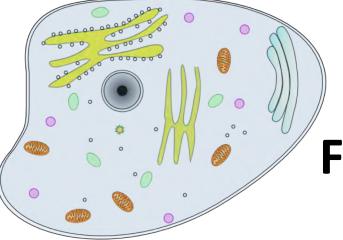
 BPA can be as potent as estradiol, an endogenous hormone Prins 2014



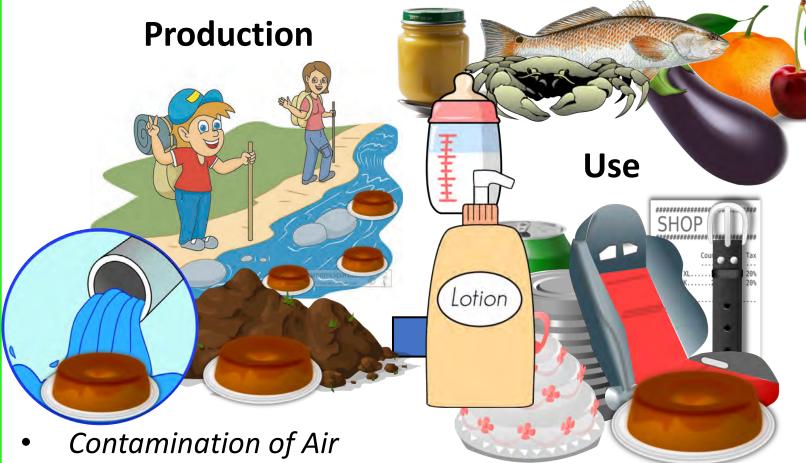
- BPA can be as potent as estradiol, an endogenous hormone Prins 2014
- Long-term exposure may significantly affect human immunity Watanabe 2003



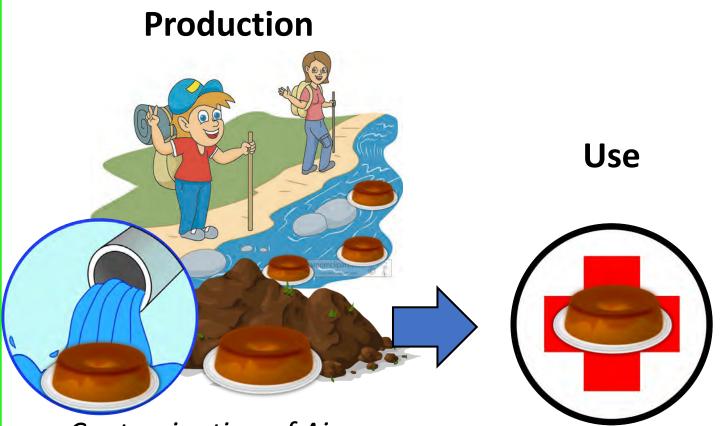
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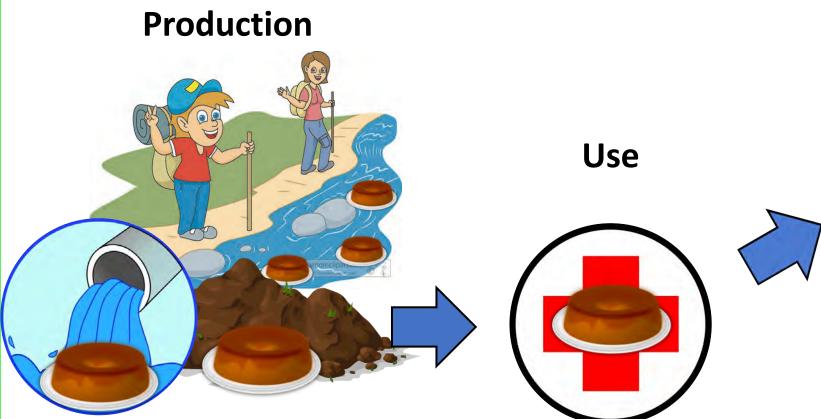
- BPA can be as potent as estradiol, an endogenous hormone Prins 2014
- Long-term exposure may significantly affect human immunity Watanabe 2003
- Exposure of the developing human prostate epithelium increases its susceptibility to hormonal carcinogenesis Prins 2014
- Exposure may adversely effect metabolic homeostasis
 Ben-Jonathan 2009, Hugo 2008



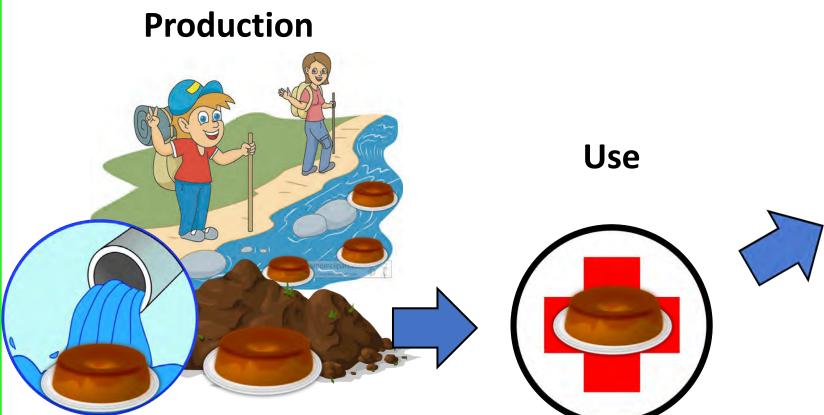
- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products
- Contamination of People



- Contamination of Air
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Disposal

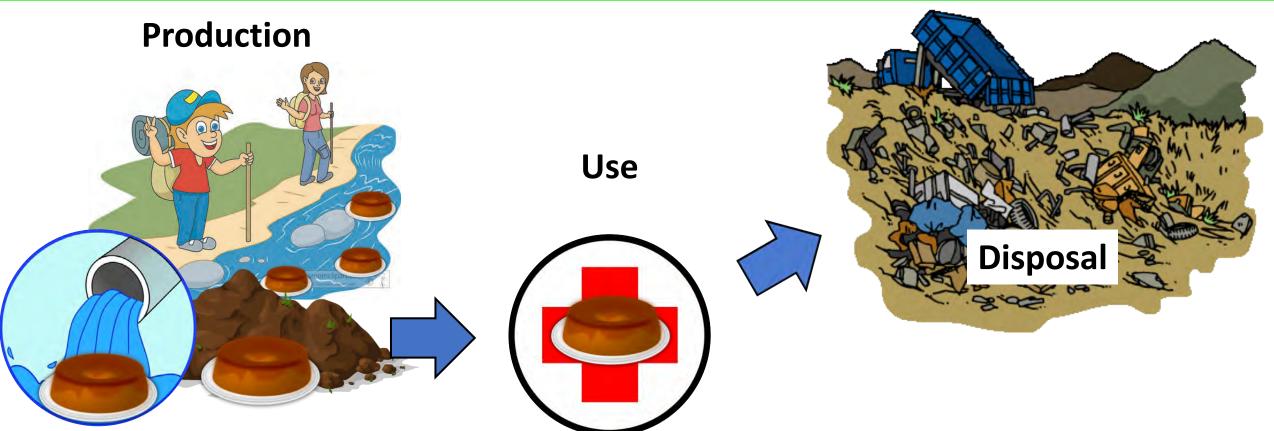
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What Is the Fate of BPA Containing Goods?

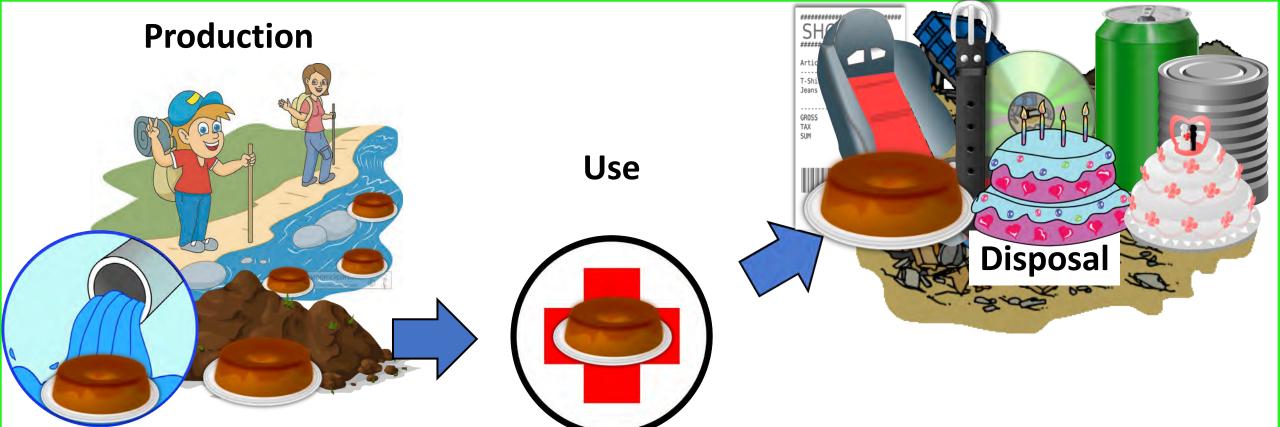
In the US in 2014, 258 million tons of municipal solid waste were generated.

Waste	Million tons (mt)	Percent Landfilled	Percent Recycled	Percent Combusted
Paper and Paperboard	69	28	65	7
Plastics	33	76	10	15
Textiles	16	65	16	19
Rubber and Leather	8	51	18	32

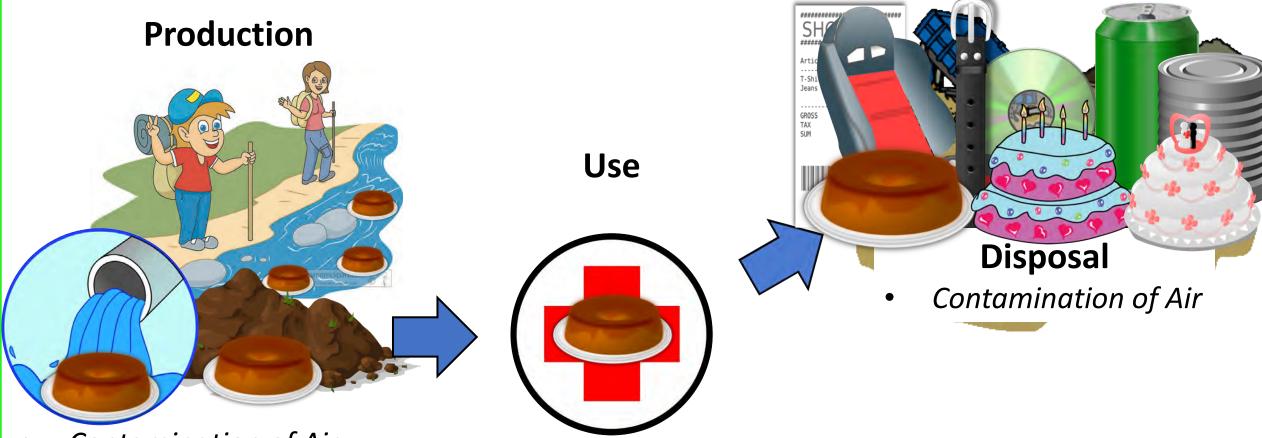
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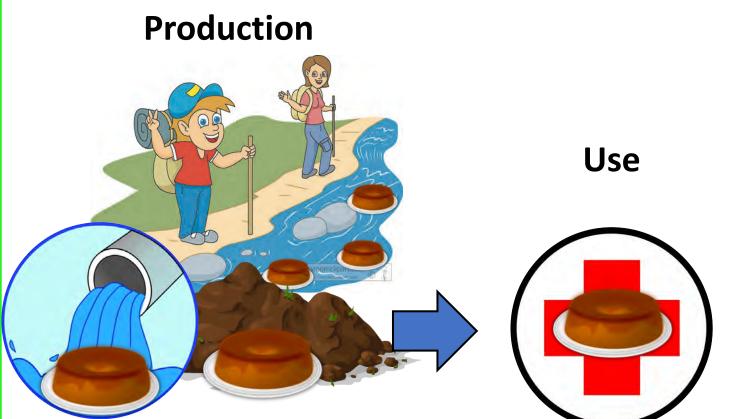
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Polycarbonate

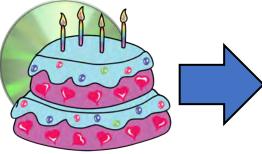
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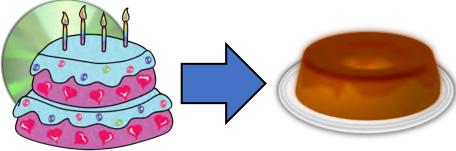
60% of the produced BPA

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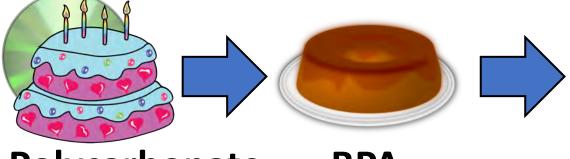
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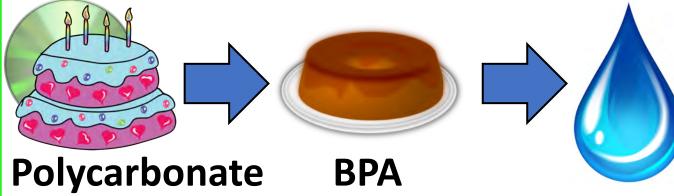
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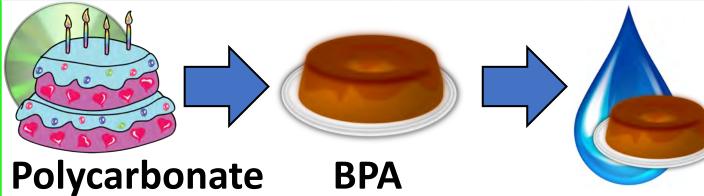
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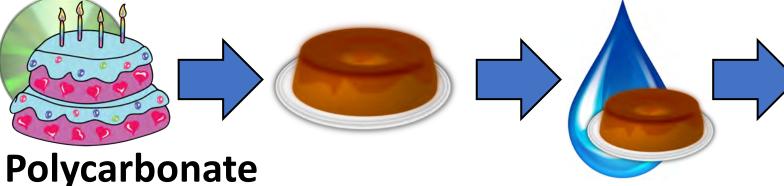
Concentrations of BPA Detected in Landfill Leachate						
Country	Year	Range (µg L ⁻¹)	Average (µg L ⁻¹)	Ref.		
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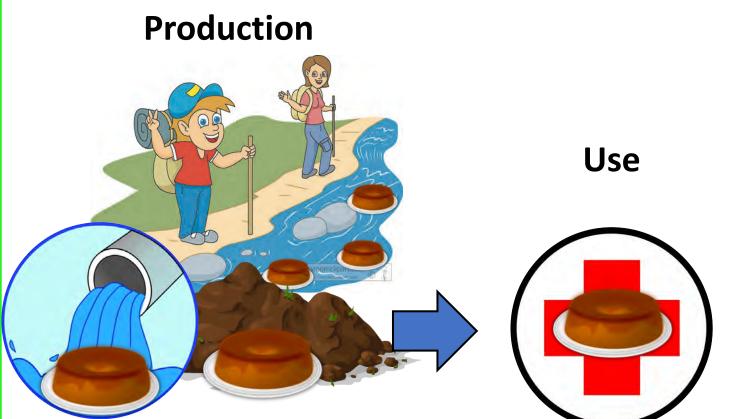


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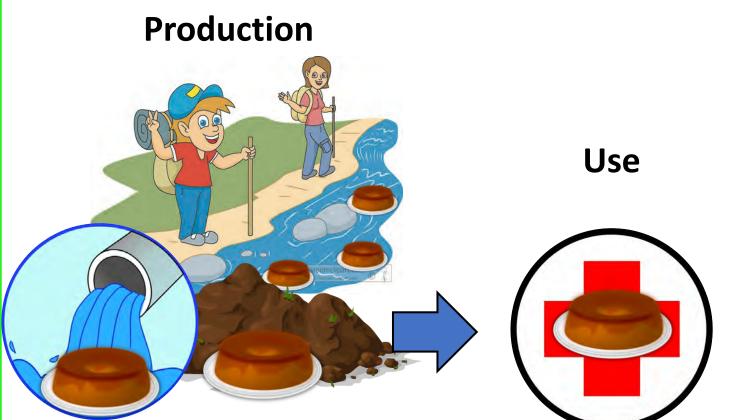
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- Contamination of Water

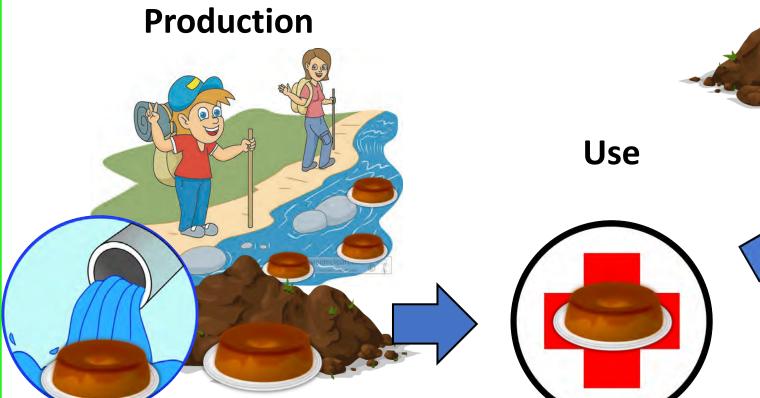
- Contamination of Air
- Contamination of Water
- Contamination of Soil
- Contamination of Foods
- Contamination of Beverages
- Contamination of Products
- Contamination of People





- Contamination of Air
- Contamination of Water
- Contamination of Soil

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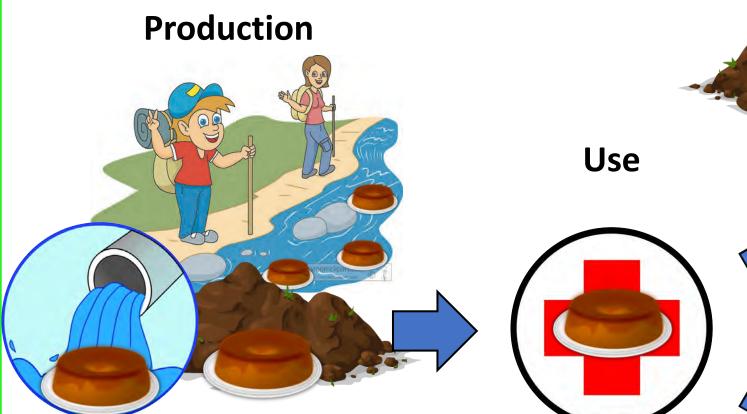
Disposal

Contamination of Air

Contamination of Soil

Contamination of Water

- **Contamination of Products**
- Contamination of People



- Contamination of Air
- Contamination of Water
- Contamination of Soil
- Contamination of Foods
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- Contamination of People

- Contamination of Air
- Contamination of Water
- Contamination of Soil

• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People

- Contamination of Air
- Contamination of Water
- Contamination of Soil



• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People

- Contamination of Air
- Contamination of Water
- Contamination of Soil



• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People



Disposal

Contamination of Air

Contamination of Recycled Cellulose Fiber

• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People

Disposal Contamination of Air Contamination of Water Contamination of Soil SHOP <u>Article Court Amount Tay</u> <u>For Form 1 Stars 200</u>

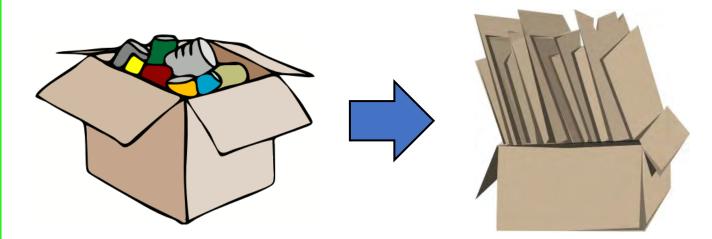
Recycling

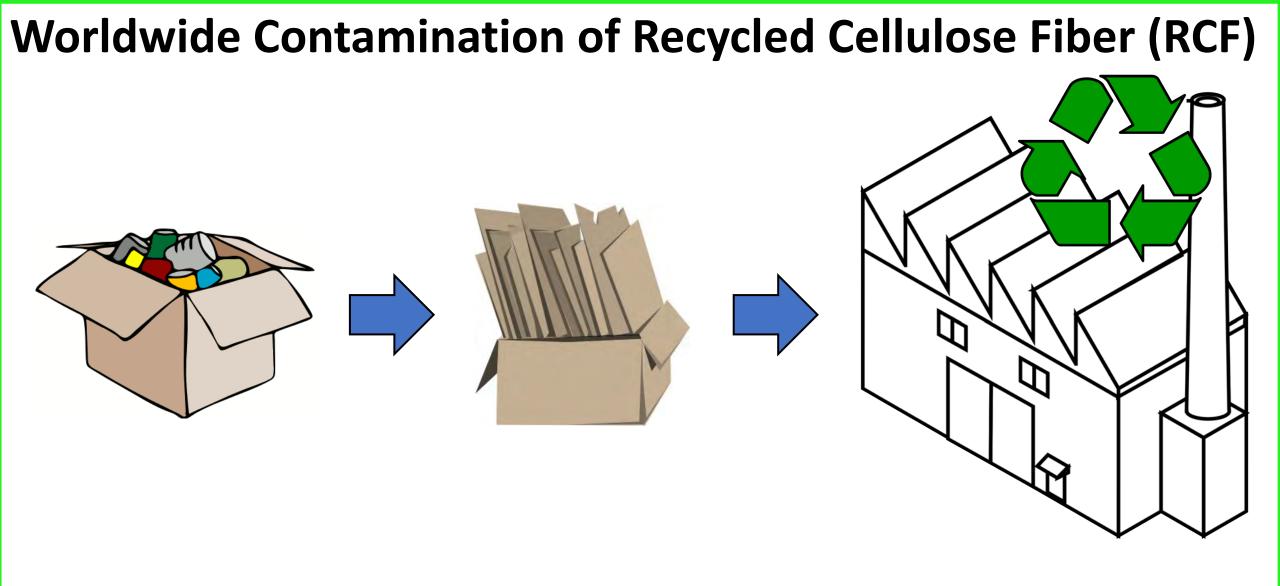
Contamination of Recycled Cellulose Fiber

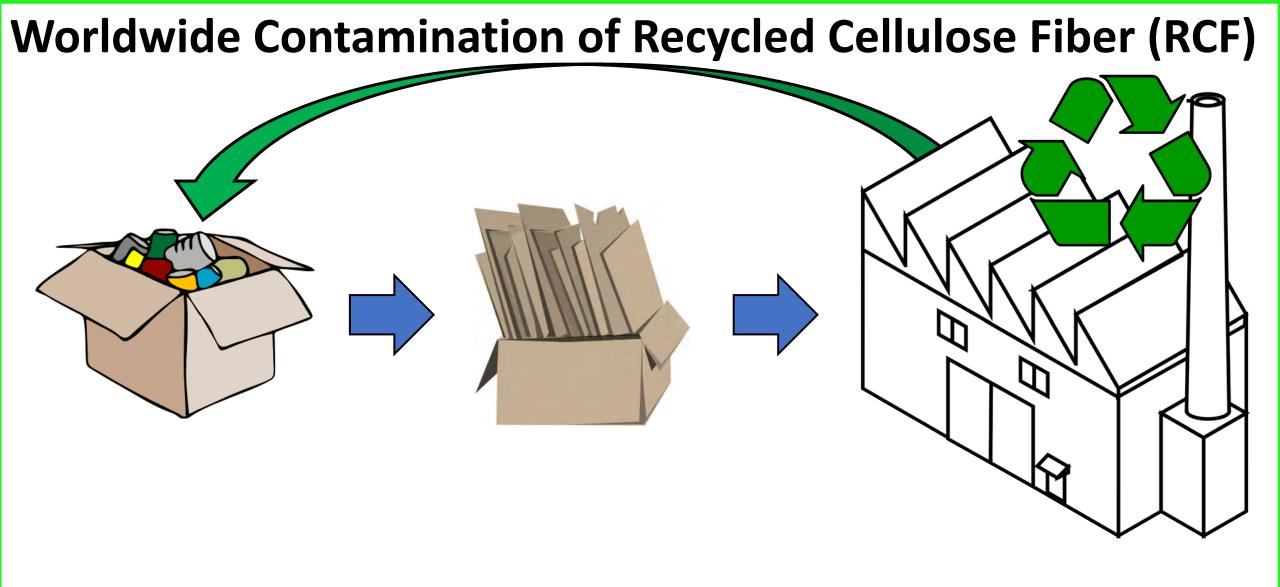
Contamination of Air, Water, and Soil

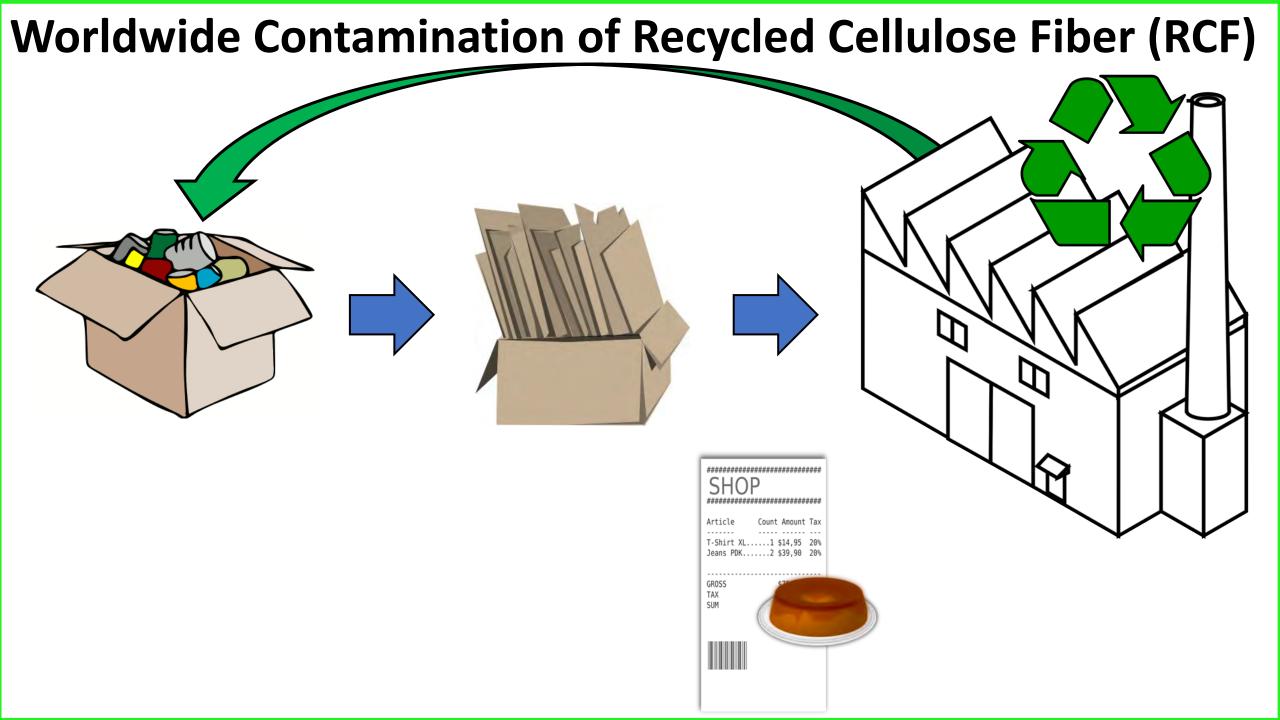


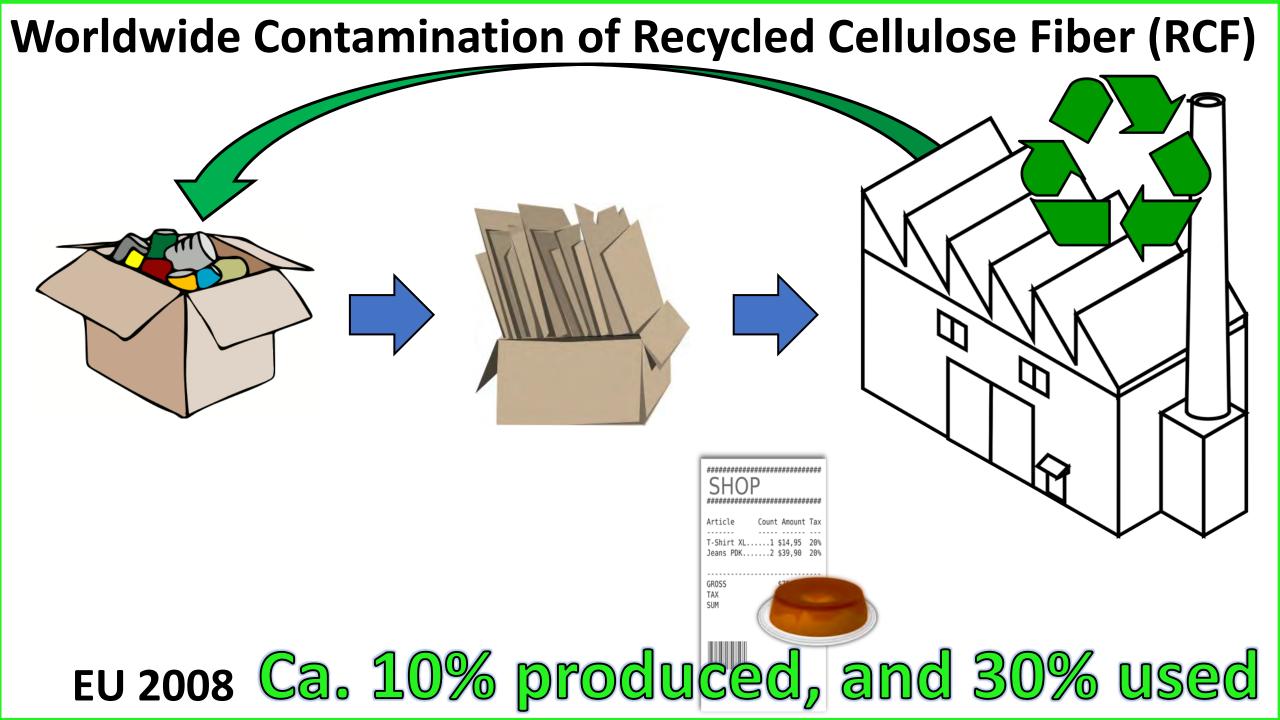


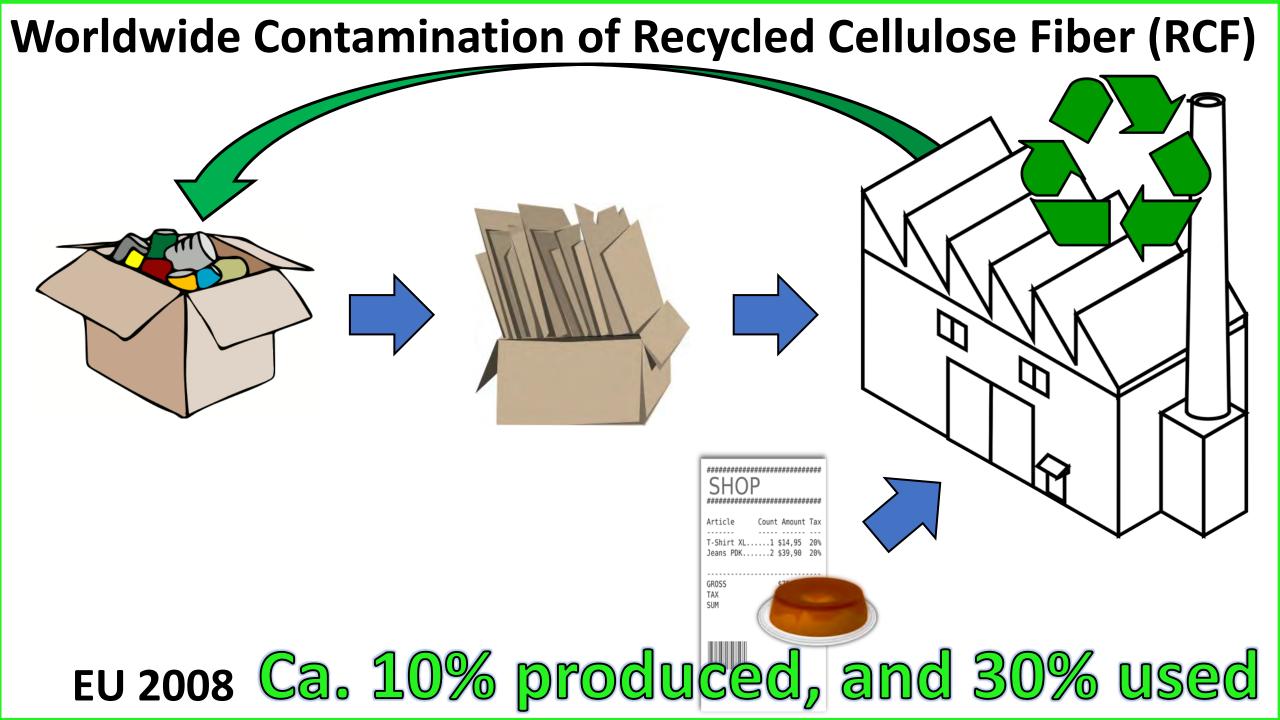


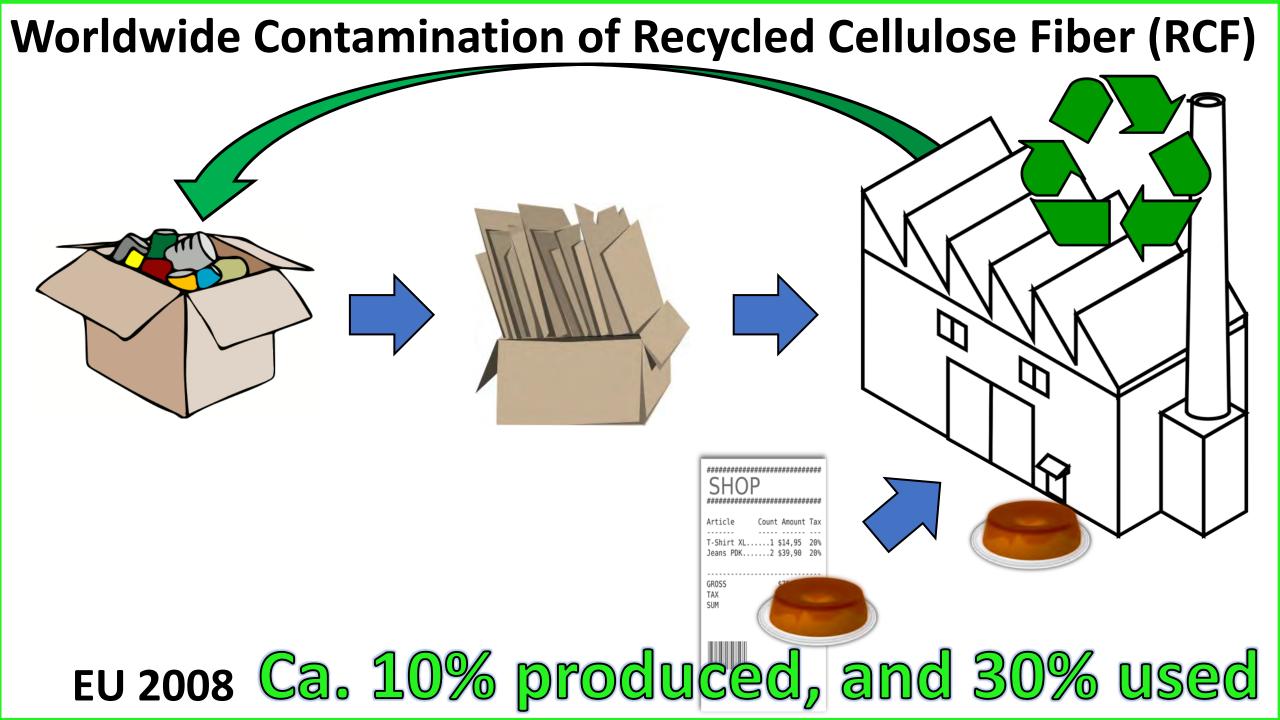


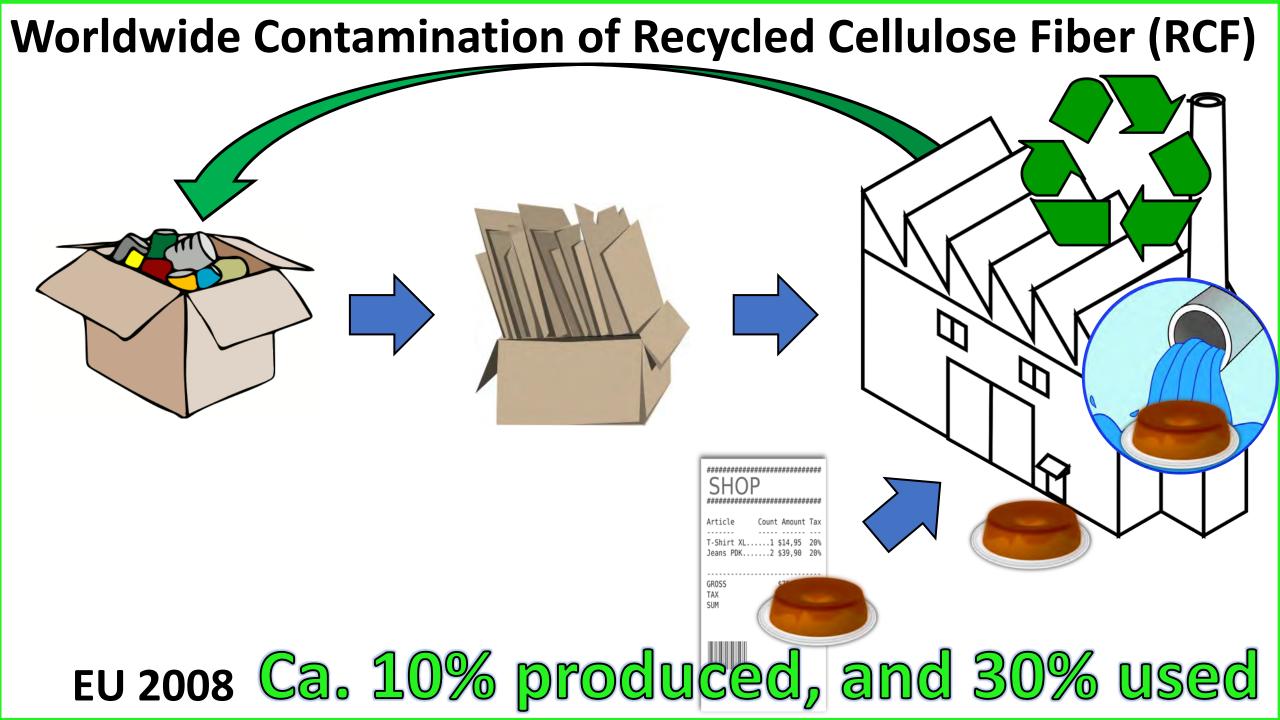


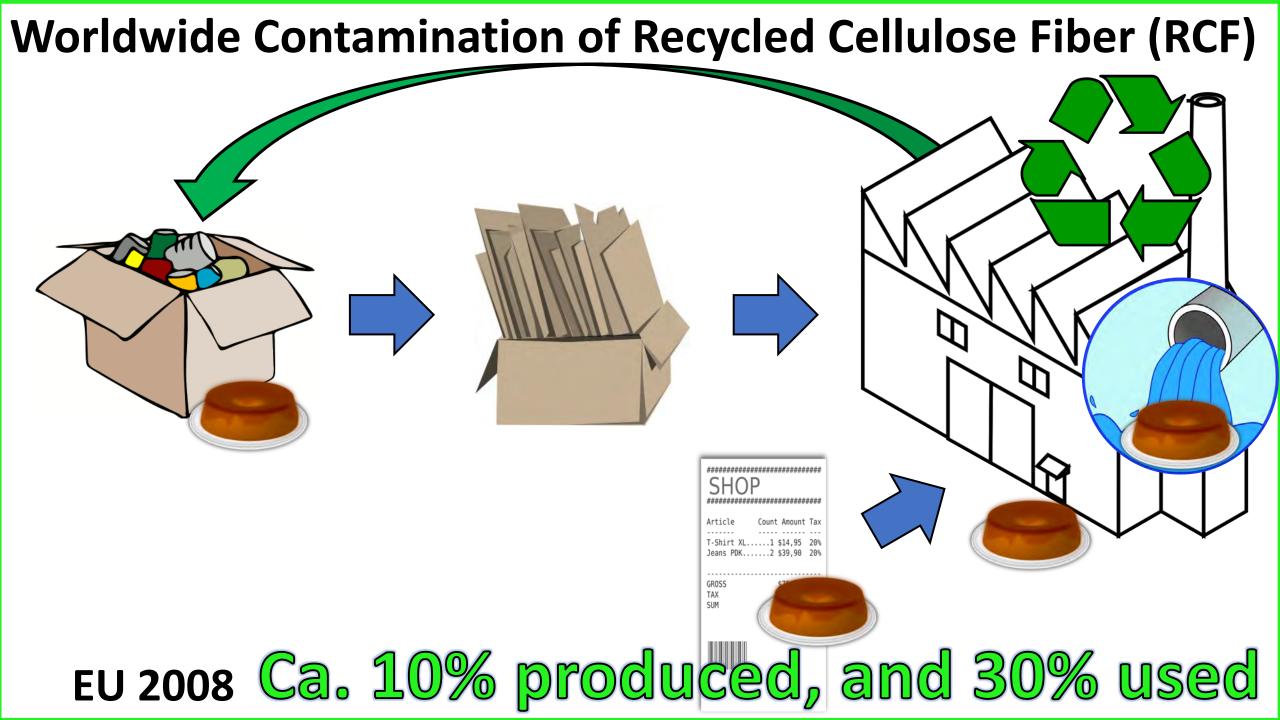


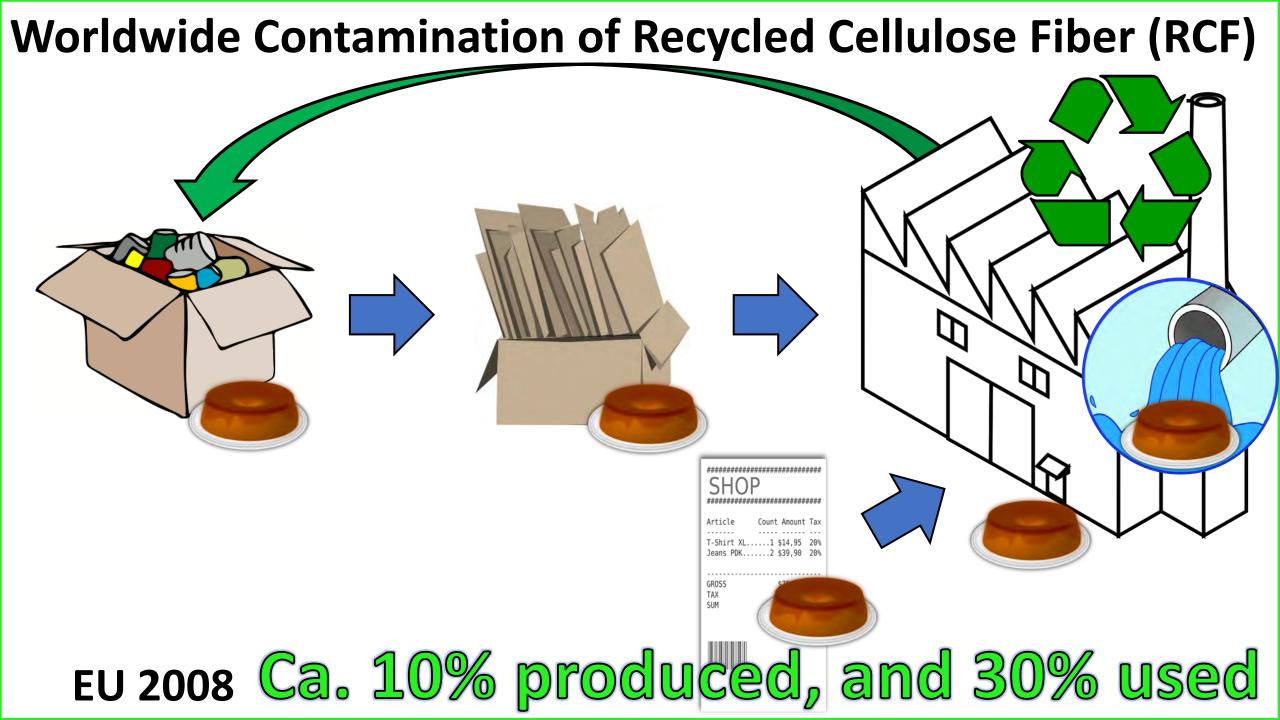












Worldwide Contamination of Recycled Cellulose Fiber (RCF) Ca. 50% of the furnish for worldwide paper and board production likely contaminated Liao 2011, Gehring 2014, Nordstrom 2013 TAX SUM EU 2008 Ca. 10% produced, and 30% used

• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People

Contamination of Air
 Contamination of Water
 Contamination of Soil
 Contamination of Soil

Recycling

- Contamination of Recycled Cellulose Fiber
- Contamination of Air, Water, and Soil

Contamination of Air

• Contamination of Water

Production

• Contamination of Soil

Contamination of Foods

Use

- Contamination of Beverages
- Contamination of Products
- Contamination of People



- Contamination of Air
- Contamination of Water
- Contamination of Soil



- Contamination of Recycled Cellulose Fiber
- Contamination of Air, Water, and Soil

- Contaminated Air
- Contaminated Water

Production

• Contaminated Soil

- Contaminated Foods
- Contaminated Beverages

Use

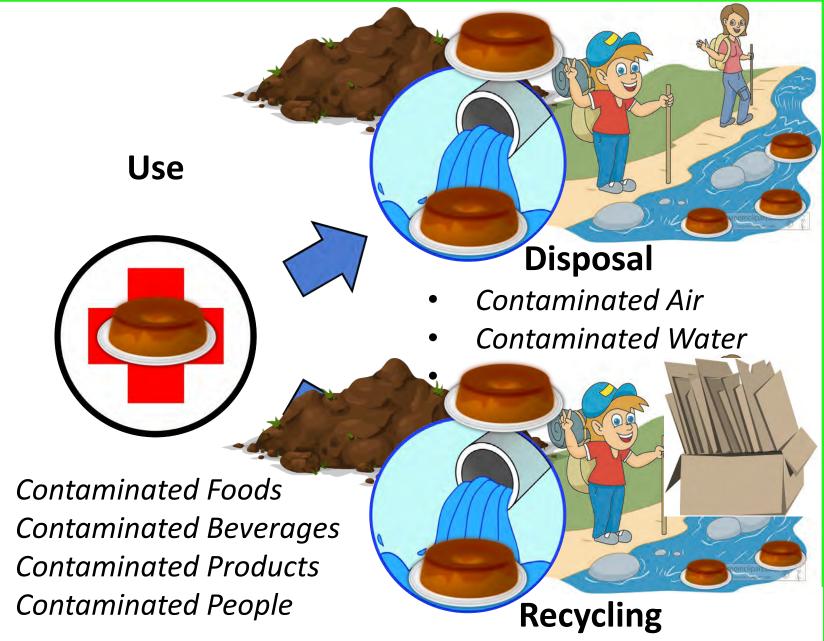
- Contaminated Products
- Contaminated People

Disposal

- Contaminated Air
- Contaminated Water

Recycling

- Contaminated Recycled Cellulose Fiber
- Contaminated Air, Water, and Soil



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- Contaminated Recycled Cellulose Fiber
- Contaminated Air, Water, and Soil



- Contaminated Air
- Contaminated Water

Recycling

- Contaminated Recycled Cellulose Fiber
- Contaminated Air, Water, and Soil





Remove BPA from landfill
 leachate



• Remove BPA from landfill leachate and WWTP influent



• Remove BPA from landfill leachate and WWTP influent



- Remove BPA from landfill
 leachate and WWTP
 influent
- Remove BPA from RCF processing solutions



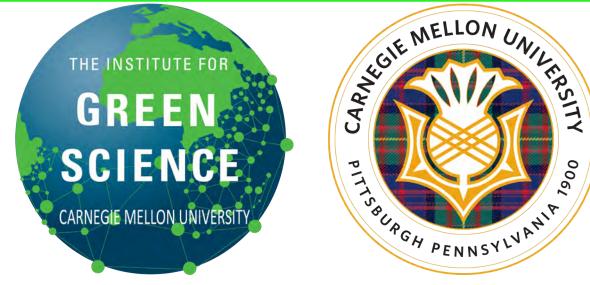
- Remove BPA from landfill
 leachate and WWTP
 influent
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00₆₁





Convert >99% of BPA in nearneutral pH lab water, a simulant of landfill leachate and WWTP influent, to an insoluble form





- Convert >99% of BPA in nearneutral pH lab water, a simulant of landfill leachate and WWTP influent, to an insoluble form
- Reduce the solution phase acute toxicity (takes EC₅₀ of 21 to >100)

THE INSTITUTE FOR **GREEN SCIENCE** CARNEGIE MELLON UNIVERSITY



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THE INSTITUTE FOR **GREEN SCIENCE** CARNEGIE MELLON UNIVERSITY



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- We have developed treatment processes that
- Convert >99.9 of BPA in pH ≥11
 lab water, a simulant of RCF
 processing solutions, to CO, CO₂,
 acetone, and trace formate



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- Convert >99.9 of BPA in pH ≥11 lab water, a simulant of RCF processing solutions, to CO, CO₂, acetone, and trace formate
 Reduce the solution phase acute
 - toxicity (takes EC_{50} of 23 to >100)

Acknowledgements

Phipps Conservatory

Richard Piacentini Sarah States

CMU's IGS *Terrence J. Collins* Colin P. Horwitz Evan S. Beach Alexander D. Ryabov Soumen Kundu Matthew R. Mills Ryan T. Malecky Bethany A. Drake

University of Auckland

Naresh Singhal L. James Wright Yusuf Onundi

Oregon State University

Robert L. Tanguay Lisa Truong Michael T. Simonich

Thank You All For Your Attention!

Thomas Zincke of the University of Marburg, Germany, first reported synthesis of bisphenol A. Zincke acknowledged in his paper that the synthesis of BPA, from phenol and acetone, was based on chemical reactions previously reported by others as well as unpublished work (from thesis dissertations) conducted at the University of Marburg. His paper reporting the synthesis of BPA and a number of related compounds was published in 1905. (Zincke, T., 1905, "Mittheilungen aus dem chemischen Laboratorium der Universitat Marburg," Justus Leibigs Annals Chemie, vol. 343, pages 75-99).

Zincke reported key physical properties of BPA (e.g., molecular composition, melting point, solubility in common solvents) but did not propose any application or use for BPA or the other materials he synthesized..

Bisphenol A Developed Commercially in the 1950s

www.bisphenol-A.org

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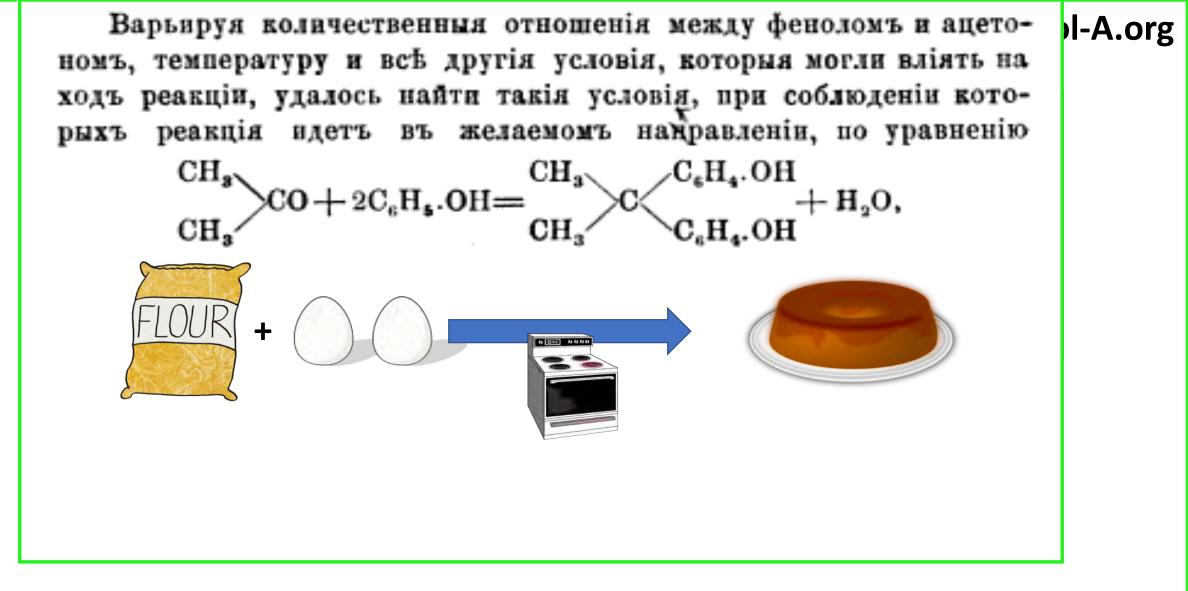
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Bisphenol A Developed Commercially in the 1950s

Варьируя количественныя отношенія между феноломъ и ацето-I-A.org номъ, температуру и всѣ другія условія, которыя могли вліять на ходъ реакціи, удалось найти такія условія, при соблюденіи котореакція идетъ въ желаемомъ найравленіи, по уравненію рыхъ CH3 CH. ′C₅H₄.OH $CO + 2C_6H_5.OH =$ $+H_{2}O$, CH,





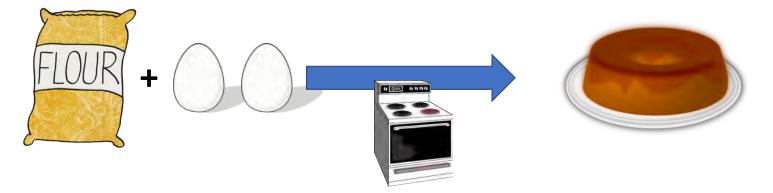
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Bisphenol A Developed Commercially in the 1950s

Bisphenol A First Synthesized in Germany

www.bisphenol-A.org



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Bisphenol A Developed Commercially in the 1950s



NATURE

JUNE 13, 1936

Synthetic Estrogenic Agents without the Phenanthrene Nucleus

	Substance	Dose in mgm.	Percentage positive
	1:2-Dihydroxy-1:2-di-α-naphthyl-	100	100*
	acenaphthene	100	100
_	1:1-Di-a-naphthyl acenaphthenone	100	100
	a-Naphthyl benzoin	100	40
	Diphenyl-a-naphthyl glycol	100	60
	Diphenyl-a-naphthyl carbinol	100	100
	4-4-Dihydroxydiphenyl methane	100	100
	Di-(p-Hydroxyphenyl) dimethyl methane	100	100
	Di-(p-Hydroxyphenyl) methyl ethyl methane	100	100
	Di-(p-Hydroxyphenyl) methyl propyl methane	100	100
	Di-(4-Hydroxy-3-methyl phenyl) di methyl methane	100	100
	Di-(4-Hydroxy-3-methyl phenyl)-1:1-cyclo- hexane	100	100
	2:4-Dihydroxy-triphenyl methane carboxylic acid lactone	100	100
	4:4 ¹ -Dihydroxy benzophenone	100	60
	4:4 ¹ -Dihydroxy diphenyl	100	100

* Rats remained in œstrus 40 days.

Substance	Dose in mgm.	Percentage positive
1:2-Dihydroxy-1:2-di-α-naphthyl-	100	100*
acenaphthene	10	100
1:1-Di-a-naphthyl acenaphthenone	100	100
a-Naphthyl benzoin	100	40
Diphenyl-a-naphthyl glycol	100	60
Diphenyl-a-naphthyl carbinol	100	100
4-4-Dihydroxydiphenyl methane	100	100
Di-(p-Hydroxyphenyl) dimethyl methane	100	100
Di-(p-Hydroxyphenyl) methyl ethyl methane	100	100
Di-(p-Hydroxyphenyl) methyl propyl methane	100	100
Di-(4-Hydroxy-3-methyl phenyl) di methyl methane	100	100
Di-(4-Hydroxy-3-methyl phenyl)-1:1-cyclo- hexane	100	100
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4:4 ¹ -Dihydroxy benzophenone	100	60
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acenaphthene	100	100
33 33 33 33	100	100
1:1-Di-a-naphthyl acenaphthenone		
a-Naphthyl benzoin	100	40
Diphenyl-a-naphthyl glycol	100	60
Diphenyl-a-naphthyl carbinol	100	100
4-4-Dihydroxydiphenyl methane	100	100
Di-(p-Hydroxyphenyl) dimethyl methane	100	100
Di-(p-Hydroxyphenyl) methyl ethyl methane	100	100
Di-(p-Hydroxyphenyl) methyl propyl methane	100	100
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1:1-Di-a-naphthyl acenaphthenone	100	100
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Substance	Dose in mgm.	Percentage positive
1:2-Dihydroxy-1:2-di-α-naphthyl- acenaphthene	100	100*
,,)	10	100
1: 1-Di-a-naphthyl acenaphthenone	100	100
a-Naphthyl benzoin	100	40
Diphenyl-a-naphthyl glycol	100	60
Diphenyl-a-naphthyl carbinol	100	100
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	-	
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acenaphthene	100	100*
57 59 77	10	100
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* Rats remained in œstrus 40 days.								
72 h 52 c) Prol	onged						
$100 \text{ mgs} \longrightarrow 6$	100 mgs $rac{100}{c}$ $rac{100$							
	(40	days)						

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1:2-Dihydroxy-1:2-di-α-naphthyl- acenaphthene	100	100*
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72h) Prol	onged
$100 \text{ mgs} \longrightarrow 6^{\circ}$	= oes	strus =
	(40	days)

Substance	Dose in mgm.	Percentage positive		1936				
1:2-Dihydroxy-1:2-di-α-naphthyl- acenaphthene	100	100*						
>> >> >> >> >> >> >> >> >> >> >> >> >>	10	100						
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a-Naphthyl benzoin	100	40						
Diphenyl-a-naphthyl glycol	100	60						
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* Rats remained in œstrus 40 days. 72 h 72 h 7								
	(40	days)	ατινιτ					



JUNE 13, 1936

Synthetic Estrogenic Agents without the Phenanthrene Nucleus



JUNE 13, 1936

Synthetic Estrogenic Agents without the Phenanthrene Nucleus

612.621.5:547.6

Molecular structure in relation to oestrogenic activity. Compounds without a phenanthrene nucleus

By E. C. Dodds and W. Lawson

Proc. Roy. Soc. 1938

Derivatives of diphenyl methane:	Method of preparation	Dose (mg.)	% positive		
4-Hydroxy-diphenyl methane	Clemmensen (1914)	100	Nil		
3:3'-Dihydroxy-diphenyl methane	Auwers and Rietz (1907)	100	40		
4:4'-Dihydroxy-diphenyl methane	Eberhardt and Welter (1894)	100	100	H	
${ m Diphenyl}$ -methane-3 : 3'-dicarboxylic acid	Schöpf (1894)	100	Nil	E.	
${ m Diphenyl}$ -methane-4 : 4'-dicarboxylic acid	Schöpf (1894)	100	Nil	Ω	
4:4'-Dihydroxy-diphenyl-methane- $3:3'$ -dicarboxylic acid	Kahl (1906)	100	Nil		
2:5-Dihydroxy-diphenyl-methane-carboxylic acid lactone		100	Nil	Dodds	
α -(2-Hydroxy-5-methyl-phenyl) phthalide		100	Nil	dc	
Benzilic acid		100	Nil		.9
2:2'-Dihydroxy-di- $lpha$ -naphthyl methane	Fries and Hübner (1906)	100	Nil	ar	
$lpha: lpha ext{-Diphenyl}$ ethylene	Klages (1902)	100	Nil	and	
Derivatives of 4 : 4'-dihydroxy diphenyl methane (Easson, Ha R R' $Methyl$ MethylMethylMethylMethyl n -PropylEthyl n -PropylEthyl n -HexylH n -HexylH n -Henyl	arrison, McSwiney and Pyman 1934): $HO = \begin{pmatrix} R \\ C \\ R' \end{pmatrix} OH$ Russanow (1889)	$ \begin{array}{r} 100 \\ 100 \\ 100 \\ 100 \\ 100 \\ 100 \\ 10 \\ 100 \\ 100 \\ $	100 100 100 100 100 100 20 60 Nil	W. Lawson	

Derivatives of diphenyl methane:	Method of preparation	Dose (mg.)	% positive		
 4-Hydroxy-diphenyl methane 3: 3'-Dihydroxy-diphenyl methane 4: 4'-Dihydroxy-diphenyl methane Diphenyl-methane-3: 3'-dicarboxylic acid Diphenyl-methane-4: 4'-dicarboxylic acid 4: 4'-Dihydroxy-diphenyl-methane-3: 3'-dicarboxylic acid 2: 5-Dihydroxy-diphenyl-methane-carboxylic acid lactone \$\alpha\$-(2-Hydroxy-5-methyl-phenyl) phthalide Benzilic acid 2: 2'-Dihydroxy-di-\$\alpha\$-naphthyl methane \$\alpha\$: \$\alpha\$-Diphenyl ethylene 	Clemmensen (1914) Auwers and Rietz (1907) Eberhardt and Welter (1894) Schöpf (1894) Schöpf (1894) Kahl (1906) — Fries and Hübner (1906) Klages (1902)	$ \begin{array}{r} 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100\\ 100 \end{array} $	Nil 40 100 Nil Nil Nil Nil Nil Nil Nil Nil	E. C. Dodds and	.9:

Derivatives of 4:4'-dihydroxy diphenyl methane (Easson, Harrison, McSwiney and Pyman 1934):

R Methyl	$R^{\prime} \ { m Methyl}$	R	100	100	Law
Methyl Methyl F nenyi	Ethyl n-Propyl Ethyl n-Hexyl Phenyl Phenyl Phenyl	HOCCCCOH R' Russanow (1889)	$ \begin{array}{r} 100 \\ $	100 100 100 100 100 20 60 Nil	nos

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JUNE 13, 1936

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612.621.5:547.6

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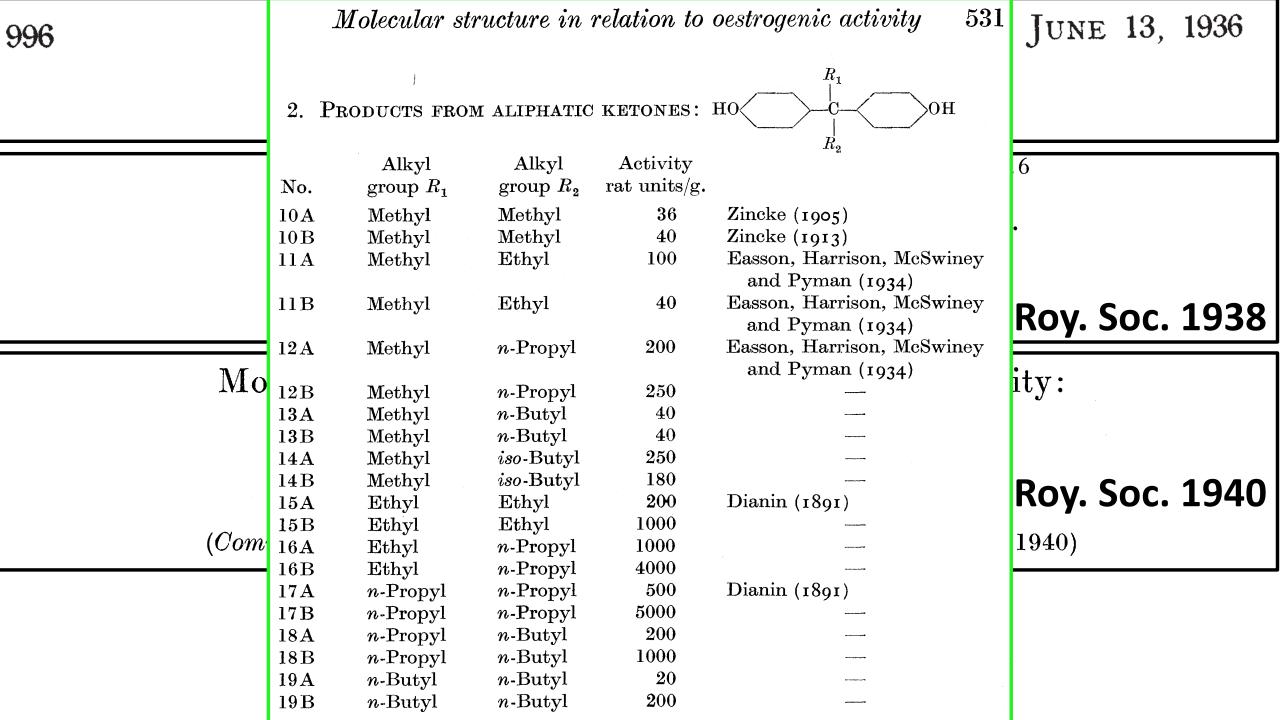
Proc. Roy. Soc. 1938

Molecular structure in relation to oestrogenic activity: derivatives of 4:4'-dihydroxydiphenylmethane

By N. R. CAMPBELL

Proc. Roy. Soc. 1940

(Communicated by Sir Robert Robinson, F.R.S.—Received 26 August 1940)



996	М	olecular str	ructure in a	relation to a	oestrogenic activity 531	JUNE 13, 1936
	2. Pro) DUCTS FROM	1 ALIPHATIC	KETONES:]	HO $($	
	No.	Alkyl group R_1	Alkyl group R_2	Activity rat units/g.		6
	10 A	${\bf Methyl}$	${f Methyl}$	36	Zý	
	$10\mathrm{B}$	Methyl	Methyl	40		
	11A	Methyl	Ethyl	100	L Swiney and J (-934)	
	11B	Methyl	\mathbf{Ethyl}	40	Easson, Harrison, McSwiney and Pyman (1934)	Roy. Soc. 1938
	12A	\mathbf{Methyl}	n-Propyl	200	Easson, Harrison, McSwiney	
${ m Mo}$		·			and Pyman (1934)	i+x7 •
	1~1	Methyl	n-Propyl	250		ity:
	13A	Methyl	n-Butyl	40		
	1 3 B	Methyl	n-Butyl	40		
	14A	Methyl	iso-Butyl	250		
	14B	Methyl	iso-Butyl	180	 D: : (0)	Roy. Soc. 1940
	15A	Ethyl	Ethyl	200	Dianin (1891)	NOY. 30C. 1340
(Com)	15B	Ethyl	Ethyl	1000		1940)
$(\bigcirc m)$		Ethyl Ethal	n-Propyl	1000		1010)
	16B	Ethyl ^m Propyl	n-Propyl	4000 500	Dianin (1801)	
	17A 17B	<i>n</i> -Propyl	n-Propyl n-Propyl	500 5000	Dianin (1891)	
	17B 18A	$n\operatorname{-Propyl} n\operatorname{-Propyl} $	$n\operatorname{-Propyl} onumber n\operatorname{-Butyl} onumber n$	200		
	18A 18B	<i>n</i> -Propyl	n-Butyl n -Butyl	1000		
	18 D 19 A	<i>n</i> -Propyr <i>n</i> -Butyl	<i>n</i> -Butyl	20		
	19A 19B	<i>n</i> -Butyl	<i>n</i> -Butyl	200		

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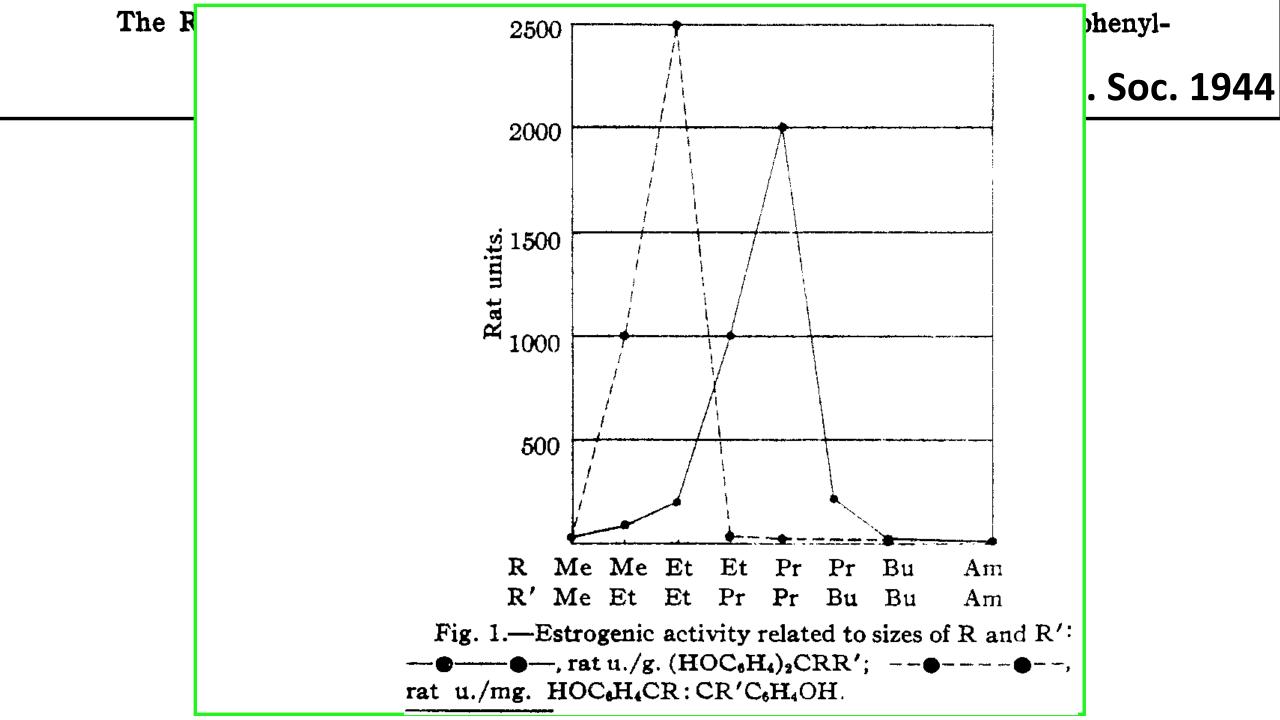
By N. R. CAMPBELL

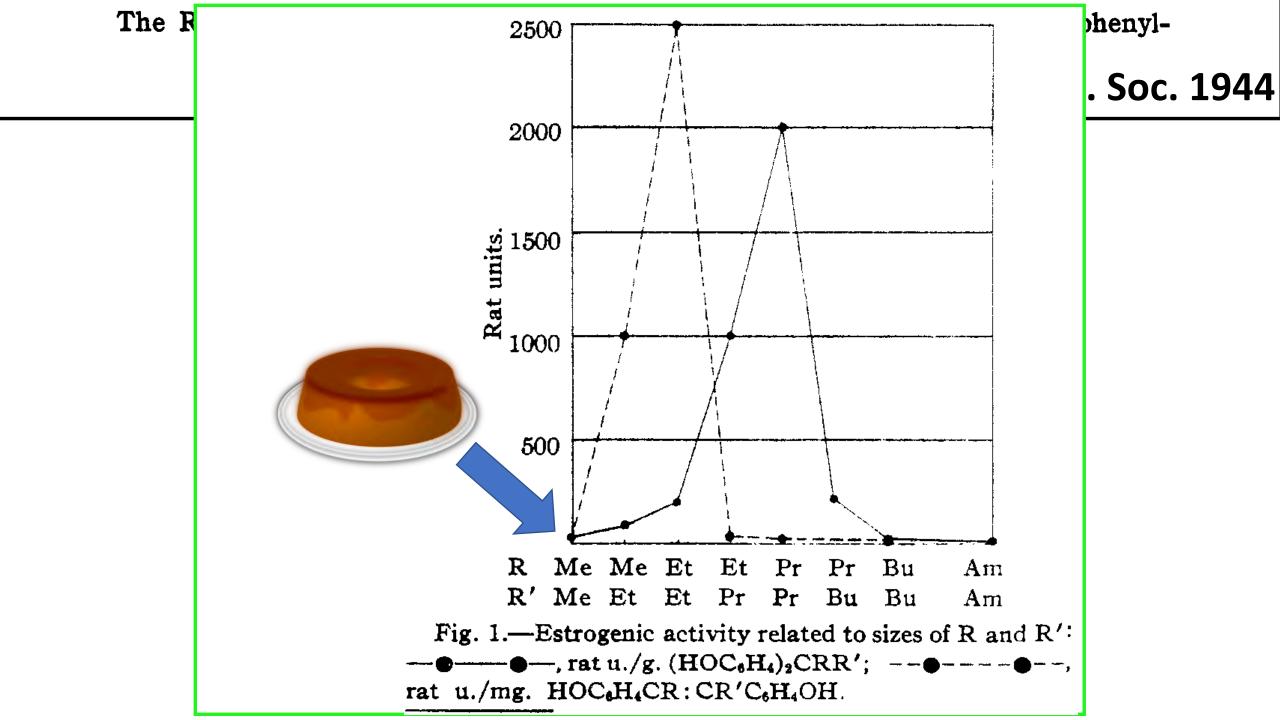
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The Relation of Estrogenic Activity to Structure in Some 4,4'-Dihydroxydiphenylmethanes¹

By E. Emmet Reid and Edith Wilson² J. Am. Chem. Soc. 1944





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SYNTHETIC ESTROGENS AND THE RELATION BETWEEN THEIR STRUCTURE AND THEIR ACTIVITY ULRICH V. SOLMSSEN

Research Laboratories of Hoffman-La Roche, Inc., Nutley 10, New Jersey

Received April 13, 1945

Chem. Rev. 1945

	TABLE 15 Diphenylmethane derivatives							
			MELTING Point	NAME	ESTROGENIC ACTIVITY SUBCUTANEOUSLY (100 RESPONSE UNLESS IN OTHERWISE)	PER CENT DICATED	REFERENCES	1944
-	R'	R'			Minimum effective dose	Rat units		
-	Н	И	°C.	Bis(4-hydroxyphenyl)methane	micrograms 100,000	20	(61, 192) (21)	-
	Н	CH ₃	122	1,1-Bis(4-hydroxyphenyl)ethane		20	(21, 89)	
	CH_3	CHI3	155	2,2-Bis(4-hydroxyphenyl)propane	100,000	36 28	(61, 64) (21, 231) (158)	
	II	C_2H_5	129	1,1-Bis(4-hydroxyphenyl)propane	100,000	16	(61) (21, 89)	.945
	CH3	C_2H_5	124	2,2-Bis(4-hydroxyphenyl)butane	100,000	124 100	(61, 64, 67, 120) (158) (21)	
	Н	n-C ₃ H ₇	137	1,1-Bis(4-hydroxyphenyl)butane		40	(21, 89)	
	н	Iso-C ₃ H ₇	152	1,1-Bis(4-hydroxyphenyl)-2-methyl- propane		36	(21)	
	CH3	<i>n</i> -C ₃ H ₇	149	2,2-Bis(4-hydroxyphenyl)pentane	100,000	200	(61, 64, 67, 120) (21, 158)	

TABLE 15												
				Diphenylmethane derivatives				-				
_							MELTING Poi nt	NAME	ESTROGENIC ACTIVITY SUBCUTANEOUSLY (100 RESPONSE UNLESS IN OTHERWISE)	PER CENT DICATED	REFERENCES	1944
-	R'	R'			Minimum effective dose	Rat units per gram						
-	Н	И	°C.	Bis(4-hydroxyphenyl)methane	micrograms 100,000	20	(61, 192) (21)					
	Н	CH_3	122	1,1-Bis(4-hydroxyphenyl)ethane		20	(21, 89)					
	CH3	CH ₃	155	2,2-Bis(4-hydroxyphenyl)propane	100,000	36 28	(61, 64) (21, 231) (158)					
	п		129	1,1-Bis(4-hydroxyphenyl)propane	100,000	16	(61) (21, 89)	.945				
	CH3		4	2,2-Bis(4-hydroxyphenyl)butane	100,000	124 100	(61, 64, 67, 120) (158) (21)					
	н	76-Vore1	137	1,1-Bis(4-hydroxyphenyl)butane		40	(21, 89)					
	н	$Iso-C_3H_7$	152	1,1-Bis(4-hydroxyphenyl)-2-methyl- propane		36	(21)					
	CH3	<i>n</i> -C ₃ H ₇	149	2,2-Bis(4-hydroxyphenyl)pentane	100,000	200	(61, 64, 67, 120) (21, 158)					

www.bisphenol-A.org



A. Dianin, **1891**, J. Russian Physical Chemistry Society, **23**: 488-517.

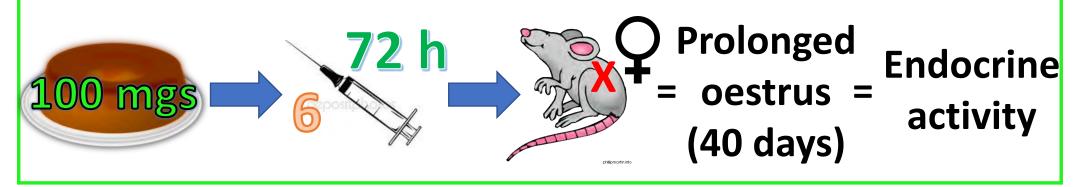
Zincke reported key physical properties of BPA (e.g., molecular composition, melting point, solubility in common solvents) but did not propose any application or use for BPA or the other materials he synthesized..

Bisphenol A Developed Commercially in the 1950s

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A. Dianin, 1891, J. Russian Physical Chemistry Society, 23: 488-517.

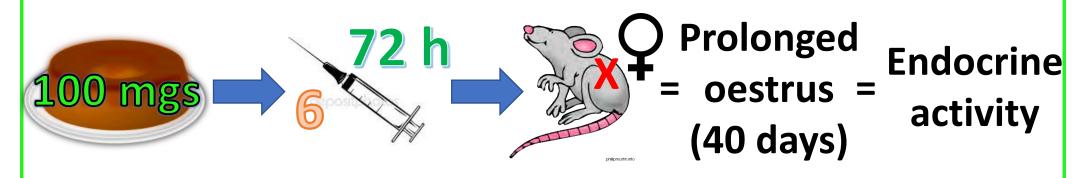


Bisphenol A Developed Commercially in the 1950s

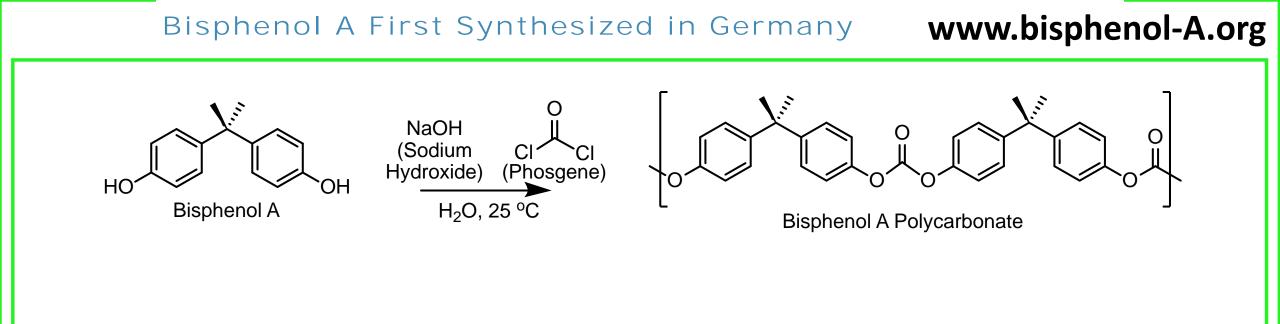
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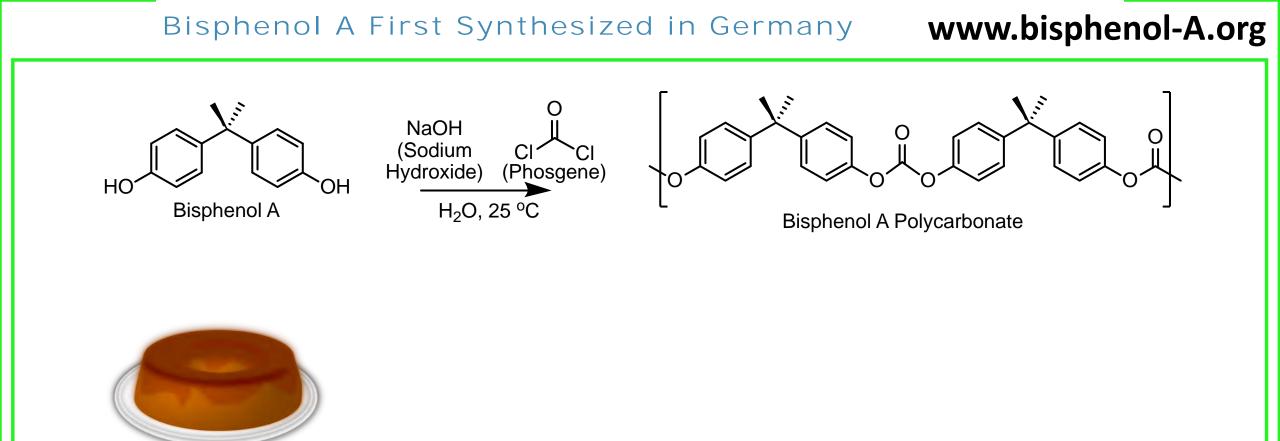


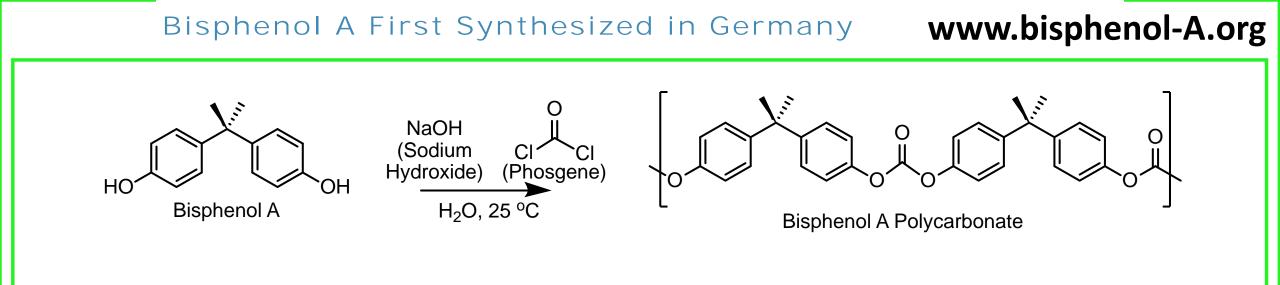
A. Dianin, 1891, J. Russian Physical Chemistry Society, 23: 488-517.



Bisphenol A Developed Commercially in the 1950s

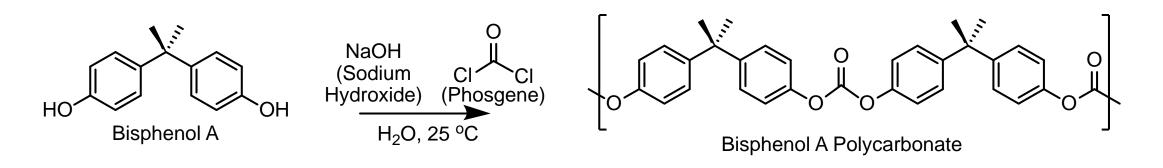


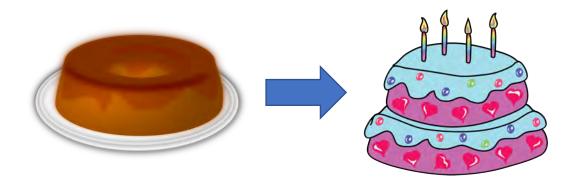




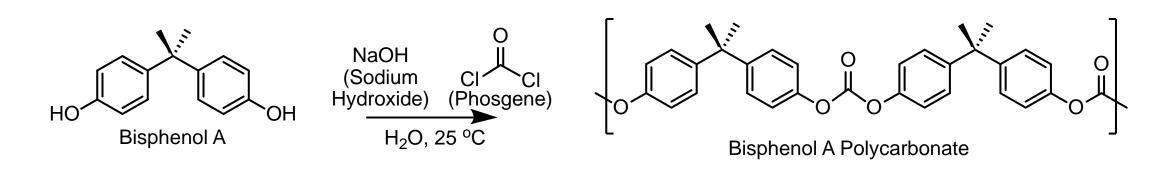


Bisphenol A First Synthesized in Germany www.bisphenol-A.org





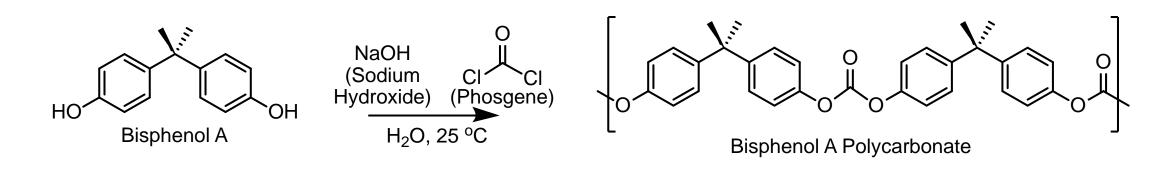
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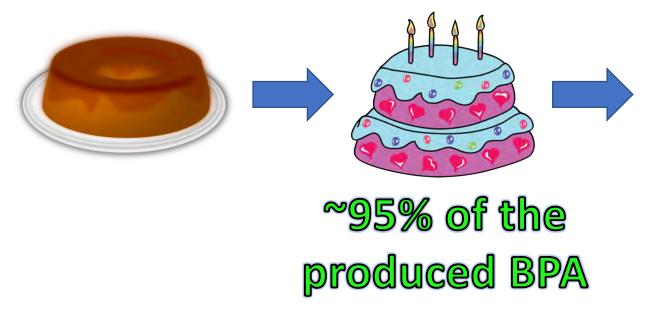




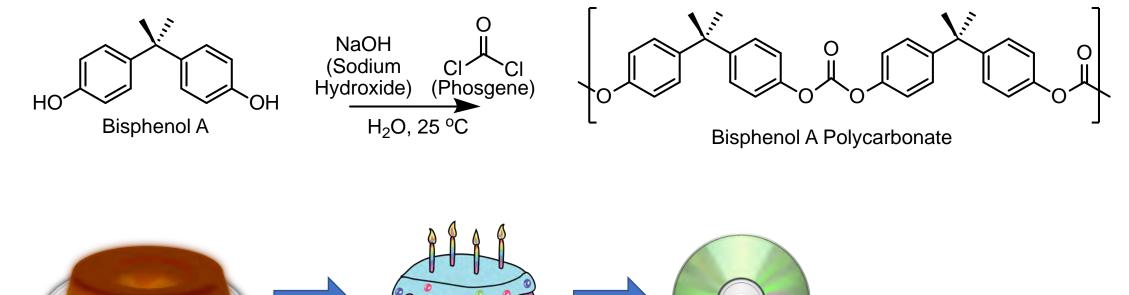
produced BPA

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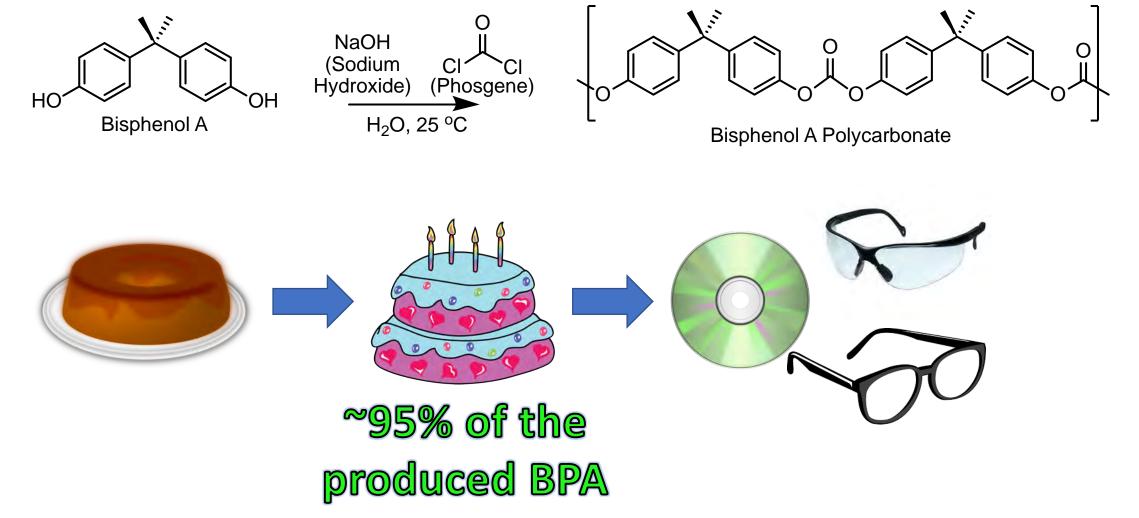


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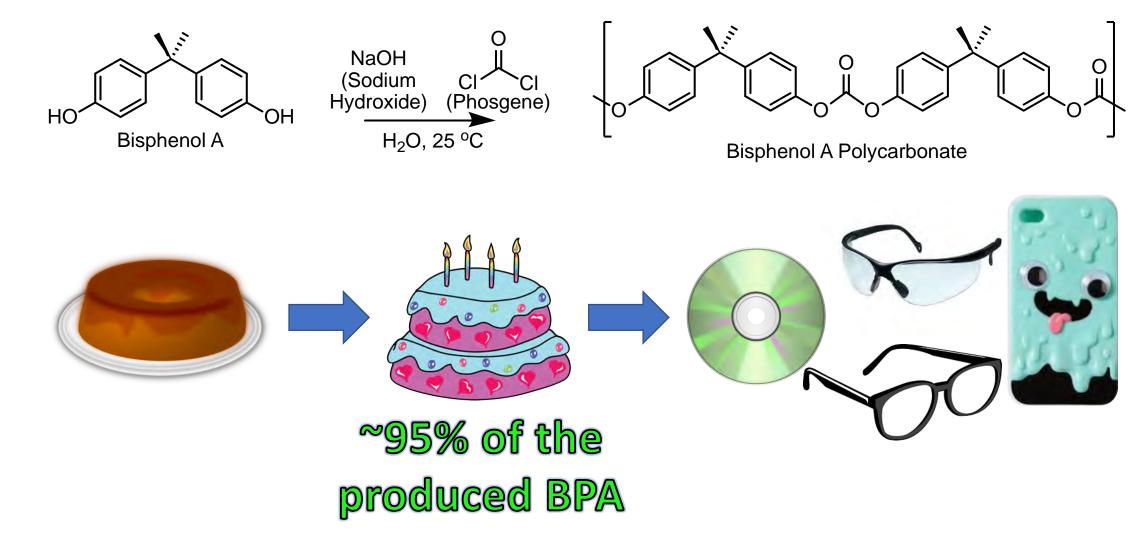


Solution of the produced BPA

Bisphenol A First Synthesized in Germany www.bisphenol-A.org



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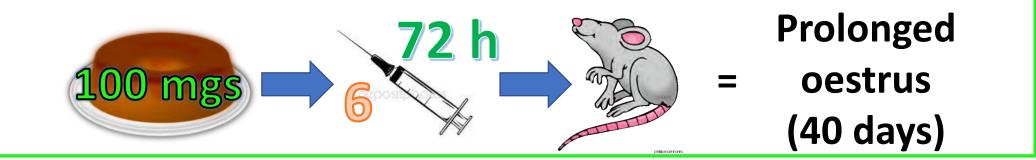




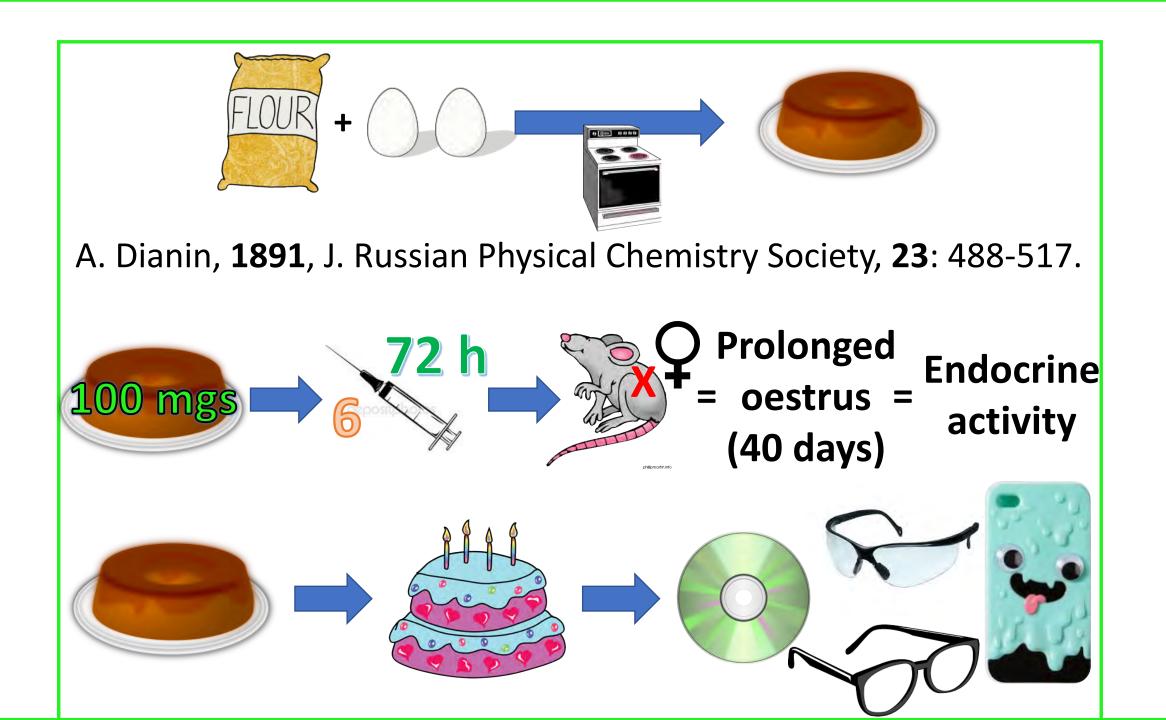


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A. Dianin, 1891, J. Russian Physical Chemistry Society, 23: 488-517.



Bisphenol A Developed Commercially in the 1950s



• Newspapers



- Newspapers
- Toilet Paper



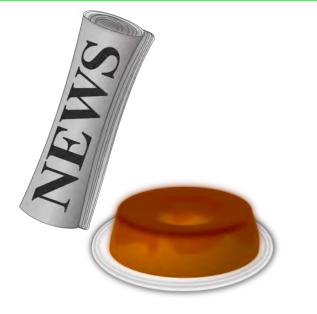


- Newspapers
- Toilet Paper



Contributes 36,000 Ibs of BPA to wastewater per year

- Newspapers
- Toilet Paper
- Pizza Boxes



Contributes 36,000 lbs of BPA to wastewater

- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes



Contributes 36,000 lbs of BPA to wastewater



- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes



Contributes 36,000 lbs of BPA to wastewater

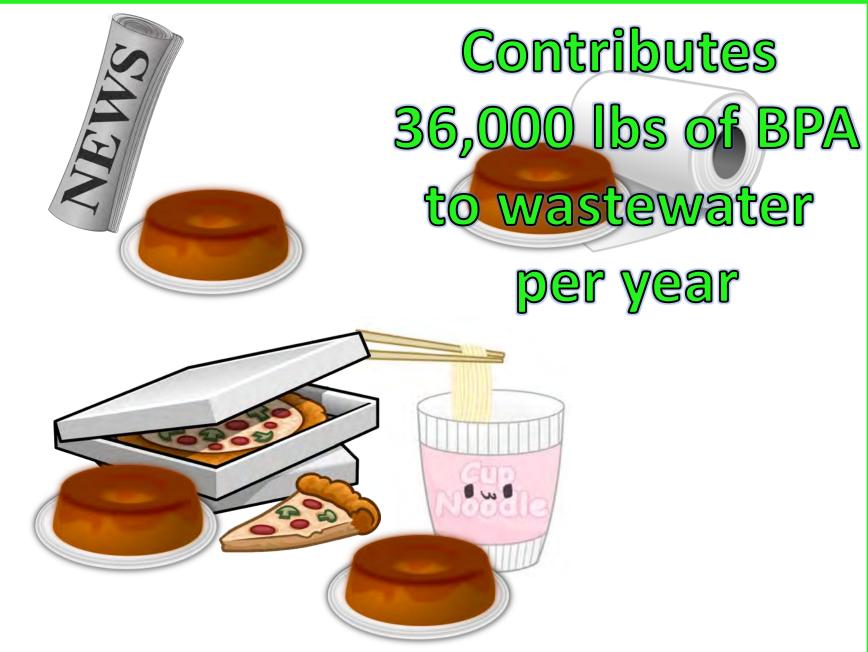


- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes
- Sandwich Boxes



Contributes 36,000 lbs of BPA to wastewater

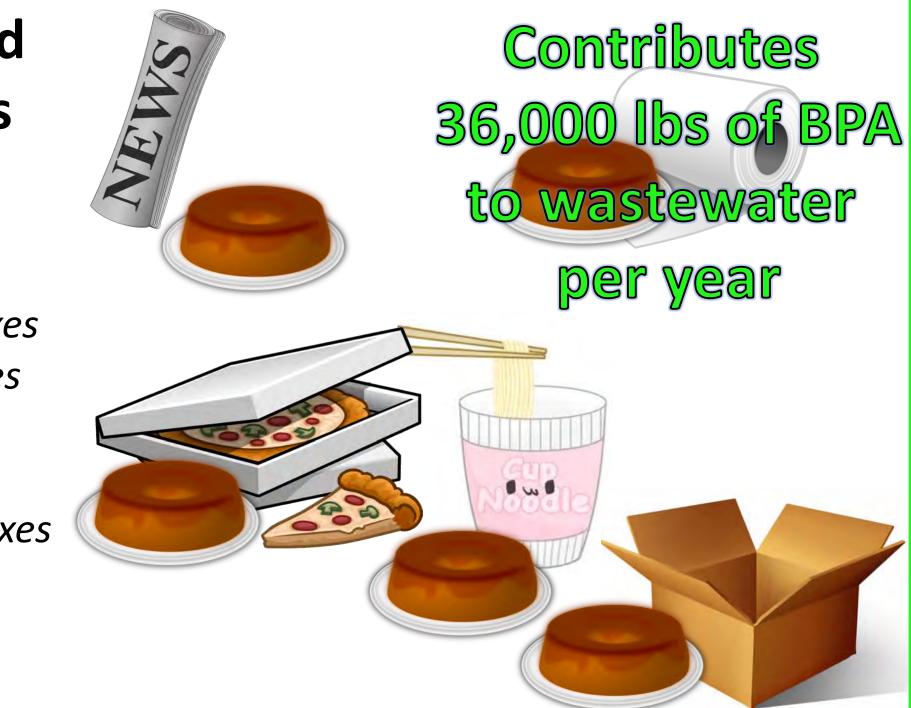
- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes
- Sandwich Boxes
- Noodle Cups



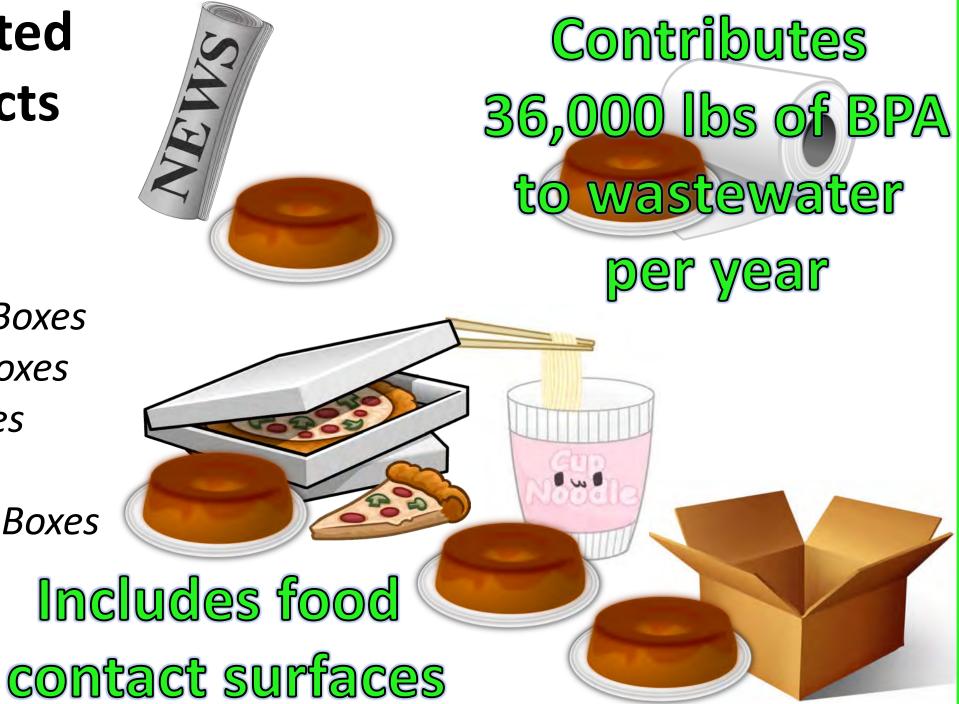
- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes
- Sandwich Boxes
- Noodle Cups
- Confectionary Boxes



- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes
- Sandwich Boxes
- Noodle Cups
- Confectionary Boxes
- General Food Storage Boxes



- Newspapers
- Toilet Paper
- Pizza Boxes
- Fried Chicken Boxes
- Fried Potato Boxes
- Sandwich Boxes
- Noodle Cups
- Confectionary Boxes
- General Food
 Storage Boxes



Liao 2011 Gehring 2004 Ozaki 2004 Lopez-Espinosa 2007 USEPA 2015